

CONGRESS OF NEUROLOGICAL SURGEONS SYSTEMATIC REVIEW AND EVIDENCE-BASED GUIDELINE ON INTRAOPERATIVE CRANIAL NERVE MONITORING IN VESTIBULAR SCHWANNOMA SURGERY

Sponsored by: Congress of Neurological Surgeons (CNS) and the Section on Tumors

Endorsed by: Joint Guidelines Committee of the American Association of Neurological Surgeons (AANS) and the Congress of Neurological Surgeons (CNS)

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Abbreviations

ABR: Auditory brainstem response CMAP: Compound muscle action potential DENM: Direct eighth cranial nerve monitoring EMG: Electromyogram FN: Facial nerve GR: Gardner-Robertson facial function grading system HB: House-Brackmann facial function grading system ICNM: Intraoperative cranial nerve monitoring NF: Neurofibromatosis **PPV:** Positive predictive value PTA: Pure tone average SMS: Supramaximal stimulation SRS: Speech recognition score SRT: Speech reception threshold VS: Vestibular schwannoma WRS: Word recognition score

ABSTRACT

Facial Nerve Monitoring

Question 1

Does intraoperative facial nerve monitoring during vestibular schwannoma surgery lead to better long-term facial nerve function?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery regardless of tumor characteristics.

Recommendation

Level 3: It is recommended that intraoperative facial nerve monitoring be routinely utilized during vestibular schwannoma surgery to improve long-term facial nerve function.

Question 2

Can intraoperative facial nerve monitoring be used to accurately predict favorable long-term facial nerve function after vestibular schwannoma surgery?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery.

Recommendation

Level 3: Intraoperative facial nerve monitoring can be used to accurately predict favorable longterm facial nerve function after vestibular schwannoma surgery. Specifically, the presence of favorable testing reliably portends a good long-term facial nerve outcome. However, the absence of favorable testing in the setting of an anatomically intact facial nerve does not reliably predict poor long-term function and therefore cannot be used to direct decision-making regarding need for early reinnervation procedures.

Question 3

Does an anatomically intact facial nerve with poor electromyogram electrical responses during intraoperative testing reliably predict poor long-term facial nerve function?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery.

Recommendation

Level 3: Poor intraoperative electromyogram electrical response of the facial nerve should not be used as a reliable predictor of poor long-term facial nerve function.

Cochlear Nerve Monitoring

Question 4

Should intraoperative eighth cranial nerve monitoring be used during vestibular schwannoma surgery?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery with measurable preoperative hearing levels and tumors smaller than 1.5 cm.

Recommendation

Level 3: Intraoperative eighth cranial nerve monitoring should be used during vestibular schwannoma surgery when hearing preservation is attempted.

Question 5

Is direct monitoring of the eighth cranial nerve superior to the use of far-field auditory brain stem responses?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery with measurable preoperative hearing levels and tumors smaller than 1.5 cm.

Recommendation

Level 3: There is insufficient evidence to make a definitive recommendation.

1 INTRODUCTION

2 Rationale

3 The surgical management of VSs has experienced a significant evolution since its 4 inception by Harvey Cushing, MD, and other early pioneering surgeons of the late 5 nineteenth and early twentieth centuries. In the 1960s, further progress was made with the 6 implementation of the operating microscope and surgical drill, the use of which is largely 7 credited to William House, MD. The coadvancement of surgical techniques and 8 technology has led to a significant decline in the morbidity and mortality profile of VS 9 surgery. Today, the neurologic deficits once considered acceptable sequelae are no longer 10 commonplace, and mortality from surgery is reported at <1% when performed by 11 experienced surgical teams.¹ 12

13 Early on, facial paralysis and deafness were thought to be inevitable and acceptable 14 consequences of tumor resection, particularly because most patients were diagnosed with 15 large and often life-threatening tumors. In today's practice, however, the expectation is to 16 preserve facial function in the vast majority of cases. As a result, patient quality of life 17 following VS surgery has improved, and the sequelae of ophthalmologic complications 18 and the need for invasive dynamic facial rehabilitation procedures have been reduced. A 19 systematic review of the literature published in 2010 found an overall microsurgical FN preservation rate of 74%.² The value placed in preserving seventh cranial nerve 20 21 functional integrity is high and can motivate subtotal resection in select large tumors, 22 with or without the use of postoperative radiation therapy.

23

24 Hearing preservation surgery is the latest chapter in the evolution of VS management. 25 The advent and widespread availability of contrast-enhanced magnetic resonance 26 imaging has allowed for earlier diagnosis, producing a population of patients with smaller 27 tumors and better baseline hearing. Patients with small- to medium-sized tumors and 28 serviceable hearing are now being offered hearing preservation surgery at higher rates 29 than ever before. Whereas the translabyrinthine approach commits the patient to 30 permanent ipsilateral deafness, the retrosigmoid and middle fossa approaches offer 31 opportunities to preserve acoustic function in select tumors. Currently, postoperative FN

32 function and hearing preservation are 2 primary benchmarks consistently reported by

33 high-volume VS surgical centers. These 2 measures have been enhanced significantly by

34 improvements in surgical technique and the development and refinement of

35 intraoperative cranial nerve monitoring (ICNM).

36

37 Delgado et al³ introduced ICNM of the facial nerve in the late 1970s, which has become a 38 mainstay for most VS surgeons.² A consensus statement published by the National Institutes of Health in the 1990s recommended the routine use of neuromonitoring during 39 40 VS surgery.⁴ The existing literature on this subject primarily consists of large 41 retrospective case series from high-volume surgical centers – large prospective 42 comparative studies are generally lacking. Furthermore, heterogeneous reporting and the 43 use of inconsistent electroprognostic testing parameters are variable, rendering interstudy comparisons challenging. Recently these parameters have garnered a more elaborate role. 44 45 Whereas the initial role of ICNM was for the identification and intraoperative mapping of 46 the FN, there is a new focus on electrical factors that could potentially serve as 47 electroprognostic indicators of long-term facial function. The utility of testing in this 48 manner may have a profound impact on how to counsel patients with immediate 49 postoperative paresis and an anatomically preserved FN. In addition, it would offer the 50 treating physician an objective basis for proceeding with watchful waiting and 51 conservative measures versus a recommendation that a patient undergo early dynamic 52 facial reanimation procedures. An example would be timing VS surgery with 53 hypoglossal-facial anastomosis (where an earlier intervention is associated with improved 54 functional outcomes), as opposed to enrolling the patient into an observation period for 55 spontaneous recovery that can last anywhere from 12 to 18 months. 56

In contrast to facial nerve monitoring, the role of ICNM for hearing preservation is less well defined and is not uniformly used. This may be the result of a more technically challenging and cumbersome process than what is required with FN monitoring. It may also have to do with differences in treatment philosophy for smaller tumors between surgeons and between centers.

- 63 There is currently a need to assess the existing literature for VS surgery outcomes,
- 64 specifically as it relates to the use of ICNM and its impact on postoperative FN function
- and hearing preservation.

66 **Objectives**

The objective of this systematic review is to critically assess the existing literature and
provide an evidence-based clinical practice guideline regarding the use of ICNM during
VS surgery. Specifically, this systematic review focuses on intraoperative monitoring
techniques and eletroprognostic parameters as they relate to posttreatment function of the
seventh and eighth cranial nerves.

72 **METHODS**

73 Process Overview

- 74 The evidence-based clinical practice guideline taskforce members and the Tumor Section
- 75 of the American Association of Neurological Surgeons and the Congress of Neurological
- 76 Surgeons (CNS) conducted a systematic review of the literature relevant to the
- 77 management of VSs. The PubMed, Embase, and Web of Science databases were queried.
- 78 The keywords used during our search of the medical literature databases cited above are
- 79 documented in Tables 1 and 2. Additional details of the systematic review are provided
- 80 below and within the introduction and methodology chapter of the guideline
- 81 (https://www.cns.org/guidelines/guidelines-management-patients-vestibular-
- 82 <u>schwannoma/chapter_1</u>).

83 Article Inclusion/Exclusion Criteria

84 Citations were manually reviewed by the team with specific inclusion and exclusion

- 85 criteria as outlined below. The duplicates from the search were eliminated. Two
- 86 independent reviewers reviewed and abstracted full-text data for each article, and the 2
- 87 sets of data were compared for agreement by a third party. Inconsistencies were re-
- 88 reviewed and disagreements were resolved by consensus. The evolution of the article
- 89 selection is illustrated with flow diagrams (Figures 1 and 2). All citations that focused on
- 90 adult patients and surgical treatment of VSs were broadly considered. For literature to be
- 91 included for further consideration, papers had to meet the following criteria:

92	General	
93	•	Investigated patients suspected of having vestibular schwannomas
94	•	Was of humans
95	•	Was not an in vitro study
96	•	Was not a biomechanical study
97	•	Was not performed on cadavers
98	•	Published between January 1, 1990 and December 31, 2014
99	•	Published in a peer-reviewed journal
100	•	Was not a meeting abstract, editorial, letter, or commentary
101	•	Was published in English
102	•	Included quantitatively presented results
103		
104	Specific	
105	•	Used an established FN function grading system, such as the House-
106		Brackmann (HB) ⁵ scale or the Sunnybrook (SB) ⁶ scale.
107	•	Used the 1995 American Academy of Otolaryngology-Head and Neck
108		Surgery (AAO-HNS) ⁷ or Gardner–Robertson (GR) ⁸ hearing classification
109		system OR presented data using word recognition score (WRS) and pure tone
110		average (PTA) for defining hearing status or had individual patient data
111		presented such that the latter criteria could be applied and analyzed
112	•	Included pre- and postoperative audiometric data
113	•	Included a median or mean follow-up of 12 months following treatment when
114		assessing long-term facial outcomes
115	•	Included only studies evaluating intraoperative electrophysiological testing of
116		the facial and cochlear nerves
117	•	Used electrically evoked testing with EMG
118	٠	NF status was collected when available but was not an exclusion criterion
119		
120	The au	thors did not include systematic reviews, guidelines, or meta-analyses
121	conduc	ted by others. These documents were developed using different inclusion

122 criteria than those specified in our guideline. Therefore, they may have included

studies that do not meet the inclusion criteria listed above. These documents were

- 124 recalled if their abstract suggested that they might address one of the recommendations
- set forth in this guideline. The authors searched their bibliographies for additional
- studies.

127 Search Strategies

The task force collaborated with a medical librarian to search for articles published between January 1, 1990 and December 31, 2014. Three electronic databases were searched: PubMed, EMBASE, and Web of Science. Strategies for searching electronic databases were constructed by the evidence-based clinical practice guideline taskforce members and the medical librarian using previously published search strategies to identify relevant studies (Tables 1 and 2).

134

135 Searches of electronic databases were supplemented with manual screening of the

136 bibliographies of all retrieved publications. Bibliographies of recent systematic reviews

137 and other review articles for potentially relevant citations were also searched. All articles

- 138 identified were subject to the study selection criteria listed above. The guideline
- 139 committee also examined lists of included and excluded studies for errors and omissions.

140 The guideline task force went to great lengths to obtain a complete set of relevant articles

141 to ensure guideline recommendations are not based on a biased subset of articles. Two

142 datasets were constructed, one for FN monitoring and another for cochlear nerve

143 monitoring.

144 Facial Nerve Monitoring

145 The search of the 3 above mentioned databases yielded a total of 2853 candidate articles.

146 One thousand nine hundred and eighty-four remained after duplicates were removed and

- 147 date range criteria were applied. The abstracts were reviewed, and after the
- 148 aforementioned general and specific inclusion/exclusion criteria were applied, 21 articles

149 remained and were included in the final analysis (Table 1, Figure 1).

150

151 Cochlear Nerve Monitoring

- 152 The search of the 3 abovementioned databases yielded a total of 1849 articles. Eight
- 153 hundred and three remained after duplicates were removed and date range criteria
- applied. The abstracts were reviewed, and after the aforementioned general and specific
- 155 exclusion criteria were applied, 7 articles remained and were included in the final
- 156 analysis (Table 2, Figure 2).

157 Data Analysis

- 158 Evidence tables for the use of intraoperative cochlear nerve monitoring and FN
- 159 monitoring were constructed using key study parameters as outlined above.
- 160
- 161 Facial Nerve: Data extraction included study design, level of evidence, total number of
- 162 patients, pre- and posttreatment facial function, study selection parameters, tumor
- 163 characteristics, mean or median follow-up, neurofibromatosis type 2 status, and
- 164 prognostic parameters associated with short- and long-term facial function.
- 165
- 166 *Cochlear Nerve:* Data extraction included study design, level of evidence, total number
- 167 of patients, pre- and posttreatment hearing status, study selection parameters, tumor
- 168 characteristics, mean or median follow-up, neurofibromatosis type 2 status, and
- 169 prognostic features associated with postoperative hearing preservation.

170 Classification of Evidence and Guideline Formulation

- 171 The concept of linking evidence to recommendations has been further formalized by the
- 172 American Medical Association (AMA) and many specialty societies, including the
- 173 American Association of Neurological Surgeons (AANS), the CNS, and the American
- 174 Academy of Neurology (AAN). This formalization involves the designation of specific
- 175 relationships between the strength of evidence and the strength of recommendations to
- avoid ambiguity. In the paradigm for prognostication used in this guideline, evidence is
- 177 classified into 1 of 3 tiers based on the degree at which the study fulfills the 5 technical178 criteria listed below:
- 179 180
 - Was a well-defined representative sample of patients assembled at a common (usually early) point in the course of their disease?
- Was patient follow-up sufficiently long and complete?

182	• Were objective outcome criteria applied in a "blinded" fashion?
183	• If subgroups with different prognoses were identified, was there adjustment for
184	important prognostic factors?
185	• If specific prognostic factors were identified, was there validation in an
186	independent "test set" group of patients?
187	
188	Class I evidence is used to support recommendations of the strongest type, defined as
189	Level 1 recommendations, and require that all 5 technical criteria are satisfied. Class II
190	evidence supports intermediate strength recommendations, defined as level 2
191	recommendations, and requires that 4 of the 5 technical criteria be met. Class III evidence
192	supports Level 3 recommendations, comprising all remaining studies that satisfy ≤ 3 of
193	the 5 technical criteria. A basis for these guidelines can be viewed in Haines SJ and
194	Nicholas JS (2006). Evidence-Based Medicine: A Conceptual Framework. In Haines SJ
195	and Walters BC (Eds.), Evidence-Based Neurosurgery: An Introduction (Pages 1-17).
196	New York: Thieme Medical Publishers.

197 **RESULTS**

198 FACIAL NERVE MONITORING

Question 1

Does intraoperative facial nerve monitoring during vestibular schwannoma surgery lead to better long-term facial function?

Target population

This recommendation applies to all adult patients undergoing vestibular schwannoma surgery regardless of tumor characteristics.

Recommendation

Level 3: It is recommended that intraoperative facial nerve monitoring be routinely utilized during vestibular schwannoma surgery to improve long-term facial nerve function.

199 STUDY SELECTION AND CHARACTERISTICS

200 A total of 2853 candidate studies were screened and assessed for eligibility per the

201 previous criteria, and 21 studies were included in the final review.^{9–29} Postoperative FN

- 202 function with the use of intraoperative electrically evoked testing with EMG versus facial 203 function in unmonitored surgery was the basis of the recommendations in this section. To 204 be included as a part of this recommendation, a study had to provide a cohort of patients 205 with assessment of pre- and postoperative FN function using an established FN function 206 grading system, such as the HB scale or the SB scale. Furthermore, the method of 207 intraoperative FN monitoring had to be clearly delineated with a comparison between 208 monitored and unmonitored cohorts. Using these criteria, a final total of 3 studies were included for analysis (Table 3A).^{14,16,25} 209
- 210

211 In cases where an authoring center published multiple papers that met these criteria, only

the study with the largest number of subject patients was used to avoid duplicate

213 reporting of patient data if the patient recruitment dates overlapped. Data extraction

214 included study design, level of evidence, number of patients, tumor characteristics,

215 method of ICNM, and long-term FN function.

216 **RESULTS OF INDIVIDUAL STUDIES, DISCUSSION OF STUDY**

217 LIMITATIONS, AND RISK OF BIAS

218 Three studies met the inclusion criteria for this recommendation.^{14,16,25} All 3 studies

219 represent Class III data, primarily due to lack of blinded assessment and the absence of a

220 validation set. The key results of individual studies are outlined in evidence Table 3A and

are summarized within the guideline recommendations. All 3 studies performed a

retrospective analysis of postoperative FN function between unmonitored and monitoredcohorts.

224

In 1994, Lenarz and Ernst¹⁶ performed a retrospective review of 64 VS patients who

226 underwent microsurgical resection by the same surgeon at a single institution between

1986 and 1991. The goal of the study was to compare postoperative facial function

between monitored (n = 30) and unmonitored groups (n = 34). The 2 groups were

229 comparable with respect to tumor size, surgical time, and surgical approach (middle fossa

230 or translabyrinthine). ICNM consisted of facial muscle EMG via needle electrodes, and

- 231 electrical stimulation of the nerve was performed with bipolar forceps using constant
- current pulses of 100 microseconds (µs) and current strength between 0.05 and 0.8

233 milliamps (mA). The average tumor size in the monitored group was $1.5 \text{ cm} (\pm 0.5 \text{ cm})$ 234 and in the unmonitored group was $1.7 \text{ cm} (\pm 0.7 \text{ cm})$. They could correlate intraoperative 235 tonic (train) activity per hour of surgery, as well as postresection threshold current with 236 immediate postoperative facial function. An increase in train activity and an increase in 237 threshold current (mA) with decreasing wave amplitude at the end of the case correlated 238 with worse immediate facial function. The lack of intraoperative stimulation at the end of 239 the case was predictive of a complete immediate postoperative facial paralysis. The use 240 of monitoring improved immediate and long-term FN outcomes (P < .05). This was 241 especially true in tumors >1.5 cm: HB grade I to II at 1 year 87% (monitored) versus 74% 242 (unmonitored); grade III to VI at 1 year 13% (monitored) versus 26% (unmonitored).

243

In 1993, Silverstein²⁵ performed a retrospective analysis of 121 VS patients who 244 245 underwent microsurgical resection by the same surgeon at a single center between 1974 246 and 1991. Postoperative facial function was assessed immediately and at >1 year in 247 monitored (n = 65) and unmonitored cases (n = 56). Surgery consisted of retrosigmoid 248 and translabyrinthine approaches. EMG facial monitoring was applied using various 249 techniques over the course of the study, in line with advancement in software and 250 hardware developments. Electrical stimulation of the nerve was performed with insulated 251 stimulator probe tips and insulated micro instruments. Electrical pulsed currents ranged 252 from 0.05 to 3 mA. Facial function results were reported for the entire cohort and 253 subanalyzed by surgical approach. Subgroup analysis for surgical approach found no 254 statistical difference between the monitored and unmonitored groups. A distinction was 255 made between the translabyrinthine group, subtotal versus total resection. The authors 256 found statistically worse function after total tumor resection via the translabyrinthine 257 approach when compared to subtotal resection via the translabyrinthine approach or 258 retrosigmoid approach. There were more patients with the FN transected at surgery in the 259 unmonitored group (P < .05). Analysis of the entire cohort found that patients had 260 statistically better facial function in the monitored group than in the unmonitored group 261 (P < .02) immediately and at 1-year follow-up. Assessment of both monitored and 262 unmonitored groups found that large tumors (>3 cm) had poorer FN outcomes when 263 compared to small (<1.5 cm) or medium-sized (1.5–3 cm) tumors (P < .01). This study

covers a large span of time (17 years). Improvements in monitoring technology along
with increased surgeon expertise over the time span contribute bias to the analysis. In
addition, the authors report a recent trend to perform subtotal resection in larger tumors in
efforts to preserve the anatomical integrity of the nerve.

268

In 1991, Kwartler¹⁴ performed a retrospective analysis of 244 VS patients who underwent

270 microsurgical resection at a single institution between 1986 and 1987. All patients had

tumor resection via the translabyrinthine approach. Eighty-nine patients were monitored,

and 155 patients were unmonitored. EMG was measured using bipolar hookwire

273 electrodes in the facial musculature and direct electrical stimulation using a monopolar

probe with constant-current stimulus from 0.05 to 3 mA. Monitored patients had a

statistically significant better FN outcome in the perioperative period; however, this

advantage was not seen at 1 year of follow-up. Subanalysis performed with tumor size

found worse facial function in tumors >2.5 cm (P < .01).

278 SYNTHESIS OF RESULTS

279 Level 3 data suggests the use of ICNM of the FN during VS surgery leads to better facial

280 function outcomes. The 3 studies assessed postoperative facial function in patients

- 281 undergoing microsurgical resection of VSs with or without use of ICNM. Electrical
- stimulation offers the ability to help localize and map the course of the FN and may alert

the surgeon to stretch injury by way of eliciting train or tonic activity. Larger tumors had

- an overall worse prognosis for postoperative FN function even with use of FN
- 285 monitoring. Increased train or tonic activity along with elevated threshold currents
- 286 following tumor resection were poor prognostic indicators for postoperative FN function.

287 **DISCUSSION**

- 288 The benefits of ICNM in VS surgery has been widely reported over the last few decades,
- and again, supported by this analysis. Interestingly, the 3 studies used in this
- 290 recommendation were published in the 1990s. This reflects the paucity of surgical

- 291 literature providing direct comparison between monitored and unmonitored surgeries due
- to the now common use of ICNM during VS tumor resection.

Question 2

Can intraoperative facial nerve monitoring be used to accurately predict favorable long-term facial nerve function after vestibular schwannoma surgery?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery.

Recommendation

Level 3: Intraoperative facial nerve monitoring can be used to accurately predict favorable long-term facial nerve function after vestibular schwannoma surgery. Specifically, the presence of favorable testing reliably portends a good long-term facial nerve outcome. However, the absence of favorable testing in the setting of an anatomically intact facial nerve does not reliably predict poor long-term function and therefore cannot be used to direct decision-making regarding need for early reinneravation procedures.

293 STUDY SELECTION AND CHARACTERISTICS

294 A total of 2853 candidate studies were screened and assessed for eligibility per the previous criterion and 21 studies were included in the final review.⁹⁻²⁹ The 295 296 electroprognostic value of ICNM in determining good long-term postoperative facial 297 function was the basis of the recommendation in this section. To be included as a part of 298 this recommendation, a study had to provide a cohort of patients with assessment of pre-299 and postoperative FN function using an established FN function grading system, such as 300 the HB scale or the SB scale. In addition, the method of intraoperative FN monitoring 301 had to be described and a minimum of 1-year follow-up was required for determination 302 of long-term outcomes. There were 15 studies that met the inclusion criteria for this recommendation.^{9,10,12,17–24,26–29} In cases where an authoring center published multiple 303 304 papers that met these criteria, only the study with the largest number of subject patients 305 was used to avoid duplicate reporting of patient data if the patient recruitment dates 306 overlapped. Data extraction included study design, level of evidence, number of patients, 307 method of ICNM, electrical characteristics of the ICNM that correlated with 308 postoperative facial function, and assessment of FN function at ≥ 1 year postoperatively.

309 RESULTS OF INDIVIDUAL STUDIES, DISCUSSION OF STUDY

310 LIMITATIONS, AND RISK OF BIAS

311 All 15 studies represent Class III data, primarily due to the lack of blinded assessment 312 and the absence of a validation set. The key results of individual studies are outlined in 313 evidence Table 3B and are summarized within the guideline recommendations. Of the 15 studies used in this analysis. 12 studies^{9,10,17–21,23,24,26–28} identified intraoperative 314 315 eletrophysiologic parameters that were predictive of "good" postoperative facial function 316 at ≥ 1 year. Good function in this analysis was defined as HB I-II. Heterogeneous 317 electroprognostic parameters were used between studies; however, all authors provided 318 details on the criteria applied for their assessments. All the ICNM techniques made use of 319 continuous or electrically evoked EMG activity.

320

321 In 2013, Schmitt et al²³ described a decade of experience with the use of monopolar

322 pulsed constant-current stimulation at supramaximal levels that were tested medial and

323 distal to tumor resection. These 2 measurements were used to create an amplitude ratio,

324 which is reported as a percent dropoff. A percent dropoff of $\leq 69\%$ yielded a predictive

325 value of 94% for postoperative HB I-II function. This method was not reliable in

- 326 predicting poor functional outcomes and marginal in predicting moderate function.
- 327

328 Also in 2013, Arnoldner et al¹⁰ reported on the predictive value of using percent

329 maximum values, defined as current level stimulus/maximum muscle response (see Table

330 3 for specifics). A percent maximum of >50 had a 0.9 PPV for HB I-II function. The

responses obtained with 0.3-mA current at the brainstem yielded the best predictive

results for HB I-II function compared to the other studied currents of 0.05, 0.1, and 0.2

333 mA. The authors recommended this monitoring method as complementary when evoked

responses do not conform to more conventional predictors.

335

336 Marin et al's 2011 study¹⁹ described 100% success in determining HB I-II outcome at 1

337 year after surgery when the brainstem stimulation threshold was <0.05 mA. This dropped

to 93% if the threshold was 0.05 mA. Also in 2011, Amano et al⁹ reported a high

339 predictive value by using amplitude ratios gathered from continuous evoked EMG (refer

340 to Table 3 for formula specifics). This method was heralded as a real-time assessment 341 able to facilitate warning criteria that could influence the surgeon to stop tumor 342 dissection. An amplitude preservation ratio of >50% had a 95% probability of 343 maintaining HB I-II at 1 year. The biggest limitation with this method would be 344 identifying the FN root at the start of surgery to place the probe, something that may not 345 be possible with larger tumors. Once in place, there is a need to repeatedly check for 346 probe migration (the authors verified probe position a minimum of every 30 minutes). 347 This type of monitoring requires a demanding continuous assessment and interpretation 348 of the various compound muscle action potentials (CMAPs) by an experienced 349 electrophysiologist. Finally, facial muscle groups were assessed individually as opposed 350 to the more conventional composite assessment of facial function (results reported for 351 558 muscles in 216 patients). The concept of using A-train time, a reflection of 352 neurotonic discharge activity, as a prognosticator of facial function has also been reported. In 2007, Prell et al²² found that an A-train time longer than 10 seconds was 353 354 correlated with long-term deficits in facial function with a specificity of 81%. For the patients with A-train times of <10 seconds and normal preoperative facial function, 81% 355 356 regained normal function at 1 year. Amplitude at the minimum stimulus threshold (MST) was used by Neff et al^{21} as a prognostic indicator of function at 1 year after surgery. 357 358 Applying a logistic regression model, the probability of achieving HB I-II was 98% when 359 MST was ≤ 0.05 mA and response amplitude was $>240 \mu$ V. Independently, the 2 360 parameters were not as sensitive. The authors cautioned that amplitude results were 361 technique-dependent, with responses varying according to the contact established between probe and the FN. In 2002, a study by Nakao et al²⁰ found that ordinary or 362 363 irritable patterns on EMG during the last step of tumor removal predicted 85% and 95% 364 HB I-II function at 1 year, respectively. The last step was in reference to dissection of 365 tumor around the porus of the internal auditory canal, which they leave for last. Silent 366 patterns, on the other hand, were more likely associated with poor long-term outcome 367 (HB III-VI).

368

In 2002, Fenton et al¹² provided a follow-up on a prior report on the utility of using the
 minimum stimulation intensity medial to the tumor after excision (MIMAE) and facial

function at 2-year follow-up. Consistent with their prior report,¹³ MIMAE was again not 371 372 found to be an independent predictor of long-term facial function. Another studied 373 parameter that ultimately lacked electroprognostic value involved amplitudes responses 374 in evoked facial muscle responses when the FN was stimulated at the brainstem. 375 Yokoyama et al²⁹ described how this method was better at predicting time to recovery rather than ultimate functional outcome. Mandpe et al¹⁸ found that by combining 376 377 postresection stimulation thresholds and response amplitudes (these were obtained distal 378 and proximal to tumor resection), the 2 had superior prognostic value than when they 379 were used independently. By using these 2 parameters, they had a 12% false positive rate when predicting good HB function. Magliulo et al¹⁷ compared 3 previously reported 380 381 electroprognostic methods (see Table 3 for specifics) and found the most reliable to be 382 ratios of stimulation intensity over the amplitude evoked responses when compared to 383 amplitudes of train activity or amplitude of evoked response at the brainstem 384 postresection of tumor. The ratios were helpful in predicting HB I-II outcomes at 1 year; 385 however, they were not reliable in predicting poor outcomes. The study was limited by small size and retrospective analysis. A prospective analysis of 109 patients by Zeitouni 386 et al²⁸ in 1997 found good prognostic value in the minimum stimulus thresholds obtained 387 388 at the brainstem post-tumor resection. A stimulus threshold of <0.1 mA predicted good 389 facial function at 1 year in 87% of their cohort. Conversely, higher thresholds were not predictive of poor outcomes. Selesnick et al²⁴ grouped meningioma and VS patients in a 390 391 retrospective study in 1996. A stimulation threshold of ≤ 0.2 mA was predictive of good 392 long-term facial function. Again, poor function was not reliably measured using this 393 parameter. The results do not differentiate between the 2 different pathologies included in 394 the study cohort, VS and meningioma. The authors mention that meningiomas comprised 395 14% of tumors, but no further differentiation in electrical activity was reported between the 2 groups. Taha et al²⁷ used amplitude ratios measured postresection at the brainstem 396 397 and distally at the internal auditory canal, and determined that a ratio of 2:3 was 398 predictive of good long-term function. Statistical analysis, however, was not reported, and the study cohort was small at 20 patients. Silverstein et al²⁶ used the minimum 399 400 current levels needed to elicit a response at the brainstem after tumor resection and found 401 a strong correlation with good facial function when responses were elicited at ≤ 0.1 mA.

- 402 The poor outcomes, however, could not be predicted. Lalwani et al¹⁵ concluded that good
- 403 long-term FN function correlated well with thresholds of 0.2 volts (V) or less at the
- 404 brainstem, posttumor resection.

405 SYNTHESIS OF RESULTS

Level 3 data suggest wide variability in ICNM protocols with multiple different
electroprognostic parameters found to accurately predict good long-term postoperative
FN function following VS surgery. Successful parameters included postresection
stimulation currents or thresholds, response amplitudes, and continuous EMG patterns.
Ratios using a combination of these parameters were also successfully applied. Of the
various stimulating probes used for direct nerve stimulation, monopolar devices were
preferred or reported as most consistent by various groups.^{9,11,12,15,17–19,23,24,26–30}

413 **DISCUSSION**

414 Evidence suggests that various methods can be successfully used to predict good postoperative FN function following VS resection. While several electroprognostic 415 416 parameters were identified as positive predictors of good functional outcome, none of 417 them could consistently predict poor long-term function. The lack of consistency in 418 methods by authors was driven by institutional experience, comfort level of the surgical 419 team, availability of specific equipment, and ultimately, the presence of an independent 420 electrophysiology service. Continuous EMG monitoring, such as when evaluating for 421 tonic or train activity, is laborious and requires a dedicated team member for continuous 422 assessment throughout tumor resection. This is also a task that requires a specific skillset 423 for interpretation. Other methods, such as postresection thresholds at the brainstem, are 424 not as laborious; however, even these measurements are afflicted by confounding factors, 425 such as variability in equipment and their specific electrical settings. The desired benefit 426 of using electroprognostic parameters to predict good functional outcome lies in the ability to counsel patients on the timing of surgical intervention for facial rehabilitation. 427 428 An observation period of 12 to 18 months is typically adhered to in patients with 429 postoperative paresis or paralysis and anatomically intact FN to allow for spontaneous 430 return of function. If a reliable electrical parameter predictive of ultimate good facial 431 outcome is possessed, the clinician can confidently counsel patients to proceed with

- 432 conservative management and postpone early surgical dynamic facial rehabilitation.
- 433 Conversely, none of the parameters proved to successfully predict poor functional
- 434 outcome. This is a reflection on the limitation of electrical currents in distinguishing
- 435 neuropraxia from axonotmesis or neurotmesis at a single time point, following resection.

Question 3

Does an anatomically intact facial nerve with poor electromyogram electrical responses during intraoperative testing reliably predict poor long-term facial nerve function?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery.

Recommendation

Level 3: Poor intraoperative electromyogram electrical response of the facial nerve should not be used as a reliable predictor of poor long-term facial nerve function.

436 STUDY SELECTION AND CHARACTERISTICS

- 437 A total of 2853 candidate studies were screened and assessed for eligibility per the
- 438 previous criterion and 21 studies were included in the final review.^{9–29} The
- 439 electroprognostic value of ICNM in determining poor long-term postoperative FN
- 440 function was the basis of the recommendation in this section. To be included as a part of
- this recommendation, a study had to provide a cohort of patients with assessment of pre-
- 442 and postoperative FN function using an established FN function grading system, such as
- the HB scale or the SB scale. In addition, the method of intraoperative FN monitoring
- had to be described and a minimum of 1 year of follow-up was required for the
- 445 determination of long-term outcomes.

446 **RESULTS OF INDIVIDUAL STUDIES, DISCUSSION OF STUDY**

447 LIMITATIONS, AND RISK OF BIAS

- 448 All studies were thought to represent Level 3 data, primarily due to the lack of blinded
- 449 assessment and the absence of a validation set. The key results of individual studies are
- 450 outlined in evidence Table 3C and are summarized within the guideline
- 451 recommendations. Of the 21 studies used in this analysis, 4 studies discussed
- 452 intraoperative eletrophysiologic parameters with electroprognostic value for "poor"

453postoperative facial function at ≥ 1 year.9,11,20,22Poor function in this analysis was defined454as HB grade IV or greater. There was a heterogeneous methodology used in455electroprognostic parameters; however, all authors provided details on the criteria applied456for their assessment. All the ICNM techniques made use of continuous or electrically457evoked EMG activity. Most studies listed below were described in detail in the earlier458sections of this paper; therefore, only highlights pertaining to the question at hand will be459included in this section.

460

In 2012, Carlson et al¹¹ specifically evaluated long-term facial outcomes in patients with 461 462 poor electrical response after tumor resection in anatomically intact nerves. They could 463 effectively prove that absence of electrical response did not definitively imply poor 464 functional outcome, which was defined as HB IV-VI. Although the study cohort was 465 small, at 11 patients, only 36% of the patients with electrical silence ultimately developed 466 poor function and 18% of patients (n = 2) reached HB II status. These results speak 467 against committing patients to immediate intraoperative FN grafting because of the possibility for spontaneous recovery. In 2011, Amano et al⁹ used a logistic regression 468 469 analysis of amplitude preservation ratios as a risk assessment tool for surgeons. Ratios of 470 <40% carried a higher risk of poor facial function at 1 year and was the authors' own 471 personal indicator to stop tumor resection to reduce the chances of a severe facial palsy. 472 This indicator, however, did not seem to be part of a strict protocol, and further details or 473 statistical analysis were not provided. Duration of A-train activity was a negative predictive factor as discussed by Prell et al^{22} in 2007. A-train time >10 seconds was 474 475 associated with a minimum of a 2-grade drop in HB function in the early and late 476 postoperative period (P < .001 and P < .015, respectively). A sensitivity of 57.1% and specificity of 81% was reported for poor long-term facial outcomes. In 2002, Nakao et 477 al²⁰ found that silent patterns on EMG were predictors of poor facial function; however, 478 479 this was also based on a very small cohort (2/11 patients or 11%). 480

481 The remaining studies were unable to identify reliable independent parameters for poor

482 long-term functional outcomes. This included supramaximal stimulation (SMS) of

483 proximal to distal ratios,²³ the concept of percent of maximum (current

- 484 stimulus/maximum muscle response),¹⁰ maximum stimulus thresholds (MST),^{21,26}
- 485 minimum stimulation thresholds (ST),^{19,24,28} minimum stimulation intensity after tumor
- 486 excision (MIMAE),¹² or voltage of evoked amplitudes^{15,29} and a combination or ratios of
- 487 the response thresholds and amplitudes.^{17,18,27}

488 SYNTHESIS OF RESULTS

Level 3 evidence suggests that A-train duration, amplitude ratios, absent electrical responses, and silent EMG patterns are potential prognosticators for poor facial function outcomes. Although a silent pattern or A-train EMG activity were prognostic indicators for poor function, patients with A-train activity and even absent electrical stimulation after tumor resection were also shown to still have opportunity for spontaneous recovery in the long term. Therefore, the absence of electrical stimulation after tumor resection

does not necessarily commit patients to permanent facial paralysis.

496 **DISCUSSION**

- 497 Level 3 data do not support the use of specific electroprognostic criteria to reliably
- 498 predict poor facial function after VS surgery. Although a handful of parameters were
- 499 presented as potential predictors, none had strong predictive value or were powered to do
- 500 so. The strongest argument against using electrical markers as predictors for poor
- 501 function was based on observation that patients with electrical silence, or absent
- 502 responses at the end of surgery, did not necessarily develop a permanent facial paralysis.
- 503 Whereas several markers can be reliably used to predict good facial function, the ability
- 504 to predict poor function is still limited. Because we cannot reliably predict poor long-term
- 505 FN function with intraoperative electroprognostic testing, early facial reanimation should
- 506 not be employed unless nerve transection is certain.

507 COCHLEAR NERVE MONITORING

Question 4

Should intraoperative eighth cranial nerve monitoring be used during vestibular schwannoma surgery?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery with measureable preoperative hearing levels and tumors <1.5 cm.

Recommendation

Level 3: Intraoperative eighth cranial nerve monitoring should be used during vestibular schwannoma surgery when hearing preservation surgery is attempted.

508 STUDY SELECTION AND CHARACTERISTICS

509 A total of 1849 candidate studies were screened and assessed for eligibility per the

510 previous criterion and 7 studies were included in the final analysis.^{31–37} The value of

- 511 ICNM in hearing preservation was the basis of the recommendation in this section. In
- 512 order to be included as a part of this recommendation, a study had to provide a cohort of
- 513 patients with assessment of pre- and postoperative hearing function using an established
- 514 system, such as the 1995 AAO-HNS or the GR hearing classification system, or
- 515 presented data using WRS and PTA for defining hearing status, or had individual patient
- 516 data presented such that the latter criteria could be applied and analyzed. In addition, the
- 517 method of intraoperative cochlear nerve monitoring had to be described. Data extraction
- 518 included study design, level of evidence, number of patients, tumor characteristics,
- 519 method of ICNM, and the electrical characteristics that correlated with postoperative
- 520 hearing function.

521 RESULTS OF INDIVIDUAL STUDIES, DISCUSSION OF STUDY

522 LIMITATIONS, AND RISK OF BIAS

523 All studies were thought to represent Level 3 data, primarily due to the lack of blinded

524 assessment and the absence of a validation set. The key results of individual studies are

- 525 outlined in evidence Table 4A and are summarized within the guideline
- 526 recommendations. Of the 7 studies noted above, 5 studies provided objective
- 527 comparisons between monitored and unmonitored surgeries^{31,33–36} and are therefore used
- 528 in this recommendation. Hearing preservation in this analysis was defined as any

measurable hearing using the AAO-HNS or GR classification systems. Each of the 5studies will be described briefly.

531

In 2008, Piccirillo et al³¹ retrospectively reviewed hearing outcomes in patients with 532 533 tumors <1.5 cm and normal preoperative hearing. They did not find an advantage in 534 hearing preservation outcomes when comparing monitored versus unmonitored cases. A 535 significant prognostic factor, however, was the presence of cranial nerve action potentials 536 (CNAP) at the end of surgery. Those patients were statistically more likely to have good 537 (AAO-HNS Class A) postoperative hearing (P < .01). The presence of CNAP at the end 538 of surgery, however, did not ensure good hearing outcomes. In their series, more than 539 half of the patients with intact CNAP after tumor removal ultimately had poor hearing 540 outcomes. The technical difficulties of direct eighth nerve monitoring (DENM), such as 541 (1) initial placement of the electrode proximal to the tumor and (2) maintaining that 542 placement throughout surgery, were highlighted and are important considerations for 543 surgeons wishing to undertake this type of monitoring. Finally, a limitation is the lack of 544 long-term data. The authors do not delineate the timeline in which postoperative hearing 545 function was assessed, thereby, limiting assessment of long-term outcomes.

546

In 1994, Nedzelski et al³³ assessed cochlear compound action potentials (CAP) as an 547 548 electroprognostic parameter for hearing preservation. Of the 80 patients included in the 549 cohort, 56 were successfully monitored. This was a retrospective review that compared 550 monitored (n = 56) to unmonitored cases (n = 20). All patients had preoperative 551 serviceable hearing and tumors ≤ 1.5 cm. Long-term hearing assessments were provided 552 at 1 year after treatment and hearing preservation rates were higher in the monitored 553 group (P < .02). Significantly better hearing preservation rates were seen in patients with 554 a measurable intraoperative CAP following tumor resection, although 1 patient with 555 absent CAP had serviceable hearing. CAP click threshold shifts of <20 dB predicted 556 serviceable hearing levels in 71% of patients. Shifts >20 dB, in turn, predicted poor 557 hearing outcomes. Eighteen of the patients with measureable CAP either had absent or 558 nonserviceable hearing, which speaks to the inconsistency of this parameter. This is 559 thought to be secondary to the persistence of cochlear microphonic potentials in the distal 560 cochlear nerve despite anatomic discontinuity or dysfunction in the proximal segment.

561

In 1992, Harper et al³⁵ experienced significant improvement in hearing preservation rates 562 563 in monitored cases using ABR. The difference was statistically significant only for small 564 tumors (≤ 1.1 cm). In their experience, preservation of Wave I and V were positive prognostic factors, with a 67% likelihood of useful hearing preservation. Postoperative 565 566 hearing was measured at 3 months, limiting long-term assessment. Similar findings were reported by Slavit et al in 1991.³⁶ In this study, a comparison was made between ABR-567 568 monitored cases versus no monitoring. Although there was no statistical advantage in the 569 ABR group, there was a trend in that direction. The most pronounced effect was seen in 570 tumors that were <1 cm, and none of the patients with tumors >3 cm had preservation of hearing. Kemink et al³⁷ found that complete loss of ABR Wave V was predictive of 571 572 profound hearing loss. However, not all patients with complete hearing loss had an 573 absence of Wave 5. In this cohort, hearing preservation was not achieved in patients with tumors >1.5 cm. In a smaller cohort, Kveton et al³⁴ did not find a significant difference 574 575 between monitored and unmonitored cases. On the contrary, the study showed improved 576 serviceable hearing preservation (AAO-HNS Class C or better) in the unmonitored group 577 compared to those monitored with ABR. However, this was not a significant difference. 578 In addition, correlation with tumor size or preoperative hearing levels was not provided.

579 SYNTHESIS OF RESULTS

580 Level 3 evidence supports the use of intraoperative cochlear nerve monitoring in hearing 581 preservation VS surgery. The most common method employed was ABR. The presence 582 or characteristics of Wave I and V, as well as the CAP, were the most useful parameters 583 discussed. The benefit of monitoring was most pronounced in tumors <1.5 cm. Hearing 584 preservation in tumors >3 cm was not observed. Long-term assessments were not 585 uniform, with several groups reporting hearing levels measured only 3 months after 586 treatments or not reporting timing at all. Such short-term assessments limit the ability to 587 assess permanent function.

Question 5

Is direct eighth cranial nerve monitoring superior to the use of far-field auditory brain stem responses?

Target population

This recommendation applies to adult patients undergoing vestibular schwannoma surgery with measurable preoperative hearing levels and tumors <1.5 cm.

Recommendation

Level 3: There is insufficient evidence to make a definitive recommendation.

588 STUDY SELECTION AND CHARACTERISTICS

589 A total of 1849 candidate studies were screened and assessed for eligibility per the previous criteria, and 7 studies were included in the final analysis.^{31–37} The utility of 590 591 ICNM in hearing preservation was the basis of the recommendation in this section. A 592 focus on 2 specific modalities, DENM and far-field ABR, was addressed. To be included 593 in this recommendation, a study had to provide a cohort of patients with assessment of 594 pre- and postoperative hearing function using an established system, such as the AAO-595 HNS or GR hearing classification system, or presented data using WRS and PTA for 596 defining hearing status, or had individual patient data presented such that the latter 597 criteria could be applied and analyzed. In addition, the method of intraoperative cochlear 598 nerve monitoring had to be described and direct comparison between DENM and ABR 599 provided. Of the seven studies noted above, one study met the inclusion criteria for this recommendation.³² Data extraction included study design, level of evidence, number of 600 601 patients, tumor characteristics, method of ICNM, electrical characteristics evaluated, and 602 pre- and postoperative hearing levels.

603

604 **RESULTS OF INDIVIDUAL STUDIES, DISCUSSION OF STUDY**

605 LIMITATIONS, AND RISK OF BIAS

606 The study used for this recommendation was thought to represent Level 3 data, primarily

607 due to the lack of blinded assessment and the absence of a validation set. The key results

608 of the study are outlined in Table 4B and summarized within the guideline

- 609 recommendations. This was the only study that provided a direct comparison between the
- 610 2 modalities of cochlear nerve monitoring.

611

In 2004, Danner et al³² retrospectively compared hearing preservation outcomes between 612 613 the use of DENM and ABR. In their series, DENM offered improved hearing 614 preservation outcomes when compared to ABR. The authors attributed superiority to the 615 larger amplitudes obtained with DENM, which in turn required less data averaging and 616 translated into faster, "real-time" assessment of nerve integrity. There was a bias in 617 choice of monitoring modality in that DENM became the preferred monitoring modality 618 after 1995 (study range 1992–2002). They felt experience bias did not affect outcomes in 619 this comparison because of the senior surgeon's established expertise at the onset of the 620 study. It was the senior surgeon's opinion that his learning curve had plateaued at the 621 onset of the study, which is a subjective assessment with risk for recall bias. Consistency 622 in their surgical technique between ABR and DENM was emphasized and discussed to 623 mitigate suspected experience bias. The numbers, however, are skewed toward the 624 DENM group, which was double the size of ABR group, 44 versus 22 patients, 625 respectively. Hearing preservation rates were overall highest amongst patients with 626 tumors ≤ 1.5 cm, regardless of monitoring modality. Again, long-term hearing outcomes 627 were not assessed, and the timing of postoperative hearing assessments was not specified.

628 DISCUSSION FOR COCHLEAR NERVE SECTION

629 The challenges of defining "hearing preservation" continue to plague the literature. 630 Hearing preservation rates vary with respect to the criteria used to report them. The 631 variability has been addressed by endorsing standardized hearing classification systems, 632 such as the AAO-HNS or GR scales. Despite these efforts, consensus lacks on what 633 characterizes useful or serviceable hearing. In the AAO-HNS system, Class A and Class 634 B represent "useful" or "serviceable" hearing and constitute successful hearing 635 preservation surgery. The equivalent in the GR scale is represented by grades I and II. 636 Due to the variability in the reports surrounding what can be classified as successful 637 hearing preservation, the authors opted to be inclusive of hearing levels, not just 638 serviceable hearing, as long they were reported using a standardized system or provided 639 PTA or WRS levels. Applying the aforementioned serviceable hearing criteria to the 640 entire analysis would have been too restrictive given the limited number of studies 641 available for this article. The questions posed in the cochlear nerve section will be

discussed in tandem as they include only 7 studies, compared to the larger amount ofliterature available for the FN section.

644

645 The data extracted from modern day reports supported using ICNM for hearing 646 preservation in patients with preoperative hearing and small tumors. The benefit was seen 647 with ABR or DENM. A tumor size cutoff of ≤ 1.5 cm was identified as being more likely 648 to provide hearing preservation than larger tumors. Hearing outcomes in larger tumors 649 were poor regardless of preoperative hearing status or monitoring modality. The biggest 650 challenge with neuromonitoring of the cochlear nerve involves the technical aspects and 651 delayed feedback. ABR is plagued by delay issues due to the data averaging that is 652 required to assess changes in function. To circumvent this, direct cochlear nerve 653 monitoring has been used instead. The technical requirements and challenges of 654 performing direct cochlear nerve monitoring, however, were made apparent in various 655 reports. They range from the inability to place electrodes at the nerve root exit zone prior 656 to tumor resection to the difficulty in keeping the probes in place throughout the duration 657 of surgery or securing the probe without causing iatrogenic damage to the nerve. Finally, 658 factors such as the presence of excess cerebrospinal fluid or blood, the stimulation 659 voltage used to elicit responses, or the interference of electrocautery stimuli have all been 660 reported to alter responses and the interpretation of results. Dedicated, well-trained 661 electrophysiologists are important members of a hearing preservation team, and most will 662 argue are a necessity.

663

664 In summary, ICNM monitoring has a role in hearing preservation VS surgery. Although 665 there are limitations, Level 3 evidence supports its use. When available, direct eighth 666 nerve monitoring should be employed as well, or in addition to ABR, because of the 667 more immediate real-time responses that can potentially alert the surgeon to noxious 668 stimuli or manipulations. Not all centers have the capability to perform DENM or the 669 electrophysiologists to properly interpret the information during surgery, which limit its 670 widespread implementation. It should also be highlighted that our assessment of the 671 superiority of DENM in hearing preservation surgery is based on 1 study, and, as such, 672 caution is advised in implementing drastic neuromonitoring changes to an already

successful surgical team. More studies and data are needed to better assess this electricalmodality.

675 CONCLUSION AND KEY ISSUES FOR FUTURE INVESTIGATIONS

676 The goals of VS surgery have shifted over the years. The safety profile of these surgeries 677 has continued to improve, and modern-day mortality is at an all-time low. As a result, a 678 great deal of focus is now placed on minimizing morbidity, including hearing loss and 679 facial paresis. The current expectation is that complete tumor resection is to be 680 undertaken with a serious intent to achieve good postoperative facial function. A similar 681 concept has been adopted in patients with existing preoperative hearing. Although the 682 primary goal of VS surgery is still to achieve safe and complete tumor resection, a shift 683 into subtotal resections with the hope of preserving these 2 functions has become more 684 widely accepted. The benefits of using ICNM has been accepted and is supported in this 685 analysis. Despite the best of surgical techniques and electrophysiology equipment, 686 surgical outcomes are still bound by tumor characteristics, such as size. Large tumors are 687 more likely to result in facial paralysis and hearing loss when compared to small tumors. 688 689 As technology continues to evolve and the comfort level of surgical teams continues to

690 improve, clinicians will hopefully learn more about specific parameters that will help as 691 reliable prognosticators of functions. Although several factors were discussed in this 692 review, the sensitivity and specificity profile of each will need to be validated and 693 reproduced in future studies. More prospective analyses will be needed to help with this 694 endeavor.

695 Conflict of Interest (COI)

The Vestibular Schwannoma Guidelines Task Force members were required to report all possible COIs prior to beginning work on the guideline, using the COI disclosure form of the AANS/CNS Joint Guidelines Committee, including potential COIs that are unrelated to the topic of the guideline. The CNS Guidelines Committee and Guideline Task Force Chair reviewed the disclosures and either approved or disapproved the nomination. The CNS Guidelines Committee and Guideline Task Force Chair are given latitude to approve nominations of Task Force members with possible conflicts and address this by

- restricting the writing and reviewing privileges of that person to topics unrelated to the
- possible COIs. The conflict of interest findings are provided in detail in the companion
- 705 introduction and methods manuscript (https://www.cns.org/guidelines/guidelines-
- 706 management-patients-vestibular-schwannoma/chapter_1).

707 Disclaimer of Liability

708 This clinical systematic review and evidence-based guideline was developed by a 709 multidisciplinary physician volunteer task force and serves as an educational tool designed to provide an accurate review of the subject matter covered. These guidelines 710 711 are disseminated with the understanding that the recommendations by the authors and 712 consultants who have collaborated in their development are not meant to replace the 713 individualized care and treatment advice from a patient's physician(s). If medical advice 714 or assistance is required, the services of a competent physician should be sought. The 715 proposals contained in these guidelines may not be suitable for use in all circumstances. 716 The choice to implement any particular recommendation contained in these guidelines 717 must be made by a managing physician in light of the situation in each particular patient

and on the basis of existing resources.

719 Disclosures

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736 FIGURES



737

738 **Figure 1.** Facial nerve monitoring article flow chart.



Figure 2. Cochlear nerve monitoring article flow chart.

743 **Table 1.** Facial nerve monitoring primary search strategy, results, and initial pruning

ENDNOTE PUBMED (NLM), searched on May 10, 2015:

Search 1: All Fields, Contains "Acoustic neuroma" AND, all fields, contains, "Facial nerve" AND, all fields, contains "Surgery"

Total 1392

Search 2: All Fields, Contains "Vestibular schwannoma" AND, all fields, contains "Facial nerve" AND, all fields, contains "Surgery"

Total 676

Search 3: All Fields, Contains "Acoustic neuroma" AND, all fields, contains, "Facial nerve" AND, all fields, contains "Prognostic"

Total 58

Search 4: All Fields, Contains "Vestibular schwannoma" AND, all fields, contains "Facial nerve" AND, all fields, contains "Prognostic"

Total 35

TOTAL: 2161

TOTAL with duplicates excluded: 1519

ENDNOTE EMBASE, searched on May 10, 2015:

Search 1: Abstract, Contains "Acoustic neuroma" AND, abstract, contains, "Facial nerve" AND, abstract, contains "Surgery

Total 207

Search 2: Abstract, Contains "Vestibular schwannoma" AND, abstract, contains "Facial nerve" AND, abstract, contains "Surgery"

Total 233

Search 3: Abstract, Contains "Acoustic neuroma" AND, abstract, contains, "Facial nerve" AND, abstract, contains "Prognostic"

Total 12

Search 4: Abstract, Contains "Vestibular schwannoma" AND, abstract, contains "Facial nerve" AND, abstract, contains "Prognostic"

Total 24

TOTAL 476

TOTAL with duplicates excluded: 432

ENDNOTE Web of Science, searched on May 10, 2015:

Search 1: Title/Keywords/Abstract, contains "Acoustic neuroma" AND, Title/Keywords/Abstract, contains, "Facial nerve" AND, Title/Keywords/Abstract, contains "Surgery"

Total 732

Search 2: Title/Keywords/Abstract, contains "Vestibular schwannoma" AND, Title/Keywords/Abstract, contains "Facial nerve" AND, Title/Keywords/Abstract, contains "Surgery"

Total 599

Search 3: Title/Keywords/Abstract, contains "Acoustic neuroma" AND, Title/Keywords/Abstract, contains, "Facial nerve" AND, Title/Keywords/Abstract, contains "Prognostic"

Total 78

Search 4: Title/Keywords/Abstract, contains "Vestibular schwannoma" AND, Title/Keywords/Abstract, contains "Facial nerve" AND, Title/Keywords/Abstract, contains "Prognostic"

Total 65

TOTAL 1474

TOTAL with duplicates excluded: 902

Summary of primary search: facial nerve monitoring

Combined from 3 database searches, total of 2853 candidate articles Deleted articles published before 1/1/1990 and after 12/31/2014. Deleted all duplicate articles Total number of candidate articles after primary search = 1984

744

746 **Table 2.** Cochlear nerve monitoring primary search strategy, results and initial pruning

ENDNOTE PUBMED (NLM), searched on May 10, 2015:

Search 1: All Fields, Contains "acoustic neuroma" OR All fields, Contains "vestibular schwannoma" AND All Fields, Contains "audiometric"

Total: 176

Search 2: All Fields, Contains "acoustic neuroma" OR All fields, Contains "vestibular schwannoma" AND All Fields, Contains "tinnitus"

Total: 456

Search 3: All Fields, Contains "acoustic neuroma" OR All fields, Contains "vestibular schwannoma" AND All Fields, Contains "sudden hearing loss"

Total: 183

Search 4: All Fields, Contains "acoustic neuroma" OR All fields, Contains "vestibular schwannoma" AND All Fields, Contains "asymmetry"

Total: 68

TOTAL: 883

ENDNOTE EMBASE, searched on May 10, 2015:

Search 1: Abstract, Contains "acoustic neuroma" OR Abstract, Contains "vestibular schwannoma" AND Abstract, Contains "audiometric"

Total: 108

Search 2: Abstract, Contains "acoustic neuroma" OR Abstract, Contains "vestibular schwannoma" AND Abstract, Contains "tinnitus"

Total: 253

Search 3: Abstract, Contains "acoustic neuroma" OR Abstract, Contains "vestibular schwannoma" AND Abstract, Contains "sudden hearing loss"

Total: 37

Search 4: Abstract, Contains "acoustic neuroma" OR Abstract, Contains "vestibular schwannoma" AND Abstract, Contains "asymmetry"

Total: 40

TOTAL: 438

ENDNOTE Web of Science, searched on May 10, 2015:

Search 1: Title/Keywords/Abstract, Contains "acoustic neuroma" OR Title/Keywords/Abstract, Contains "vestibular schwannoma" AND Title/Keywords/Abstract, Contains "audiometric"

Results: 112

Search 2: Title/Keywords/Abstract, Contains "acoustic neuroma" OR Title/Keywords/Abstract, Contains "vestibular schwannoma" AND Title/Keywords/Abstract, Contains "tinnitus"

Results: 243

Search 3: Title/Keywords/Abstract, Contains "acoustic neuroma" OR Title/Keywords/Abstract, Contains "vestibular schwannoma" AND Title/Keywords/Abstract, Contains "sudden hearing loss"

Results: 124

Search 4: Title/Keywords/Abstract, Contains "acoustic neuroma" OR Title/Keywords/Abstract, Contains "vestibular schwannoma" AND Title/Keywords/Abstract, Contains "asymmetry"

Results: 49

TOTAL: 528

Summary of primary search: cochlear nerve monitoring

Combined from 3 database searches, total of 1849 candidate articles Deleted all duplicate articles Total number of candidate articles after primary search = 803

747

Author/Year	Study Description	Data Class	Conclusion
Lenarz et al, 1994	Retrospectively compared FN preservation rates of monitored (<i>n</i> = 30) vs. unmonitored (<i>n</i> = 34) VS patients. Compared immediate and 1-year facial outcomes (HB) between the 2 groups. Both bipolar and monopolar probes used. Single center, same surgeon experience between 1986 and 1991. NF status not reported. HB grading system used.	III	The use of monitoring improved immediate and long-term facial nerve outcomes ($P < .05$). This was especially true in large tumors >1.5 cm. HB grade I-II at 1 year 87% (monitored) vs. 74% (no monitor). Grade III-VI at 1 year: 13% (monitored) vs. 26% (no monitor). Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients. Experience bias: same surgeon; monitoring cases occurred in later years compared to unmonitored cases.

Table 3A. Evidence table for question 1

Author/Year	Study Description	Data Class	Conclusion
Silverstein et al, 1993	Retrospective analysis of 121 VS patients. Compared FN outcome immediate and at >1 year (modified HB score) in monitored ($n = 65$) vs. unmonitored cases ($n = 56$). Single center, single surgeon experience between 1974– 1991. NF status not reported. HB grading system used.	III	There were a statistically greater number of patients with intraoperative eighth nerve transection in the unmonitored group ($P <$.05). Monitored patients had better overall early and late facial function compared to unmonitored patients ($P < .02$). Classification of evidence on prognosis, class III. Did not blind outcome measure. No validation in an independent "test set" of patients. Experience bias since monitoring became available later in surgeon experience.

Author/Year	Study Description	Data Class	Conclusion
Kwartler et al, 1991	Retrospective analysis of monitored translabyrinthine VS cases ($n = 89$) to an unmonitored translabyrinthine VS group ($n = 155$). Looked at short-term and >1-year facial function outcomes (HB score). Monopolar probe used. Single center experience between 1986–1987. NF status not reported. HB grading system used.	III	Facial nerve outcomes were better at all time points in the monitored group ($P < .05$) (immediate, time of discharge, 1 year). They found it particularly useful in the tumors >2.5 cm. Classification of evidence on prognosis, class III. Did not blind outcome measure. No validation in an independent "test set" of patients. Experience bias because monitoring became available later in the surgeon's experience.

- 750 FN, facial nerve; HB, House–Brackmann; NF, neurofibromatosis; VS, vestibular
- 751 schwannoma.
- 752
- 753

Author/Year	Study Description	Data	Conclusion
		Class	
Schmitt et al, 2013	Retrospective review of facial	III	SMS proximal to distal
	nerve function outcome after		dropoff $\leq 69\%$ at the end
	VS resection using SMS		of surgery has 94%
	proximal to distal dropoff ratio		chance of predicting HB
	to predict facial nerve function		I-II. SMS >69% had a
	at >1 year postoperatively.		56% chance of HB I-II.
	Monopolar Prass probe used.		Half the patients with
			>75% dropoff will still go
	The dropoff ratio was		on to have HB I-III,
	calculated: 1 – {distant response		therefore poor predictor
	$(\mu V)/proximal response(\mu V)\} \times$		of long-term poor
	100%		function.
	172 VS patients analyzed with		Classification of evidence
	SMS data and >1 year follow-		on prognosis class III.
	up. Only patients with		Did not blind outcome
	anatomically intact nerves were		measure. No validation in
	included.		an independent "test set"
	NF2 patients were included.		of patients.
	Single center and single surgeon		
	experience from 1999–2011.		
	HB grading system used.		

Table 3B. Evidence table for question 2

Author/Year	Study Description	Data	Conclusion
		Class	
Arnoldner et al, 2013	Prospective study. Calculated % maximum (level current stimulus/maximum muscle response) stimulation to predict facial nerve function at >1 year. %Max = SL/Mmax After skin closure, the facial nerve was stimulated	III	%Max calculated using a direct stimulus of 0.3 mA at the brainstem yielded the best predictive results of HB I-II. The facial nerve was stimulated at root exit zone with increasing stimulus intensities 0.05, 0.1, 0.2,
	transcutaneously at the stylomastoid foramen. Increasing stimulus intensities were used until the muscle response amplitude reached a plateau; a supramaximal stimulus was then further applied. The resulting muscle response amplitude was considered the MMax. Kartush bipolar stimulator was used.		 and 0.3 mA. %Max >50 had PPV of HB I-II of 0.9. Sensitivity and specificity was 0.61/0.8, respectively. For %Max of >40 PPV 0.87; >30 PPV 0.80; >20 PPV 0.80; >10 0.79. Cannot predict poor outcomes; if you get a "poor" response of %Max of 11%; you still have a
	 78 VS patients with minimum 1 year follow-up and average follow-up of 523 days. Single center experience between 2005–2010. NF2 patients excluded. HB grading system used. 		high chance (79%) of good outcome. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Marin et al, 2011	Retrospective analysis of 206 VS patients. Used stimulation threshold of 0.05 mA to predict long-term (1-year) facial nerve function. Monopolar probe used dose stimulation. Single center and multiple surgeon experience from 1996– 2008. NF2 patients excluded. Patients with abnormal facial function preoperatively were excluded. HB grading system used.	III	The facial nerve was electrically stimulated at the brainstem by a monopolar probe with a 0.5 mm tip after tumor removal. Stimulation threshold of <0.05 mA predicted HB I-II function in 100% of patients (<i>P</i> < .01). A stimulation threshold of 0.05 predicted HB I-II in 93%. A threshold of >0.05 predicted HB I-II in 82%. Cannot reliably predict poor outcomes with this method. A response was defined as >100 μ V. Classification of evidence on prognosis, level III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data	Conclusion
		Class	
Amano et al, 2011	Retrospective review of	III	Concluded that continual
	continual stimulation evoked		stimulation evoked facial
	facial nerve EMG. Calculated		nerve EMG could be used
	an amplitude preservation ratio		to determine if tumor
	(%) during and after tumor		resection should continue.
	resection and evaluated whether		An amplitude
	this could predict long-term		preservation ratio >50%
	facial function at ≥ 1 year.		or last amplitude $(Mmax) > 1000$
	The facial nerve was electrically		μV was predictive of
	stimulated with monopolar		good facial function. A
	current 0.1–3 mA at 1 Hz and		ratio >50% had 95% HB I
	CMAP continuously monitored.		or II. Unclear what >1000
	Free running spontaneous		μV predicted. A
	EMG, as well as evoked facial		discrepancy with μV
	EMG were used. The		criteria is that they
	stimulation was via monopolar		reached this cutoff with
	probe placed at the nerve's root		several tumors with a
	exit zone at the brainstem.		large amount of remnant;
			thus continued to operate.
	The amplitude preservation		
	ratio (%) = {last M-		Classification of evidence
	$max(\mu V)/Control M-max(\mu V)$		on prognosis class III.
	$\times 100$		Did not blind outcome
			measure. No validation in
	Control M-max = maximum		an independent "test set"
	CMAP amplitude at start of		of patients.
	surgery		
	1 otal patient sample of 232		
	consecutive vS patients		
	excluding 16 patients with		
	preoperative factal weakness,		
	prior surgery, or radiation $(n = 216)$		
	210).		
	Single center experience from		
	2005-2008. NF2 patients		
	excluded.		
	HB grading system used.		

Author/Year	Study Description	Data Class	Conclusion
Prell et al, 2007	Retrospective review of whether A-train duration measured from free running EMG could predict facial nerve function at 1 year postoperatively. A-train activity is defined as a close succession of ≥4 geometrically similar, mono- to triphasic discharges from baseline with an amplitude of at least double background noise. The sequence of ≥4 elements is required to build a frequency of 100–200 Hz, which must be steady over the course of any given A train. Train time is reported in seconds. 40 VS patients with a minimum of 1-year follow-up. 26 had normal preoperative facial function and 14 did not. Single center experience from 1994–2003. NF2 status not reported. HB grading system used.	III	For patients with normal preoperative function and A-train time <10 seconds, 81% had normal function at 1 year. For the entire cohort, an A-train time of >10 seconds predicted a HB II–VI (everyone but normal HB I) in 81% at 1 year of follow up. 5 of 40 (13%) with prolonged A- train times still became HB I. Sensitivity was 57.1% and specificity 81% for the 10-second threshold. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Neff et al, 2005	 Prospective evaluation of 74 consecutive VS patients. Used MST and response amplitude (at MST) to predict long-term facial function at >1 year. Results report data on 71 patients with postoperative anatomically intact facial nerves. Measurements were made at the brainstem or medial to tumor resection. Single center experience, date ranges and NF2 status not reported. HB grading system used. 	Π	Using their logistic regression model, an MST ≤ 0.05 mA with a response amplitude >240 μ V carried a 98% probability of HB I–II at 1 year. Patients with MST >0.05 threshold OR <240, or both still obtained HB I–II in 59% (10/17). <i>P</i> =.015. Predicting poor outcome was not as reliable, perhaps because of the small number of patients in this category (HB III– VI).
			Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Nakao et al, 2002	 Prospective analysis of EMG "pattern" (irritable, silent, stray, ordinary) during the critical portion of tumor removal to see if there was a correlation with long-term facial function ≥1 year. All had normal preoperative facial function. The EMG patterns were classified as follows: 1) an irritable pattern with repeated EMG responses elicited very easily and frequently by the slightest contact with the facial nerve, 2) a silent pattern with little or no EMG responses, 3) a stray pattern with persistent train responses up to 20 minutes despite temporary discontinuance of surgical manipulations, and 4) an ordinary pattern with EMG responses elicited by mechanical stimulation of the nerve but not very easily or frequently. 49 VS patients with at least 1- year follow-up (mean 18 months). Single center, single surgeon experience from April 1998–October 1999. NF2 status not reported. HB grading system used. 	III	An ordinary or irritable pattern predicted HB I–II in 85% and 95%, respectively. A silent pattern only predicted poor outcome HB III–VI in 73% (8/11). Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Fenton et al, 2002	 Prospective study of 67 VS patients with normal preoperative facial function collected from 2 centers. 35 patients met study criteria. Looked at various other predictive factors, including tumor size and surgical approach. Also evaluated MIMAE relationship with long- term facial function (HB score) at 2-year follow-up. MIMAE was obtained by using a constant current technique and a standard pulse duration of 100 seconds required to provoke a stimulus threshold event on an intact facial nerve medial to the tumor location (0.05–3.0 mA). Fisch dissector used. Multi-institution, same senior surgeon, in 1994. NF2 patients excluded. HB grading system used. 	III	Immediate facial nerve function was the only independent predictor of facial nerve function at ≥1 year. MIMAE was not found to be predictive in their multivariable logistic regression model. MIMAE was significant in univariate analysis (odds ratio = 0.57). Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Yokoyama et al, 1999	Retrospective analysis of 66 VS patients. Evaluated intraoperative EFMR at the brainstem after tumor removal. Responses were measured as amplitudes (µV). This was correlated with facial nerve function at 18 months postsurgery. Monopolar currents 0.5–0.6 mA with 100-ms pulse duration was used. Results were classified into 4 groups according to response levels. Single center study. NF2 status not reported. HB grading system used.	III	Amplitudes were not a good predictor of ultimate functional outcome (HB grade). They were a better predictor of time to recovery in patients who did recover; >150 μ V: 3 months to recovery; 100– 149 μ V: 6 months; 50–99 μ V: 9 months; <50 μ V: 12 months (Mann– Whitney <i>U</i> test <0.05). 85% of their cohort with response amplitudes of \geq 100 μ V obtained long- term function of HB I. Function was unpredictable with levels <100 μ V. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Author/Year	Study Description	Data Class	Conclusion
Mandpe et al, 1998	Prospective, nonconsecutive, 44 VS patients. Stimulation threshold at brainstem (volts) and amplitude proximal and distal to tumor resection at 0.2 V above threshold. Monopolar probe used. Correlated this with FN outcomes with \geq 1-year follow-up. Single center experience between 1994–1996. NF2 patients included ($n = 3$). HB grading system used.		Combined threshold $(\leq 0.1 \text{ V})$ and amplitude $(\geq 200 \mu \text{V})$ was superior to threshold alone at predicting early and late postoperative HB I–II FN function. False positive rate was 12% (false positive = poor HB outcome despite favorable ICNM parameters). Cannot reliably predict poor outcomes. 94% of patients with stimulation thresholds $\leq 0.1 \text{ V}$ had HB I–II at 1 year. 89% of those with amplitude >200 μV had HB I–II function. Using combined parameters, 88% had HB I–II function. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

Magliulo et al, 1998Retrospective analysis of 34 VS patients. Compared 3 methods of intraoperative testing: 1) amplitude of train activity lasting 30 seconds (less or greater than 500μ V), 2) amplitude of response at brainstem at 0.05 mA (less than or greater than 500μ V), 3) ratio or greater than 500μ V), 3) ratio methods were analyzed in relation to 10-day and 1-year HB scores. Two center experience between 1990–1994. NF status not reported.Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.Zeitouni et al, 1997Prospective study of 109 VS patients evaluating correlation between the minimum intraoperative stimulus threshold and 1-year facial function outcomes (HB score). Threshold and 1-year facial function outcomes (HB score). Threshold and 1-year facial function outcomes (HB score). Threshold and 1-year facial fush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.IIISingle center experience. NF2 status not reported.Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"	Author/Year	Study Description	Data	Conclusion
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function outcomes (HB score). Thresholds were measured using a constant current technique with a monopolar flush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.Stimulus threshold of 0.05 mA or 0.1 mA predicted HB I–II in 87%. Could not predict long- term poor function.Single center experience. NF2 status not reported.Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"		threshold and 1-year facial		= .048, respectively).
Thresholds were measured using a constant current technique with a monopolar flush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.0.05 mA or 0.1 mA predicted HB I–II in 87%. Could not predict long- term poor function.Classification of evidence on prognosis class III.Classification of evidence on prognosis class III.Single center experience. NF2 status not reported.Did not blind outcome measure. No validation in an independent "test set"		function outcomes (HB score).		Stimulus threshold of
using a constant current technique with a monopolar flush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.predicted HB I–II in 87%. Could not predict long- term poor function.Single center experience. NF2 status not reported.Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"		Thresholds were measured		0.05 mA or 0.1 mA
technique with a monopolar flush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.Could not predict long- term poor function.Single center experience. NF2 status not reported.Classification of evidence on prognosis class III.UDIIDid not blind outcome measure. No validation in an independent "test set"		using a constant current		predicted HB I–II in 87%.
flush tip Prass probe at the brainstem. Minimum intensity used 0.05 mA.term poor function.Single center experience. NF2 status not reported.Classification of evidence on prognosis class III.UDIIDid not blind outcome measure. No validation in an independent "test set"		technique with a monopolar		Could not predict long-
brainstem. Minimum intensity used 0.05 mA. Single center experience. NF2 status not reported. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"		flush tip Prass probe at the		term poor function.
used 0.05 mA.Classification of evidence on prognosis class III.Single center experience. NF2 status not reported.Did not blind outcome measure. No validation in an independent "test set"		brainstem. Minimum intensity		
Single center experience. NF2 status not reported.on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"		used 0.05 mA.		Classification of evidence
Single center experience. NF2 status not reported.Did not blind outcome measure. No validation in an independent "test set"				on prognosis class III.
status not reported. measure. No validation in an independent "test set"		Single center experience. NF2		Did not blind outcome
an independent "test set"		status not reported.		measure. No validation in
HK grading system used		HB grading system used		of patients

Author/Year	Study Description	Data	Conclusion
		Class	
Selesnick et al, 1996	Retrospective analysis of 49 VSs or CPA meningiomas patients. 14% of cohort had CPA meningioma. Stimulation of FN at root entry zone after tumor resection. Constant current method starting at 0.1 mA with 50 µsec pulse duration. Measured stimulation threshold at 0.1-mA increments. Monopolar stimulators were used (Kartush and Prass probe). Compared this to early and 1- year facial function (HB score. Two center, single surgeon experience between 1991–1995. NF2 status not reported.	III	A stimulation of $\leq 0.2 \text{ mA}$ predicted good facial nerve function (HB I or II) at 1 year ($P < .01$). Could not predict poor function. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.
	HB grading system used.		

Author/Year	Study Description	Data Class	Conclusion
Taha et al, 1995	Retrospective analysis of 20 VS patients. Evaluated postresection proximal to distal ratios of amplitude of muscle action potential. They used the lowest intensity required to elicit a response (not a supramaximal stimulation). Compared this with short- and long-term (1 year) HB scores. Mean follow-up was 18 months. The nerve was stimulated at the brainstem and at the internal auditory meatus after tumor resection. Starting at 0.05 mA to a max of 1 mA. Monopolar stimulator probe was used with constant current at 4 pulses per second for 100 msec. Single center experience between 1992–1993. NF status not reported.	III	A proximal to distal ratio of 2:3 was predictive of good long-term function (HB I). Although a ratio of 1:3 predicted poor function (HB IV or more), there were too few patients for adequate statistical analysis ($n = 5$), which was not performed. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.
Silverstein et al, 1994	Retrospective analysis of 44 VS patients. Minimum brainstem threshold response level (constant current, square wave pulse stimulation) in mA used to predict long-term facial function (HB score) at 1 year. Silverstein probe used (monopolar). Single center experience between 1984–1991. NF status not reported.	III	Minimum threshold response of ≤0.1 mA predicted good long-term facial function (HB I) in 95% of patients at least 1 year after surgery. Could not predict poor outcomes. Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set"

755 CMAP, compound muscle action potential; CPA, cerebellopontine angle; EFMR, evoked

756 facial muscle response; EMG, electromyogram; FN, facial nerve; HB, House-

757 Brackmann; MMax, maximum muscle response; MIMAE, medial to the tumor after

- excision; MST, minimum stimulus threshold; NF/NF2, neurofibromatosis; PPV, positive predictive value; SL, current stimulus; SMS, supramaximal stimulation.

Author/Year	Study Description	Data Class	Conclusion
Carlson et al,	Retrospective review of 11 VS	III	"No electrical response"
2012	patients with no measured		was unable to predict
	electrical response at the end of		poor (HB IV–VI) long-
	VS surgery and anatomically		term facial function at >1
	intact nerves. These patients were		year. Only 36% (4/11)
	followed for >1 year to see if "no		had a poor outcome. 18%
	response" parameter could		(2/11) had a HB II
	predict poor facial nerve function.		recovery which would be
	Monopolar Prass probe was used.		superior to current facial
			reinnervation procedure
	Single center, experience from		results.
	2000–2010, mean follow-up of		
	81.8 months. NF2 patients		Classification of evidence
	excluded.		on prognosis class III.
			Did not blind outcome
	HB grading system used.		measure. No validation in
			an independent "test set"
			of patients.

Table 3C. Evidence table for question 3

Author/Year	Study Description	Data Class	Conclusion
Author/Year Amano et al, 2011	Study DescriptionRetrospective review of continualstimulation evoked facial nerveEMG. Calculated an amplitudepreservation ratio (%) during andafter tumor resection andevaluated whether this couldpredict long-term facial functionat ≥ 1 year. The facial nerve waselectrically stimulated withmonopolar current $0.1-3$ mA at 1Hz and CMAP continuouslymonitored. Free runningspontaneous EMG, as well asevoked facial EMG were used.The stimulation was viamonopolar probe placed at thenerve's root exit zone at thebrainstem.The amplitude preservation ratio(%) = {last M-max(μ V)/ControlM-max (μ V)} × 100Control M-max = maximumCMAP amplitude at start ofsurgeryTotal patient sample of 232consecutive VS patientsexcluding 16 patients withpreoperative facial weakness,prior surgery, or radiation ($n = 216$).	Data Class III	ConclusionConcluded that continual stimulation evoked facial nerve EMG could be used to determine if tumor resection should continue.An amplitude preservation ratio >50% or last amplitude
	 216). Single center experience from 2005–2008. NF2 patients excluded. HB grading system used. 		

Author/Year	Study Description	Data Class	Conclusion
Prell et al, 2007	Retrospective review of whether	III	For the patients with
	A-train duration measured from		normal preoperative
	free running EMG could predict		function and A-train time
	facial nerve function at 1 year		<10 seconds, 81% had
	postoperatively. A-train activity		normal function at 1 year.
	is defined as a close succession of		For the entire cohort, an
	at least 4 geometrically		A-train time of >10
	similar, mono- to triphasic		seconds predicted a HB
	discharges from baseline		II–VI (everyone but
	with an amplitude of at least		normal HB I) in 81% at 1
	double background noise. The		year of follow-up. 5 of 40
	sequence of at least four elements		(13%) with prolonged A-
	is required to build a frequency of		Train times still became
	100–200 Hz, which must be		HB I. Sensitivity
	steady over the		was 57.1% and
	course of any given A train. Train		specificity 81% for the
	time is reported in seconds.		10-second threshold.
	40 VS patients with a minimum		Classification of evidence
	of 1-year follow-up. 26 had		on prognosis class III.
	normal preoperative facial		Did not blind outcome
	function and 14 did not.		measure. No validation in
			an independent "test set"
	Single center experience from		of patients.
	1994–2003. NF2 status not		-
	reported.		
	HB grading system used.		

Author/Year	Study Description	Data Class	Conclusion
Nakao et al, 2002	 Prospective analysis of EMG "pattern" (irritable, silent, stray, or ordinary) during the critical portion of tumor removal to see if there was a correlation with long- term facial function ≥1 year. All had normal preoperative facial function. The EMG patterns were classified as follows: 1) an irritable pattern with repeated EMG responses elicited very easily and frequently by the slightest contact with the facial nerve, 2) a silent pattern with little or no EMG responses, 3) a stray pattern with persistent train responses up to 20 minutes despite temporary discontinuance of surgical manipulations, and 4) an ordinary pattern with EMG responses elicited by mechanical stimulation of the nerve but not very easily or frequently. 49 VS patients with at least 1- year follow-up (mean 18 months). Single center, single surgeon experience from April 1998– October 1999. NF2 status not reported. HB grading system used. 	III	An ordinary or irritable pattern predicted HB I–II in 85% and 95%, respectively. A silent pattern only predicted poor outcome HB III–VI in 73% (8/11). Classification of evidence on prognosis class III. Did not blind outcome measure. No validation in an independent "test set" of patients.

- 762 CMAP, compound muscle action potential; EMG, electromyogram; FN, facial nerve;
- 763 HB, House–Brackmann; MMax, maximum muscle response; NF2, neurofibromatosis.

Author/Year	Study Description	Data Class	Conclusion
Piccirillo et	Retrospective analysis of	III	Across all surgeries, those
al, 2008	hearing preservation (modified		with ICNM (ABR,
	Sanna class A–B) in patients		DENM, or both) did not
	undergoing surgical resection of		have a statistically greater
	VSs with or without use of		chance of hearing
	ICNM at a single center from		preservation, though it
	1998–2005 by a single surgeon.		trended in this direction
			(26.7% vs 20.8%, <i>P</i> =
	99 cases of tumor <1.5 cm, NF2		.79).
	excluded, and with Sanna class		
	A–B (AAO-HNS class A)		Surgical approach, either
	preoperative hearing.		MCF or RS/RL, did not
	Fast-ABR (5 sec) and direct		have a significant effect.
	cochlear nerve action potentials		
	were obtained. These were		The only statistically
	measured by placing an		significant parameter was
	electrode directly on the		prognostic value to the
	cochlear nerve.		presence of continued,
			appropriate stimulation
	There were 2 groups. Group I		with ICNM. Meaning, if
	consisted of patients with either		there was a positive
	DENM or auditory brain stem		response at the end of the
	response. Group 2 included		case, you were
	monitoring		statistically likely to have
	monitoring.		preserved hearing ($P < 0.1$ Eicher exact test)
			.01, Fisher exact test).
			Author conclusions.
			ICNM does not help
			preserve hearing, but it
			may have prognostic
			value.
			Classification of evidence
			on prognosis class III. Did
			not blind outcome
			measure. No validation in
			an independent "test set"
			of patients.

Table 4A. Evidence table for question 4

Nedzelski et	Retrospective analysis of	III	Significantly better rates
al 1994	hearing preservation after VS		of hearing preservation
ui, 1991	resection via suboccipital		were seen with patients in
	approach at a single center		whom CAP was measured
	between 1975_1993 Number		intraoperatively (38% vs
	of surgeons involved not		15% P = 02 Results
	specified		1570, T = .02). Results
	specified.		tumor size
	80 cases were evaluated in		tumor size.
	which hearing preservation was		In cases where CAP was
	attempted The		present and unchanged
	magurament/prognostic value		52% of patients had
	of cochloar CAPs vs po		sorviceable bearing. One
	monitoring was evoluated 56		serviceable hearing. One
	notion to had CAP		of CAR had some patho
	manufacture parts during the case		basering
	and the remainder did not (the		nearing.
	remainder also including 4		CAD threshold shifts of
	remainder also including 4		CAP threshold shifts of
	CAD magazine anta from the		≥20 dB predicted
	CAP measurements from the		successful serviceable
	start). CAP was obtained by		nearing preservation
	placing a silver ball electrode		postoperatively in $/1\%$ of
	directly on the promontory via		cases ($P = .001$). >20 dB
	myringotomy. Threshold shifts		shift predicted loss of
	were calculated as the		serviceable hearing ($P < 0.02$)
	difference between threshold		.003).
	measurements at the beginning		
	and end of surgery.		Author conclusions:
			Intraoperative CAP
	All patients had tumors ≤ 1.5 cm		monitoring can be useful
	and SRT <50 and WRS $>60\%$.		for hearing preservation
			attempts.
	Postoperative hearing was		
	followed for I year after		Classification of evidence
	surgery.		on prognosis class III. Did
			measure. No validation in
			an independent "test set"
			of patients.

Harper et al,	Retrospective comparison of	III	When comparing the
1992	hearing preservation rates using		groups across all tumor
	ABR vs no monitoring during		sizes, the ABR group
	VS resection via suboccipital		trended towards better
	approach at a single center		hearing preservation and
	between 1986–1991 by multiple		better useful hearing
	surgeons.		preservation, but the
	501800100		differences were not
	There were 90 consecutive		statistically significant.
	patients who underwent a		5
	hearing preservation attempt		When comparing the
	with use of ABR A control		$\frac{1}{3}$ groups for tumors <1 1
	group of 90 patients who were		cm the ABR groun had
	matched for age tumor size		better hearing
	and preoperative hearing were		preservation (79% vs
	included		42%) and better rates of
	included.		useful preservation (47%
	All patients had preoperative		$v_{s} 21\%$) This was the
	PTA < 65 dB and WRS of		only size category in
	$\sim 10\%$ Erequencies used to		which there was a
	24070. Frequencies used to		statistically significant
	calculate FTA not specified.		difference $(B < 05)$ If
	Postoporative bearing assessed		APP way as L and V
	Postoperative hearing assessed		ABR waves I allu v
	at 5 months. Preserved		preserved, then 67%
	nearing = PIA < 65 dB;		chance of useful hearing.
	"useful" preserved hearing =		A .1 . 1 ·
	WRS >40%.		Author conclusions:
			ABR 1s better than no
			monitoring, particularly
			when tumors are <1.1 cm.
			Classification of evidence
			on prognosis class III. Did
			not blind outcome
			measure. No validation in
			an independent "test set"
			of patients.

Slavit et al,	Retrospective analysis of	III	No tumor >3 cm had
1991	hearing preservation after VS		hearing preserved.
	resection via posterior fossa		
	approach at a single center		Rates of preservation
	between 1986–1989.		trended towards higher
	Comparison included use of		with ABR, but this was
	cochlear nerve monitoring by		not statistically significant
	ABR vs no monitoring. All		(30% with ABR, 20%
	procedures performed by same		without ABR). <i>P</i> values
	surgical team.		not provided.
	60 patients with some		The difference became
	preoperative hearing and use of		more pronounced when
	intraoperative ABR were		focusing on tumors <1 cm
	matched with 60 patients with		(82% preservation with
	no ABR on the basis of tumor		ABR, 36% without) but
	size (within 2 mm),		still not statistically
	preoperative PTA (at 500,		significant. P values not
	1000, and 2000 Hz), word		provided.
	discrimination scores, and year		
	of operation		The difference for the
			preservation of useful
	1 umor size classified as		nearing was also not
	sinally when < 2 cm, medium		The level of preoperative
	when $2-4$ cm, and large when >4 cm		hearing did not seem to
	>4 cm.		matter in terms of rates of
	Follow up included 1 week and		preservation in the 2
	3-month postoperative		groups
	audiogram.		Stoups.
	5		Author conclusions:
	Preservation = anything		Definite trend favoring
	measurable. Useful preservation		the monitored group to
	= PTA <60 dB, WRS >40%		improved rates of hearing
			preservation, but nothing
			statistically significant.
			ABR could not always
			reliably tell when the
			cochlear nerve was cut.
			Classification of evidence
			on prognosis class III. Did
			measure No validation in
			measure. No validation in

			an independent "test set"
			of patients.
Kveton, 1990	Retrospective analysis of	III	No significant difference
	hearing preservation after VS		in preoperative tumor size
	resection via suboccipital-		or postoperative hearing
	transmeatal approach by a		outcome between the
	single surgeon at a single center		monitored and
	between 1987–1989. A		unmonitored groups.
	comparison was performed		
	between cases with		Postoperative serviceable
	intraoperative ABR to ones in		hearing preservation was
	which no monitoring was used.		greater in the non-ABR
			than monitored group
	16 cases were evaluated in		(57% vs 44%, not
	which there was a hearing		significant).
	preservation attempt. Nine		
	patients had intraoperative		No formal analysis of age
	ABR, and 7 had no form of		or tumor size or
	monitoring. All tumors were ≤ 2		preoperative hearing
	cm.		status as potential
			confounders.
	The cochlear nerve was		
	anatomically intact in all cases.		Author conclusions:
			ABR not very helpful.
	All patients had preoperative		
	AAO-HNS class B hearing or		Classification of evidence
	better. Pre- and postoperative		on prognosis class III. Did
	SRT/WRS are listed for		not blind outcome
	comparison. 50/50 criterion		measure. No validation in
	used to define "serviceable"		an independent "test set"
	hearing. Postoperative		of patients.
	audiograms measured at		
	variable intervals (from 2		
	months to 1 year		
	postoperatively).		

- AAO-HNS, American Academy of Otolaryngology-Head and Neck Surgery; ABR,
- auditory brainstem response; DENM, direct eighth nerve monitoring; ICNM, intracranial
- 768 cochlear nerve monitoring; MCF, middle cranial fossa; PTA, pure tone average; RS/RL,
- retrosigmoid-retrolabyrinthine; SRT, speech recognition threshold; VS, vestibular
- 770 schwannoma; WRS, word recognition score.
- 771

Author/Year	Study Description	Data	Conclusion
		Class	
Danner et al,	Retrospective comparison of	III	Analysis of all cases, regardless
2004	hearing preservation rates using		of tumor size (0–2.5 cm), found
	either ABR or DENM		the use of DENM had a
	following VS resection via		statistically greater chance of
	retrosigmoid approach by a		hearing preservation than ABR
	single surgeon between 1992–		(P = .03). However, the
	2002.		differences between tumor sizes
			in the 2 groups (ABR and
	66 patients were included in the		DENM) is not well addressed.
	study for comparison of DENM		,
	and ABR. 22 patients were		Hearing preservation
	monitored with ABR and 44		analysis by size subcategories
	with DENM.		(<1 cm, 1–1.5 cm, 1.5–2 cm, and
			2-2.5 cm), found no statistical
	Patients with a tumor >2.5 cm		difference in any group between
	were excluded from the analysis		DENM and ABR.
	as none had hearing		
	preservation achieved.		DENM had improved rates of
	I		hearing preservation that trended
	All patients had preoperative		towards significance with an
	AAO-HNS class B hearing or		exception of the $2-2.5$ cm group
	better (SRT <50 dB, WRS		where rates were equal.
	>50%).		1
			The type of eighth nerve
	Unclear when postoperative		monitoring did not affect
	audiogram was performed.		postoperative facial nerve
			preservation or CSF leak rates.
			1
			Author conclusions:
			DENM gives better rates of
			hearing preservation in tumors
			<2 cm. DENM is shown to have
			a statistically significant
			advantage when comparing all
			study patients with tumors <2.5
			cm.
			Classification of evidence on
			prognosis class III. Did not blind
			outcome measure. No validation
			in an independent "test set" of
			patients.

Table 4B. Evidence table for question 5

- AAO-HNS, American Academy of Otolaryngology-Head and Neck Surgery; ABR,
- auditory brainstem response; CSF, cerebrospinal fluid; DENM, direct eighth nerve
- 775 monitoring; SRT, speech recognition threshold; VS, vestibular schwannoma; WRS, word
- recognition score.

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