ELSEVIER



# World Neurosurgery: X



journal homepage: www.journals.elsevier.com/world-neurosurgery-x

# Long-term quality of life outcomes in patients undergoing microsurgical resection of vestibular schwannoma $\stackrel{\star}{\sim}$

Loren N. Riedy<sup>a</sup>, Rachyl M. Shanker<sup>b</sup>, Dayna C. Sloane<sup>c</sup>, Daniel M. Heiferman<sup>d</sup>, Elhaum G. Rezaii<sup>e</sup>, Sarah E. Finucane<sup>f</sup>, Perry Veras<sup>c</sup>, John P. Leonetti<sup>f</sup>, Douglas E. Anderson<sup>g,\*</sup>

<sup>a</sup> Committee Neurobiology, University of Chicago, Hyde Park, IL, USA

<sup>b</sup> Department of Surgery, Loyola University Medical Center, Maywood, IL, USA

<sup>c</sup> Stritch School of Medicine, Loyola University Chicago, Maywood, IL, USA

<sup>d</sup> Department of Neurological Surgery, Edward-Elmhurst Health, Naperville, IL, USA

<sup>e</sup> Department of Neurological Surgery, University of Nebraska Medical Center, Omaha, NE, USA

<sup>f</sup> Department of Otolaryngology, Loyola University Medical Center, Maywood, IL, USA

<sup>g</sup> Department of Neurological Surgery, Loyola University Medical Center, Maywood, IL, USA

ARTICLE INFO	A B S T R A C T			
Keywords: Vestibular schwannoma PANQOL Quality of life Surgical outcomes Skull base Brain tumor	<i>Background</i> : While previous studies have assessed patient reported quality of life (QOL) of various vestibular schwannoma (VS) treatment modalities, few studies have assessed QOL as related to the amount of residual tumor and need for retreatment in a large series of patients. Objective: To assess patient reported QOL outcomes following VS resection with a focus on extent of resection and retreatment. <i>Methods</i> : A retrospective chart review was performed using single-center institutional data of adult patients who underwent VS resection by the senior authors between 1989-2018 at Loyola University Medical Center. The Penn Acoustic Neuroma Quality of Life (PANQOL) survey was sent to all patients via postal mail. <i>Results</i> : Fifty-five percent of 367 total patients were female with a mean age of 61.6 years (SD 12.63). The mean period between surgery and PANQOL response was 11.4 years (IQR: 4.74-7.37). The median tumor size was 2 cm (IQR: 1.5-2.8). The mean total PANQOL score was 70 (SD 19). Patients who required retreatment reported lower overall scores ( $\mu$ diff = -10.11, 95% CI: -19.48 to -0.74; p = 0.03) and face domain scores ( $\mu$ diff = -20.34, 95% CI: -29.78 to -10.91; p < .001). There was no association between extent of resection and PANQOL scores in any domain. <i>Conclusion</i> : In an analysis of 367 patients who underwent microsurgical resection of VS, extent of resection did not affect PANQOL scores in contrast to previous reports in the literature, while the need for retreatment and facial function had a significant impact on patient-reported outcomes.			

#### 1. Introduction

Vestibular schwannomas (VS), formerly termed acoustic neuromas, are benign tumors of the cranial base with a natural history resulting in hearing loss, ataxia, hydrocephalus, and brainstem compression, all of which have been shown to negatively affect quality of life (QOL).<sup>1–3</sup> Historically, VS resection carried a high risk of morbidity and mortality. However, with technological advancements and the development of surgical practice, outcomes have dramatically improved while balancing extent of resection with optimized neurological outcomes,

including facial function and hearing preservation.<sup>4</sup> Furthermore, analysis of these measures has resulted in improved preoperative treatment planning.<sup>5–11</sup> Patient-reported metrics have the ability to shed light onto the most clinically important outcomes that influence QOL, which can guide our focus and management of VS. The Penn Acoustic Neuroma Quality of Life (PANQOL) questionnaire has been validated as a method to assess QOL in patients with VS, and stratifies QOL outcomes into an overall score and subdomains relevant to various area of life functions.<sup>12</sup>

The PANQOL questionnaire has been applied in several studies

https://doi.org/10.1016/j.wnsx.2024.100294

Received 6 April 2023; Accepted 20 February 2024

Available online 24 February 2024

2590-1397/© 2024 The Authors. Published by Elsevier Inc. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).

<sup>\*</sup> Portions of this work were presented in part at the Congress of Neurological Surgeons Annual Meeting, Austin, Texas, October 16–20, 2021.

<sup>\*</sup> Corresponding author. Department of Neurological Surgery Loyola University Medical Center 2160 S. 1st Ave Maguire Center, Rm 1900 Maywood, IL 60153, USA.

E-mail address: Dander1@lumc.edu (D.E. Anderson).

analyzing QOL outcomes in patients between different VS treatment groups, such as microsurgery, stereotactic radiosurgery, and observation, and authors have suggested various factors that influence patient QOL.<sup>13–21</sup> Here, we utilize the PANQOL questionnaire in patients who elected to receive microsurgical resection of VS to determine the baseline and surgical characteristics that most significantly impact QOL outcomes. The present study not only analyzes QOL outcomes in a large microsurgical cohort, but also provides analysis of how facial and hearing preservation, extent of resection, and retreatment vary in their association with QOL outcomes.

# 2. Methods

# 2.1. Patient population

A retrospective review was conducted of patients aged 17 years and older who underwent microsurgical resection of VS by senior authors (D. E.A. and J.P.L) between 1989 and 2018 at Loyola University Medical Center. PANQOL surveys were mailed to patients' homes along with a return envelope with postage and a letter from corresponding author, D. E.A., detailing the study. Patients who did not respond within six weeks received a single follow-up phone call. This study was approved by the Loyola University Stritch School of Medicine Institutional Review Board (reference number 211402), and informed consent was received from all survey respondents.

# 2.2. Patient characteristics

A retrospective chart review was conducted to collect baseline demographic information and tumor characteristics, including tumor lateralization and size in the greatest dimension as seen on magnetic resonance imaging (MRI). Pre- and postoperative symptoms were recorded, including tinnitus, imbalance, headache, cerebrospinal fluid leak, and postoperative House-Brackmann (HB) grade.

#### 2.3. PANQOL survey

The PANQOL questionnaire, developed in 2010 by Shaffer et al,<sup>12</sup> consists of 26 questions grouped into 7 domains determined to be most pertinent to quality of life (QOL): hearing, balance, facial, anxiety, energy, pain, and general health. Patients self-reported answers on a scale of 1–5, with 1 representing strong discordance and 5 representing strong accordance. Individual scores from each question were then converted to a 100-point scale as follows: for all questions except for question 25, a strong accordance reflected a lower quality of life. As such, a score of 5 was assigned a value of 0, a score of 4 was assigned a value of 25, and so forth. For question 25, a strong accordance reflected a higher quality of life, so a score of 5 was assigned a value of 100, a score of 4 was assigned a value of 75, and so forth. For each domain, the scores for the associated questions were averaged, with a higher score indicating a better QOL. The primary outcomes were individual domain and total PANQOL scores.

#### 2.4. Extent of resection

The presence of residual tumor was assessed intraoperatively using a custom 1 cm graduated bayoneted micro-measuring instrument. Extent of tumor resection was divided into two groups: gross total resection (GTR) and less than GTR. GTR was defined as total removal by the surgeon's operative note with no observable residual tumor identified on postoperative MRI. Less than GTR was characterized by any presence of a residual tumor placode either measurable intraoperatively or postoperative imaging.

#### 2.5. Retreatment

Retreatment was defined as requiring at least one additional microsurgical intervention for resection of residual or recurrent tumor. Decision for retreatment was made between operating surgeon and patient in the setting of residual or recurrent tumor causing return or continuation of symptoms associated with VS.

#### 2.6. Hearing preservation

Postoperative hearing preservation was graded based on audiogram at most recent follow up according to the Gardner-Robertson hearing scale (GR).<sup>22</sup> Pure tone average (PTA) was calculated as a mean of 500, 1000, 2000, and 3000 Hz. Serviceable hearing was defined as GR grade 1 and 2.<sup>22</sup> Patients who were not candidates for hearing preservation surgery due to complete hearing loss prior to surgery or whose data were lost were excluded from analysis.

# 2.7. Facial outcome

Facial outcomes were graded using the House-Brackmann scoring system.<sup>23</sup> Grade I describes normal facial function, grade II describes mild dysfunction, grade III describes moderate dysfunction notably with complete eye closure with effort, grade IV describes moderately severe dysfunction with incomplete eye closure, grade V describes severe dysfunction, and grade VI describes total paralysis. For the purpose of our study, grade I and II were considered ideal facial nerve functional outcome.

# 2.8. Year of operation

Based on our previous study looking at the learning curve of VS resections of the operating neurosurgeons from 1988 to 2018, we looked at PANQOL scores from 1988-2004, 2005–2009, and 2010–2018. The probability of attaining a HB score of I was twofold higher in 2005–2009 and 2010–2018 as compared with 1988–2004.<sup>24</sup>

# 2.9. Statistical analysis

Continuous variables were summarized using mean with standard deviation or median with interquartile range (IQR). Nominal and ordinal variables were summarized using counts and proportions. A linear regression model was used to test whether PANQOL scores were associated with age, months since symptoms began, and tumor size. In this model, the assumptions of linearity, normality, and homoscedasticity were assessed using residual plots. An independent samples t-test was used to test whether the distribution of PANQOL scores varied by sex, residual symptoms, retreatment, hearing preservation status, presence of a CSF leak, and House-Brackmann grade. A general linear model was used to test whether the distribution of PANQOL scores varied by tinnitus, balance, and headache symptoms. A Kruskal-Wallis test was used to test for differences in the distribution of each PANQOL score among patients with a HB grade of I, II, III, and IV-VI; when the overall significance value was less than .05, all possible post-hoc pairwise comparisons were tested using the Dwass (1960), Steel (1960), Critchlow-Fligner method (1991).<sup>25–27</sup>

Following univariable analysis, multivariable general linear models were used to estimate adjusted PANQOL scores. In these models, covariates were included if they improved model fit as measured by Akaike's Information Criterion (AIC). In this study, a *p*-value less than .05 was considered statistically significant. All analyses were completed using SAS version 9.4 (Cary, NC).

#### 3. Results

#### 3.1. Baseline demographics

Of the 881 patients who underwent VS microsurgical resection, 175 patients were either deceased or lost to follow-up with no valid address or phone number available. Of the 706 remaining patients, 367 PANQOL surveys were returned (52%). Fifty-five percent of patients were female with a mean age of 61.6 years (SD 12.63). The mean time between surgery and PANQOL survey was 11.4 years (median 10.53; IQR: 4.74–17.37), and the median tumor size was 2 cm (IQR: 1.5–2.8). Two-hundred and four patients (55.6%) had a left-sided tumor, while the remaining 163 (44.4%) had a right-sided tumor. Zero patients had bilateral VS. The retrosigmoid approach was more commonly performed than the translabyrinthine or combined retrosigmoid/translabyrinthine approach (49% vs 37% vs 10%). A minority of patients underwent middle fossa approach. The mean period of follow-up between surgery and last clinical encounter was 81.3 months (median 40.05 months; IQR: 13.10–91.77) (Table 1).

#### 3.2. Functional outcomes

House-Brackmann (HB) scores at most recent follow-up were recorded for 359 patients. Seventy-six percent of patients reported a HB grade I, 12.8% HB grade II, 8.4% HB grade III, .8% HB grade IV, .8% HB grade V, and 1.1% HB grade VI. HB scores were broken down into patients with ideal HB grade (I-II) (n = 319, 88.9%) and poor HB grade (III-VI) (n = 40, 11.1%) (Tables 2–5), in addition to further stratification (I vs II vs III vs IV-VI) in Table 6.

Gardner-Robertson (GR) scores were used to determine degree of hearing preservation, with serviceable hearing defined as GR I-II. Serviceable hearing at most recent follow-up was reported in 28% of patients with follow-up audiometric data.

# 3.3. PANQOL scores

#### 3.3.1. Total

The mean total PANQOL score was 70/100 (SD 19). Controlling for sex, residual status, and retreatment status, patients with a poor HB grade (III-VI) reported lower overall quality of life ( $\mu$ diff = -7.85, 95% CI: -14.76 to -.93; *p* = 0.03). Similarly, controlling for all other

#### Table 1

Baseline characteristics among 367 patients who underwent microsurgical resection of vestibular schwannoma.

Characteristic	N=367					
Age at survey years						
Mean (SD)	61.55 (12.63)					
Median (IQR)	61.96 (53.32–71.21)					
<b>Sex</b> <i>n</i> (%)						
Female	202 (55)					
Male	165 (45)					
Tumor size cm						
Mean (SD)	2.23 (1.03)					
Median (IQR)	2 (1.5–2.8)					
Laterality n (%)						
Left	204 (55.6)					
Right	163 (44.4)					
Extent of resection n (%)						
Gross total	334 (91)					
Subtotal	33 (9)					
Retreatment n (%)	20 (5.4)					
Follow-up years						
Mean (SD)	6.78 (13.04)					
Median (IQR)	3.34 (1.09–7.65)					
Period from surgery to PANQOL survey years						
Mean (SD)	11.49 (7.50)					
Median (IQR)	10.53 (4.74–17.37)					

Table 2			
PANQOL total scores as a	function	of patient	characteristics.

	Valid	Unadjusted		Adjusted	
	Ν	β (95% CI)	р	β (95% CI)	р
Age (per 1-year increase)	367	06 (22 to .09)	.41		
Sex: Male vs Female	367	4.27 (.43–8.10)	.03	5.25 (.90–9.61)	.02
Months since symptoms (per 1-month increase)	367	.01 (01 to	.48	.03 (01 to	.15
Residual: Yes vs No	366	-4.39 (-9.67 to .89)	.10	-1.91 (-7.45 to 3.63)	.50
Retreatment: Yes vs No	261	-8.75 (-16.96 to 55)	.04	-10.11 (-19.48 to 74)	.03
Tinnitus	364		.30		
Only post-op versus no post-op		-7.23 (-19.23 to 4.77)	.33		
Pre-and-post op versus no post-op		-2.18 (-8.94 to 4.58)	.73		
Pre-and-post op versus only post-op		5.05 (-8.24 to 18.35)	.64		
Balance	253		.33		
Only post-op versus no post-op		-2.32 (-10.19 to 5.54)	.77		
Pre-and-post op versus no post-op		-5.51 (-14.84 to 3.82)	.35		
Pre-and-post op versus only post-op		-3.19 (-14.62 to 8.24)	.79		
Headache	361		.01		
Only post-op versus no post-op		-12.09 (-21.28 to -2.89)	.01		
Pre-and-post op versus no post-op		-6.15 (-22.76 to 10.46)	.66		
Pre-and-post op versus only post-op		5.93 (–12.74 to 24.61)	.74		
Hearing Preservation: Yes vs No	93	3.83 (-4.26 to 11.92)	.35		
CSF Leak: Yes vs No	364	.16 (-4.96 to 5.28)	.95		
House–Brackmann: Score 3–6 versus Score 1–2	359	-5.61 (-11.76 to .55)	.07	-7.85 (-14.76 to 93)	.03
Size (per 1 cm increase)	366	86(-2.73)	.37		

Valid N = The number of patients used for the estimates. The sample size for the adjusted estimates = 260. PANQOL total scores range from 0 to 100 with higher scores indicating greater quality of life.

variables in the model, patients who required retreatment reported lower overall quality of life ( $\mu$ diff = -10.11, 95% CI: -19.48 to -.74; *p* = 0.03). Conversely, controlling for all other variables in the model, male patients reported higher overall QOL compared to female patients ( $\mu$ diff = 5.25, 95% CI: .90 to 9.61; *p* = 0.02) (Table 2).

#### 3.3.2. Hearing

The mean score in the hearing domain was 56/100 (SD 24). PANQOL hearing domain scores were higher for males ( $\mu$ diff = 5.21, 95% CI: .26–10.15; *p* = 0.04) and for those with hearing preservation ( $\mu$ diff = 11.22, 95% CI: 1.33–21.11; *p* = 0.03). Otherwise, PANQOL hearing domain scores were not associated with remaining patient characteristics in this sample of data (Supplemental Table 1).

#### Table 3

PANQOL face scores as a function of patient characteristics.

Table 4					
PANOOL pai	n scores	as a	function	of patient	characteristics.

	Valid	Unadjusted		Adjusted	р
	N	β (95% CI)	р	β (95% CI)	
Age (per 1-year	367	09	.38		
increase)		(29 to .11)			
Sex: Male vs Female	367	4.38 (76 to 9.53)	.09		
Months since	367	03	.05		
symptoms (per 1-		(06 to			
month increase)		.00)			
Residual: Yes vs No	366	-6.84	.06	23	.94
		(-13.90  to)		(-6.68  to)	
Petreatment: Vec vc No	261	.22)	002	0.21)	< 001
Retreatment. Tes vs No	201	-23.34 (-40.43 to	.002	-20.34 (-29.78 to	<.001
		-10.25)		-10.91)	
Tinnitus	364		.34		
Only post-op versus		-4.25	.81		
no post-op		(-20.22 to			
		11.72)			
Pre-and-post op		4.85	.41		
versus no post-op		(-4.15 to			
Dro and nost on		13.85)	45		
versus only post-on		(-8.60  to)	.45		
verbub only post op		26.80)			
Balance	253		.30		
Only post-op versus		-6.05	.36		
no post-op		(-16.41 to			
		4.32)			
Pre-and-post op		-4.66	.64		
versus no post-op		$(-16.96\ to$			
Pre-and-post op		1.39	97		
versus only post-op		(-13.68 to			
		16.45)			
Headache	361		.13		
Only post-op versus		-9.54	.16		
no post-op		(-21.83 to			
Due and next on		2.74)	50		
versus no post-op		-9.27 (-31.47 to	.39		
versus no post op		12.93)			
Pre-and-post op		.27	.99		
versus only post-op		(-24.69 to			
		25.24)			
Hearing Preservation:	93	3.98	.39		
Yes vs No		(-5.10  to)			
CSE Leak: Ves vs No	364	-13.06)	21		
Cor Leak. Tes vs No	304	(-15.48  to)	.21		
		3.40)			
House-Brackmann:	359	-35.34	<.001	-35.31	<.001
Score 3-6 versus		(-42.79 to		(-43.45 to	
Score 1–2		-27.88)		-27.17)	
Size (per 1 cm increase)	366	-6.62	<.001	-2.41	.08
		(-9.03 to		(-5.17 to	
		-4.21)		.36)	

Valid N = The number of patients used for the unadjusted estimates. The sample size for the adjusted estimates = 260. PANQOL face ranges from 0 to 100 with higher scores indicating greater quality of life.

#### 3.3.3. Balance

The mean score in the balance domain was 69/100 (SD 26). For every 1-year increase in age, PANQOL balance scores declined by approximately -.35 points (95% CI: -.56 to -.14; p = 0.001). Otherwise, PANQOL balance domain scores were not associated with remaining patient characteristics in this sample of data (Supplemental Table 2).

#### 3.3.4. Face

The mean score in the face domain was 78/100 (SD 25). Controlling

	Ν	Unadjusted		Adjusted	
		OR (95% CI)	р	OR (95% CI)	р
Age (per 1-year increase)	365	.99	.16		
		(.98 - 1.00)			
Sex: Male vs Female	365	.74	.11		
		(.51 - 1.08)			
Months since symptoms	365	.94 (.89–.99)	.01	.94	.06
(per 24-month increase)				(.90 - 1.00)	
Residual: Yes vs No	364	1.84	.02	1.42	.21
		(1.11 - 3.07)		(.82–2.44)	
Retreatment: Yes vs No	260	1.14	.76		
		(.50–2.59)			
Tinnitus	362		.002		.04
Only post-op versus no		4.54	.002	3.31	.02
post-op		(1.72 - 11.98)		(1.23 - 8.90)	
Pre-and-post op versus		1.70	.06	1.45	.20
no post-op		(.99–2.93)		(.83–2.53)	
Pre-and-post op versus		.37	.07	.44	.14
only post-op		(.13–1.09)		(.15 - 1.30)	
Balance	252		.33		
Only post-op versus no		1.37	.34		
post-op		(.71–2.65)			
Pre-and-post op versus		1.64	.21		
no post-op		(.76–3.56)			
Pre-and-post op versus		1.20	.71		
only post-op		(.46–3.08)			
Headache	359		.005		.02
Only post-op versus no		3.30	.002	2.84	.01
post-op		(1.57 - 6.97)		(1.32-6.14)	
Pre-and-post op versus		2.03	.30	1.59	.50
no post-op		(.54–7.72)		(.41–6.13)	
Pre-and-post op versus		.62	.53	.59	.46
only post-op		(.14–2.76)		(.12–2.57)	
Hearing Preservation:	93	1.04	.93		
Yes vs No		(.46–2.33)			
CSF Leak: Yes vs No	362	1.05	.88		
		(.53 - 2.10)			
House–Brackmann:	357	.74	.34		
Score 3-6 versus Score		(.40–1.37)			
1–2					
Size (per 1 cm increase)	364	.96	.68		
		(.80–1.16)			

N= The number of patients used for the estimates. The sample size for the adjusted estimates = 359. The PANQOL pain item asks participants to respond to the following item using a five-point ordinal scale (1 = Strong disagree to 5 = Strongly agree): "I have problems with head pain on the side of my acoustic neuroma tumor".

for tumor size, retreatment status, and residual status, patients with poor HB grades reported lower QOL scores in the face domain than patients with ideal HB grades ( $\mu$ diff = -35.31, 95% CI: -43.45 to -27.17; p < .001). Similarly, controlling for all other variables in the model, patients who underwent retreatment reported lower quality of life scores in the face domain ( $\mu$ diff = -20.34, 95% CI: -29.78 to -10.91; p < .001) (Table 3).

#### 3.3.5. Anxiety

The mean score in the anxiety domain was 80/100 (SD 25). Patients with a postoperative headache reported lower QOL in the PANQOL anxiety domain ( $\mu$ diff = -14.68, 95% CI: -26.98 to -2.37; *p* = 0.01). Otherwise, PANQOL anxiety domain scores were not associated with remaining patient characteristics in this sample of data (Supplemental Table 3).

# 3.3.6. Energy

The mean score in the energy domain was 68/100 (SD 26). Patients with a postoperative headache reported lower QOL in the PANQOL energy domain ( $\mu$ diff = -15.31, 95% CI: -28.33 to -2.29; p = 0.02). Conversely, males reported higher PANQOL energy scores ( $\mu$ diff = 5.60, 95% CI: .17 to 11.02; p = 0.04). Otherwise, PANQOL energy domain

#### Table 5

PANQOL health scores as a function of patient characteristics.

	Ν	Unadjusted		Adjusted		
		β (95% CI)	р	β (95% CI)	р	
Age (per 1-year increase)	366	–.34 (–.51 to –.16)	<.001			
Sex: Male vs Female	366	61 (-5.12 to 3.89)	.79			
Months since symptoms (per 1-month increase)	366	01 (04 to .01)	.27			
Residual: Yes vs No	365	2.27 (-3.90 to 8.44)	.47	1.76 (–4.92 to 8.44)	.60	
Retreatment: Yes vs No	261	-8.98 (-18.55 to .59)	.07	-9.22 (-19.07 to .64)	.07	
Tinnitus	363		.69			
Only post-op versus no post-op		3.53 (–10.52 to 17.58)	.82			
Pre-and-post op versus no post-op		-1.95 (-9.87 to 5.96)	.83			
Pre-and-post op versus only post-op		-5.48 (-21.06 to 10.09)	.69			
Balance	253		.82			
Only post-op versus no		.16 (-9.07	.99			
post-op		to 9.39)				
Pre-and-post op versus		2.96 (-8.00	.80			
no post-op		to 13.91)				
Pre-and-post op versus		2.80	.88			
only post-op		(-10.62 to 16.21)				
Headache	360		.001		.02	
Only post-op versus no post-op		-9.90 (-20.59 to .80)	.08	1.58 (–10.45 to 13.61)	.93	
Pre-and-post op versus no post-op		-25.15 (-44.47 to -5.83)	.01	-23.52 (-42.99 to -4.06)	.01	
Pre-and-post op versus only post-op		-15.25 (-36.98 to	.23	-25.10 (-47.13 to	.02	
Hearing Preservation:	93	2.98 (-6.10	.52	5.07 )		
Yes vs No		to 12.06)				
CSF Leak: Yes vs No	363	-1.08 (-9.39 to 7.23)	.80			
House–Brackmann: Score 3–6 versus Score 1–2	358	-2.85 (-10.05 to 4.34)	.44	-7.78 (-16.26 to .70)	.07	
Size (per 1 cm increase)	365	.83 (-1.36 to 3.03)	.45	2.44 (41 to 5.29)	.09	

N = The number of patients used for the estimates. The sample size for the adjusted estimates = 259. PANQOL health ranges from 0 to 100 with higher scores indicating greater QOL.

Table 6	
PANQOL scores by House Brackmann	groups.

House-Brackmann (recoded) Overall p Score 1 Score 2 Score 3 Scores 4 - 6 25% 25% 75% Mdn Ν Mdn 75% Ν Mdn N Mdn 25% 75% Ν 25% 75% PANQOL Hearing .91 **PANOOL Balance** .58 PANQOL Face <.001 **PANQOL Anxiety** .24 PANQOL Energy .28 PANOOL Pain .63 PANQOL Health .44 PANQOL Total .02

*Note*: Mdn = Median. 25% & 75% = Interquartile Range (IQR).

scores were not associated with remaining patient characteristics in this sample of data (Supplemental Table 4).

#### 3.3.7. Pain

26.8% (365) of patients that responded to the pain domain question indicated that they experienced residual pain on the surgical side. Controlling for residual status, tinnitus, and months since symptoms began, patients with solely a postoperative headache were 2.8 times more likely to report higher agreement with the PANQOL pain domain (95% CI: 1.32 to 6.14; p = 0.01). Controlling for all other variables in the model, patients with solely postoperative tinnitus were 3.3 times more likely to report higher agreement with the PANQOL pain item (95% CI: 1.23 to 8.90; p = 0.02) (Table 4).

# 3.3.8. Health

The mean score in the health domain was 69/100 (SD 22). Controlling for all other variables in the model, patients with both a pre- and postoperative headache reported lower PANQOL health QOL scores than those with no postoperative headache ( $\mu$ diff = -23.52, 95% CI: -42.99 to -4.06; *p* = 0.01) and those with solely a postoperative headache ( $\mu$ diff = -25.10, 95% CI: -47.13 to -3.07; *p* = 0.02). There was no significant difference in reported scores between patients with postoperative headaches and those with no postoperative headaches ( $\mu$ diff = 1.58, 95% CI: -10.45 to 13.61) (Table 5).

#### 3.4. PANQOL scores by House-Brackmann grade

There was significant variability in the PANQOL face and total scores when analyzed by HB grade. For the face domain, patients with HB grade I had a higher QOL (median = 100, IQR: 75–100) than patients with HB grade II (median = 58, IQR: 38–75), HB grade III (median = 42, IQR: 33–58; p < .001), and HB grade IV-VI (median = 42, IQR: 25–58; p < .001). No other pairwise comparisons were statistically significant. While there was overall variability among the four HB groups in the PANQOL total score (overall p = 0.02), no post-hoc pairwise comparison was statistically significant after adjusting for inflated Type 1 error (all p > 0.05) (Table 6).

#### 3.5. PANQOL scores by extent of resection

Ninety-one percent of patients received GTR (n = 334), the remaining receiving less than GTR for purpose of neurologic preservation in the setting of closely adherent tissue. Extent of resection was not associated with PANQOL total score ( $\mu$ diff = -1.91, 95% CI: -7.45 to 3.63, *p* =0.50), hearing score ( $\mu$ diff = -1.74, 95% CI: -8.55 to 5.07, *p* =0.62), balance score ( $\mu$ diff = -3.05, 95% CI: -10.48 to 4.38, *p* =0.42), face score ( $\mu$ diff = -.23, 95% CI: -6.68 to 6.21, *p* =0.94), anxiety score ( $\mu$ diff = -3.79, 95% CI: -10.85 to 3.28, *p* =0.29), energy score ( $\mu$ diff = -5.77, 95% CI: -13.24 to 1.69, *p* =0.13), pain score ( $\mu$ diff = 1.42, 95% CI: .82 to 2.44, *p* =0.06), or health score ( $\mu$ diff = 1.76, 95% CI: -4.92 to

# 8.44, *p* = 0.60).

# 3.6. PANQOL scores by retreatment

5.4% of patients required retreatment of their tumor (n = 20). Retreatment was significantly associated with decrease in PANQOL total score (µdiff = -10.11, 95% CI: -19.48 to -.74, p = 0.03) and face score (µdiff = -20.34, 95% CI: -29.78 to -10.91, p < 0.001). Retreatment was not associated with hearing score (µdiff = -3.95, 95% CI: -14.86 to 6.97, p = 0.48), balance score (µdiff = -9.76, 95% CI: -21.69 to 2.17, p = 0.11), anxiety score (µdiff = -5.75, 95% CI: -17.14 to 5.63, p = 0.32), energy score (µdiff = -7.89, 95% CI: -19.67 to 3.89, p = 0.19), pain score (µdiff = 1.14, 95% CI: .50 to 2.59, p = 0.76), or health score (µdiff = -9.22, 95% CI: -19.07 to .64, p = 0.07).

# 3.7. PANQOL scores by year of operation

From 1989 to 2004 (n = 133), the mean PANQOL score was 75.88. From 2005 to 2009 (n = 82) and 2010–2018 (n = 152), the mean PANQOL scores were 81.50 and 78.95, respectively.

#### 4. Discussion

While most studies utilizing PANQOL have assessed how treatment modality impacts QOL in patients with VS,<sup>13–21</sup> we sought to determine the specific baseline characteristics and surgical outcomes that influence the QOL of patients who undergo microsurgical resection. Expectedly, we identified that functional outcomes, such as hearing preservation and HB grade, had a significant impact on self-reported scores within the hearing and facial domains. Also as expected, scores within the facial domain between 1988 and 2004 were, on average, lower than the facial domain scores between both 2005–2009 and 2010–2018, given that the chance of a patient having a postoperative HB I score was higher during those times.<sup>24</sup> In concordance with prior literature, postoperative headache was associated with lower QOL scores in the anxiety, energy, pain, and health domains.<sup>15</sup> Total PANQOL scores were negatively influenced by poor HB grade, retreatment status, and female sex.

Our results demonstrate that extent of resection had no significant impact on PANQOL domain or overall scores in our patient population, which differs from a pivotal study by Link et al who showed that greater extent of resection is associated with improved self-reported QOL. Their study analyzed long term QOL between 143 patients who received either GTR (85%) or less than GTR (15%) from microsurgical removal of VS, demonstrating that those receiving GTR reported better facial, energy, health, and total PANQOL scores than those receiving less than GTR.<sup>28</sup> While Link et al reported on extent of resection, the effect of the need for retreatment on QOL was not evaluated.

Significantly, we found that when controlling for extent of resection and HB grade, retreatment was significantly associated with lower face domain and total PANQOL scores. Thus, we suggest that retreatment impacts QOL while extent of resection does not. This idea builds upon suggestions by Link et al that there is a psychological component to overall QOL outcomes among patients undergoing microsurgery, and there may be an interplay between preoperative expectations, patient perception of residual tumor, and requirement of retreatment influencing satisfaction and QOL despite functional outcomes.<sup>28</sup> Although Link et al demonstrated that extent of resection was positively associated with improved QOL, differences in study design may explain our varying results. Our present study consisted of a larger patient population (367 patients vs. 143 patients) and longer mean period between surgery and PANQOL survey (11.4 years vs. 7.7 years), which may elucidate trends in patient perception of importance of extent of resection on QOL over time.

The approach to VS resection remains variable among the neurosurgical and otolaryngologic communities. Some surgeons approach VS management with tumor debulking followed by planned stereotactic

radiosurgery, and others plan for complete resection with willingness to sacrifice neurological function.<sup>10</sup> Our previous study looking specifically at surgical approach and PANQOL scores showed that patients who had the retrosigmoid approach have higher PANQOL scores than those who underwent the translabyrinthine approach.<sup>29</sup> The two senior authors (DEA and JPL) aim for GTR while recognizing that a small amount of residual tumor as a means of functional preservation is acceptable. Inclusion of various tumor sizes in this analysis highlights a potential difference in expectations between patients with small and large tumors; patients with larger tumors have a trend towards reporting improved PANQOL health and face scores, although these were not statistically significant. Reasonably, we agree with Link et al in their proposition that patients who elect to have microsurgery for VS less than 3 cm may be psychologically biased to expect to have their tumors completely removed, thus are less satisfied when discovering residual tumor was left.<sup>27</sup>

Limitations of this study include the retrospective nature of chart review, in addition to self-report bias on PANQOL. Additionally, this study was performed at a single center by one interdisciplinary team, which may limit generalizability. Our response rate was 52%, with a lower response rate in earlier years, which may contribute significant response bias, and was a notably lower response rate than other PAN-QOL studies.<sup>15,28</sup> Some patients included comments on their surveys indicating that they were unsure whether problems indicated by the survey were due to their VS or due to aging and other health conditions. Furthermore, due to the wide time range of this study, there were minor differences in reporting and data collection between patients with paper versus electronic medical records. It is also important to note that there have been advancements in technology, intraoperative monitoring, and surgical techniques over the time range of this study; however, there are numerous studies examining the effect of these advancements on the variables measured in our study, such as facial and hearing preservation and extent of resection. Therefore, one can draw obvious conclusions about the effect of these advancements on our results.

# 5. Conclusion

Our results demonstrate that patients who undergo surgical retreatment of VS following initial microsurgical intervention have lower patient reported QOL than those who do not. In contrast to previously reported studies, we found that extent of resection does not impact QOL. Additionally, poor HB grade and female sex were negatively associated with total PANQOL scores, while postoperative headache was negatively associated with anxiety, energy, pain, and health domain scores. These results suggest that technological advancements should continue to focus on maximizing facial and cochlear nerve preservation and minimizing the chances of reoperation and postoperative headache. Furthermore, stratification of HB grades revealed a significant decrease in PANQOL face domain scores between HB grade I, II, and III, suggesting that favorable facial function outcomes definitions should be revisited.

# CRediT authorship contribution statement

Loren N. Riedy: Conceptualization, Data curation, Writing – original draft. Rachyl M. Shanker: Conceptualization, Writing – original draft. Dayna C. Sloane: Writing – review & editing. Daniel M. Heiferman: Conceptualization, Writing – review & editing. Elhaum G. Rezaii: Data curation, Writing – review & editing. Sarah E. Finucane: Data curation. Perry Veras: Data curation. John P. Leonetti: Supervision. Douglas E. Anderson: Conceptualization, Supervision.

#### Declaration of competing interest

The authors do not have any disclosures.

#### Acknowledgments

We thank Dr. William Adams (Loyola University Chicago) for performing the statistical analysis.

# Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10.1016/j.wnsx.2024.100294.

# References

- Babu R, Sharma R, Bagley JH, Hatef J, Friedman AH, Adamson C. Vestibular schwannomas in the modern era: epidemiology, treatment trends, and disparities in management: clinical article. J Neurosurg. 2013;119(1):121–130. https://doi.org/ 10.3171/2013.1.JNS121370.
- Ostrom QT, Gittleman H, Liao P, et al. CBTRUS Statistical Report: primary brain and other central nervous system tumors diagnosed in the United States in 2010–2014. *Neuro Oncol.* 2017;19(suppl\_5):v1-v88. https://doi.org/10.1093/neuonc/nox158.
- Brackmann DE. Acoustic neuroma: surgical approaches and complications. Ann Acad Med Singapore, 1991;20(5):674–679.
- Goldbrunner R, Weller M, Regis J, et al. EANO guideline on the diagnosis and treatment of vestibular schwannoma. *Neuro Oncol.* 2020;22(1):31–45. https://doi. org/10.1093/neuonc/noz153.
- Samii M, Gerganov VM, Samii A. Functional outcome after complete surgical removal of giant vestibular schwannomas: clinical article. *J Neurosurg.* 2010;112(4): 860–867. https://doi.org/10.3171/2009.7.JNS0989.
- Ahn J, Ryu NG, Lim J, Kang M, Seol HJ, Cho YS. Prognostic factors of facial nerve function after vestibular schwannoma removal via translabyrinthine approach. *Acta Otolaryngol.* 2019;139(6):541–546. https://doi.org/10.1080/ 00016489.2019.1592223.
- Falcioni M, Fois P, Taibah A, Sanna M. Facial nerve function after vestibular schwannoma surgery. J Neurosurg. 2011;115(4):820–826. https://doi.org/10.3171/ 2011.5.JNS101597.
- Anderson DE, Leonetti J, Wind JJ, Cribari D, Fahey K. Resection of large vestibular schwannomas: facial nerve preservation in the context of surgical approach and patient-assessed outcome. *J Neurosurg.* 2005;102(4):643–649. https://doi.org/ 10.3171/jns.2005.102.4.0643.
- Gurgel RK, Dogru S, Amdur RL, Monfared A. Facial nerve outcomes after surgery for large vestibular schwannomas: do surgical approach and extent of resection matter? *Neurosurg Focus*. 2012;33(3):E16. https://doi.org/10.3171/2012.7.FOCUS12199.
- Sughrue ME, Kaur R, Rutkowski MJ, et al. Extent of resection and the long-term durability of vestibular schwannoma surgery: clinical article. *J Neurosurg*. 2011;114 (5):1218–1223. https://doi.org/10.3171/2010.11.JNS10257.
- Bloch O, Sughrue ME, Kaur R, et al. Factors associated with preservation of facial nerve function after surgical resection of vestibular schwannoma. *J Neuro Oncol.* 2011;102(2):281–286. https://doi.org/10.1007/s11060-010-0315-5.
- Shaffer BT, Cohen MS, Bigelow DC, Ruckenstein MJ. Validation of a disease-specific quality-of-life instrument for acoustic neuroma. *Laryngoscope*. 2010;120(8): 1646–1654. https://doi.org/10.1002/lary.20988.
- Carlson ML, Tveiten ØV, Yost KJ, Lohse CM, Lund-Johansen M, Link MJ. The minimal clinically important difference in vestibular schwannoma quality-of-life assessment: an important step beyond P < .05. Otolaryngol-Head Neck Surg Off J Am Acad Otolaryngol-Head Neck Surg. 2015;153(2):202–208. https://doi.org/10.1177/ 0194599815585508.
- Kerezoudis P, Yost KJ, Tombers NM, Celda MP, Carlson ML, Link MJ. Defining the minimal clinically important difference for patients with vestibular schwannoma: are all quality-of-life scores significant? *Neurosurgery*. 2019;85(6):779–785. https:// doi.org/10.1093/neuros/nyy467.
- Carlson ML, Tveiten ØV, Driscoll CL, et al. What drives quality of life in patients with sporadic vestibular schwannoma? *Laryngoscope*. 2015;125(7):1697–1702. https://doi.org/10.1002/lary.25110.

- Carlson ML, Tveiten OV, Driscoll CL, et al. Long-term quality of life in patients with vestibular schwannoma: an international multicenter cross-sectional study comparing microsurgery, stereotactic radiosurgery, observation, and nontumor controls. J Neurosurg. 2015;122(4):833–842. https://doi.org/10.3171/2014.11. JNS14594.
- Carlson ML, Tveiten ØV, Driscoll CL, et al. Risk factors and analysis of long-term headache in sporadic vestibular schwannoma: a multicenter cross-sectional study. *J Neurosurg.* 2015;123(5):1276–1286. https://doi.org/10.3171/2014.12. JNS142109.
- Carlson ML, Barnes JH, Nassiri A, et al. Prospective study of disease-specific qualityof-life in sporadic vestibular schwannoma comparing observation, radiosurgery, and microsurgery. Otol Neurotol Off Publ Am Otol Soc Am Neurotol Soc Eur Acad Otol Neurotol. 2021;42(2):e199–e208. https://doi.org/10.1097/ MA0.00000000002863.
- Miller LE, Brant JA, Naples JG, Bigelow DC, Lee JYK, Ruckenstein MJ. Quality of life in vestibular schwannoma patients: a longitudinal study. Otol Neurotol Off Publ Am Otol Soc Am Neurotol Soc Eur Acad Otol Neurotol. 2020;41(2):e256–e261. https://doi. org/10.1097/MAO.0000000002445.
- Chweya CM, Tombers NM, Lohse CM, Link MJ, Carlson ML. Disease-specific quality of life in vestibular schwannoma: a national cross-sectional study comparing microsurgery, radiosurgery, and observation. Otolaryngol-Head Neck Surg Off J Am Acad Otolaryngol-Head Neck Surg. 2021;164(3):639–644. https://doi.org/10.1177/ 0194559820941012.
- Pruijn IMJ, Kievit W, Hentschel MA, Mulder JJS, Kunst HPM. What determines quality of life in patients with vestibular schwannoma? *Clin Otolaryngol Off J ENT-UK Off J Neth Soc Oto-Rhino-Laryngol Cervico-Facial Surg.* 2021;46(2):412–420. https://doi.org/10.1111/coa.13691.
- Gardner G, Robertson JH. Hearing preservation in unilateral acoustic neuroma surgery. Ann Otol Rhinol Laryngol. 1988;97(1):55–66. https://doi.org/10.1177/ 000348948809700110.
- House JW, Brackmann DE. Facial nerve grading system. Otolaryngol Neck Surg. 1985; 93(2):146–147. https://doi.org/10.1177/019459988509300202.
- Heiferman DM, Riedy LN, Rezaii EG, et al. Is there a plateau to the learning curve for acoustic neuroma resection?—experience and outcomes from a single interdisciplinary team over thirty years. J Neurol Surg B Skull Base. 2023. https:// doi.org/10.1055/a-1993-7906.
- Dwass M. Some K-Sample Rank-Order Tests. Stanf Univ Press Calif, 1960:198–202. Published online.
- Steel RGD. A rank sum test for comparing all pairs of treatments. *Technometrics*. 1960;2(2):197–207. https://doi.org/10.1080/00401706.1960.10489894.
- Douglas CE, Michael FA. On distribution-free multiple comparisons in the one-way analysis of variance. *Commun Stat Theor Methods*. 1991;20(1):127–139. https://doi. org/10.1080/03610929108830487.
- Link MJ, Lund-Johansen M, Lohse CM, et al. Quality of life in patients with vestibular schwannomas following gross total or less than gross total microsurgical resection: should we be taking the entire tumor out? *Neurosurgery*. 2018;82(4): 541–547. https://doi.org/10.1093/neuros/nyx245.
- North M, Weishaar J, Nuru M, Anderson D, Leonetti JP. Assessing surgical approaches for acoustic neuroma resection: do patients perceive a difference in quality-of-life outcomes? *Otol Neurotol.* 2022;43(10):1245–1251. https://doi.org/ 10.1097/MAO.00000000003720.

# Abbreviation List

- AIC: Akaike's Information Criterion
- GR: Gardner-Robertson hearing scale

GTR: gross total resection

HB: House-Brackmann

IQR: interquartile range PANQOL: Penn Acoustic Neuroma Quality of Life

PTA: pure tone average

QOL .: quality of life

VS: vestibular schwannoma