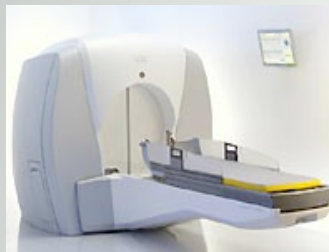




AZIENDA OSPEDALIERA UNIVERSITARIA INTEGRATA VERONA



DAI DI NEUROSCIENZE
ISTITUTO DI NEUROCHIRURGIA
UOC DI NEUROCHIRURGIA B
USO DI RADIOCHIRURGIA E NCH STEREOTASSICA

GRUPPO INTERDISCIPLINARE
NEURO-ONCOLOGICO VERONESE
(GrINOV)

Verona, Sept. 17th, 2015



University Hospital of Verona,
ENT Department
Chief Daniele Marchioni, MD



SKULL BASE ENDOSCOPY

Verona
17 September 2015



GAMMA KNIFE TREATMENT FOR ACOUSTIC NEUROMA. THE VERONA EXPERIENCE

A Nicolato, C Parisi, R Foroni, M Longhi, E Zivelonghi, N Tommasi, L Rosta, GK Ricciardi, PA Polloniato, M Rasi, A D'Ottavio, E Piovan, C Cavedon, GP Pinna, M Meglio

Verona, Sept. 17th, 2015



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VERONA



GAMMA KNIFE IN VSs

GOALS OF RS TREATMENT

**RADICAL TREATMENT OF CRITICAL
LESIONS WITH VERY LOW RISKS OF
PERMANENT COMPLICATIONS AND
HEARING PRESERVATION**

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

ADVANTAGES OF GK RS

- NO INVASIVENESS
- PREVENTION OF INOPERABLE REMNANT PROGRESSION
- PREVENTION OF FURTHER SURGERY
- EXTREMELY HIGH MECHANICAL AND RADIATION FOCUS PRECISION AND COINCIDENCE (± 0.3 mm)
- HIGHLY CONFORMAL DOSE PLANNINGS
- HIGH GRADIENT DOSE
- DIRECT STEREO-MRI ACQUISITION: volumetric and steady-state sequences



GAMMA KNIFE IN IAC-VSs

ADVANTAGES OF GK RS

- NO INVASIVENESS
- PREVENTION OF INOPERABLE REMNANT PROGRESSION
- PREVENTION OF FURTHER SURGERY
- EXTREMELY HIGH MECHANICAL AND RADIATION FOCUS PRECISION AND COINCIDENCE (± 0.3 mm)
- **HIGHLY CONFORMAL DOSE PLANNINGS**
- **HIGH GRADIENT DOSE**
- **DIRECT STEREO-MRI ACQUISITION: volumetric and steady-state sequences**



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VERONA



GAMMA KNIFE IN VSs



STEREOTACTIC RADIOSURGERY

RADIATION-INDUCED SURGICAL EFFECT

**OPTIMIZING TARGET
DOSE-EXPOSURE**

**MINIMIZING SURROUNDING
STRUCTURES EXPOSURE**



STEEP DOSE FALL-OFF

Patient Tools Plan Workspace Help

Acoustic4 y: z: LEKSELL GAMMA KNIFE C

LEKSELL GAMMA KNIFE C

Toolbox

Axial (mri)

Coronal (mri)

Dose Statistics Measurements

Line Leksell (110.5, 106.8, 113.2)-(83.8, 83.8, 113.7)
Plan = first_plan
Distance = 35.2 mm
Max = 40.0 Gy (100 %) at (95.5, 93.9, 113.5)
Mark = 40.0 Gy (100 %) at (95.5, 93.9, 113.5)

SP: 408 z: 113.4

OpenBook (mri)

Point Line Volume Histogram

Exit Clear Print... Help

Dose: Select command and mark line or point in image

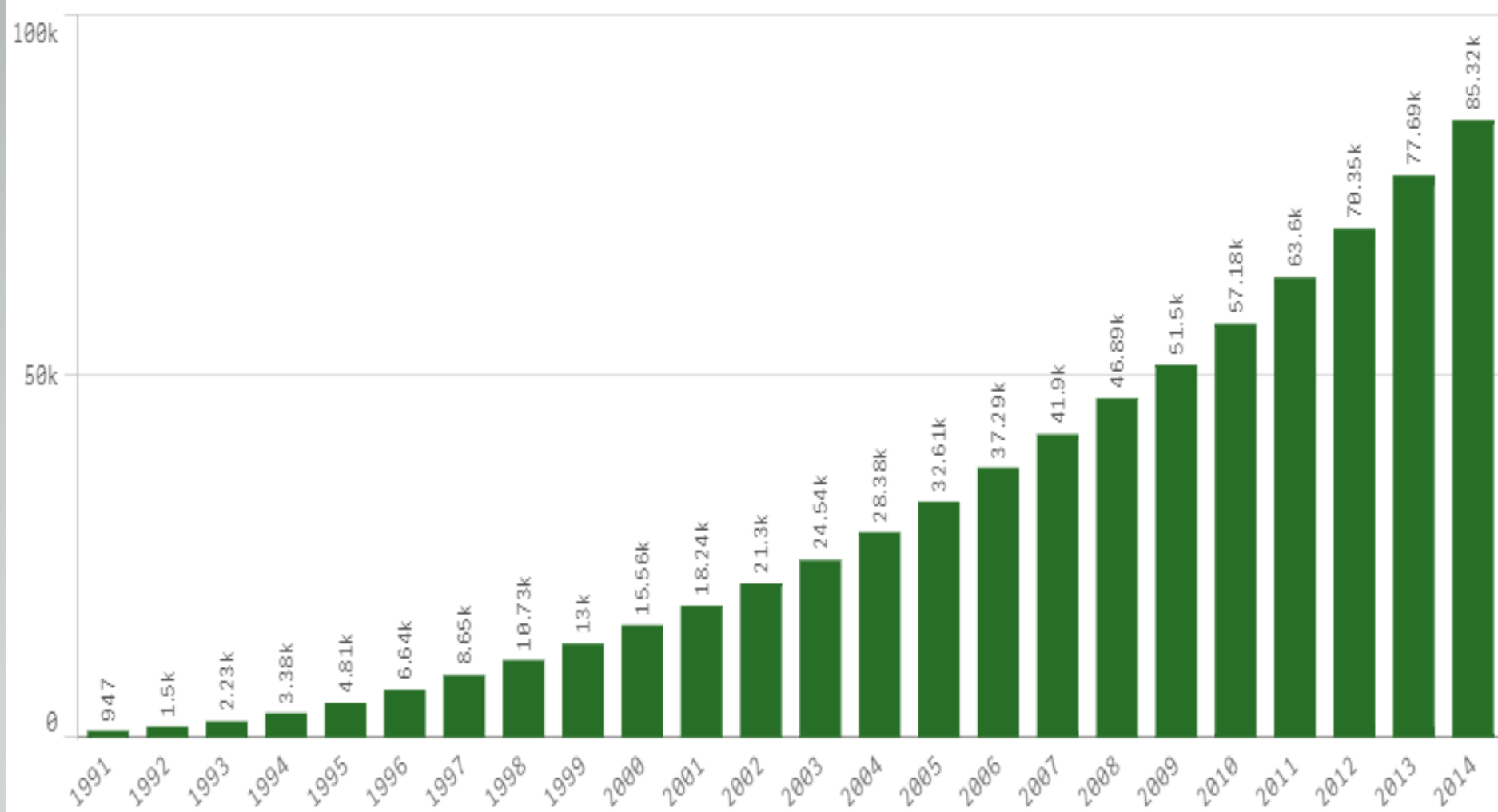


GAMMA KNIFE IN IAC-VSs

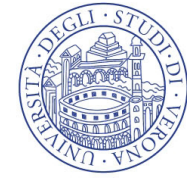
- **ADVANTAGES FOR THE PATIENT**
 - **ADVANTAGES FOR HEALTH SYSTEM**
-
- **NON-INVASIVE TECHNIQUE**
 - **DAY SURGERY PROCEDURE**
 - **HIGH NET INCOME**
(€ 7,500, 65% net income for the N.H.S.)



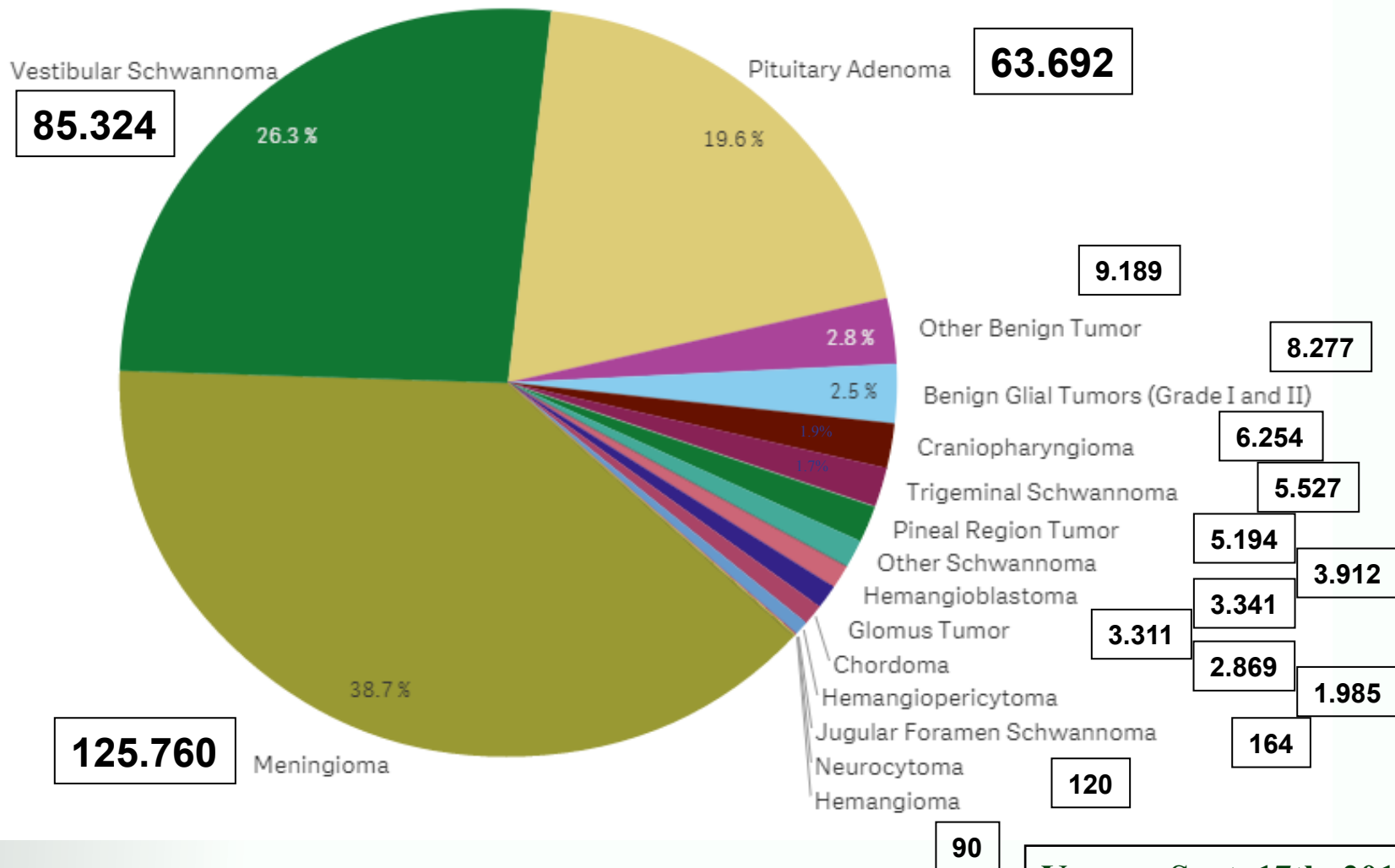
VSs – CUMULATIVE PATIENTS TREATED WORLDWIDE WITH GK



Verona, Sept. 17th, 2015



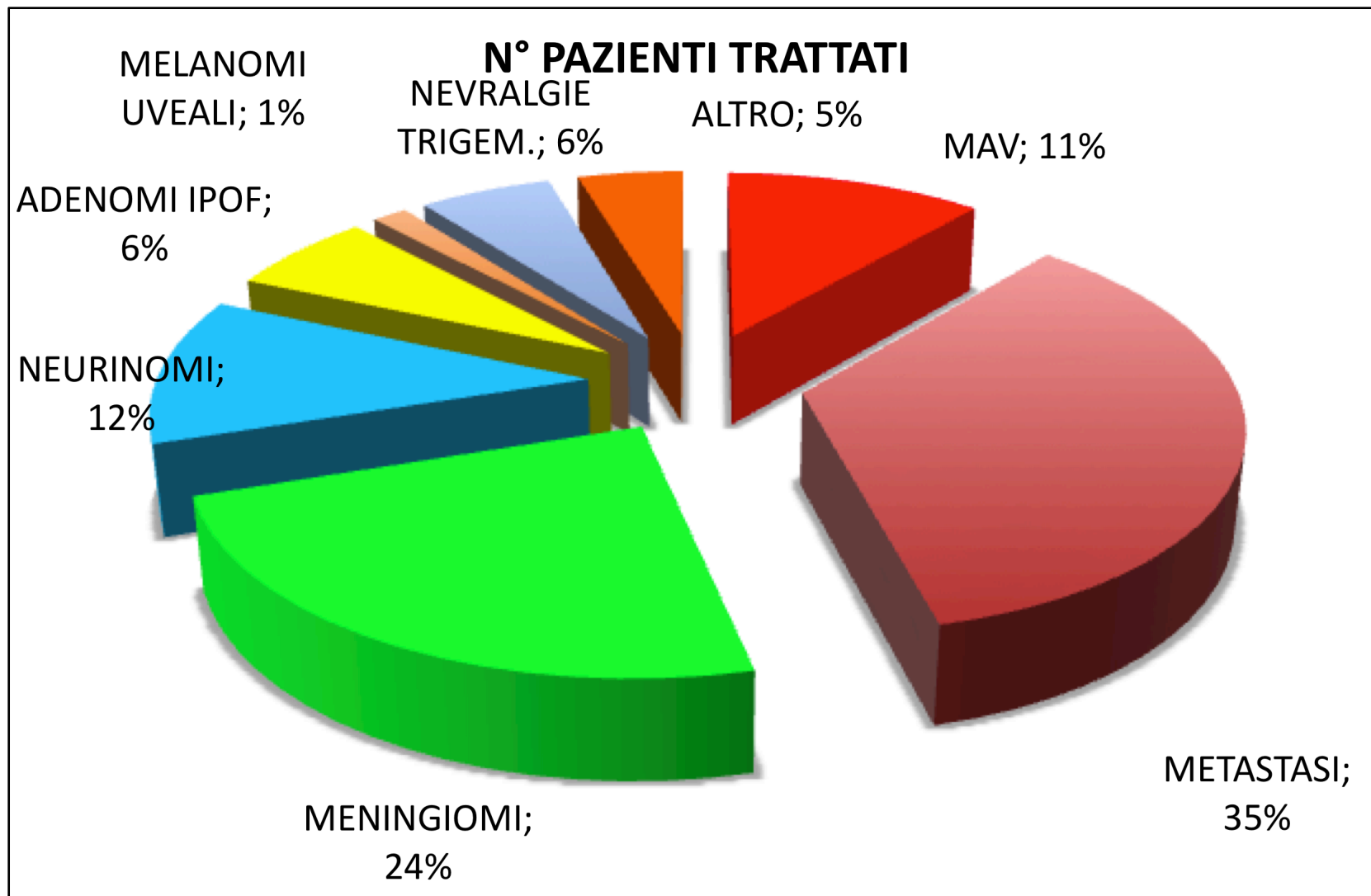
BENIGN TUMORS – CUMULATIVE PATIENTS TREATED WORLDWIDE WITH GK



Verona, Sept. 17th, 2015



GAMMA KNIFE IN VSs – AOUI VERONA





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VERONA



Otology & Neurotology
32:834–837 © 2011, Otology & Neurotology, Inc.

Stereotactic Radiosurgery for Vestibular Schwannomas: A Survey of Current Practice Patterns of Neurotologists

*Michael A. German, *Shawn Zardouz, *Mehdi K. Sina, *Kasra Ziai,
and *†Hamid Reza Djalilian

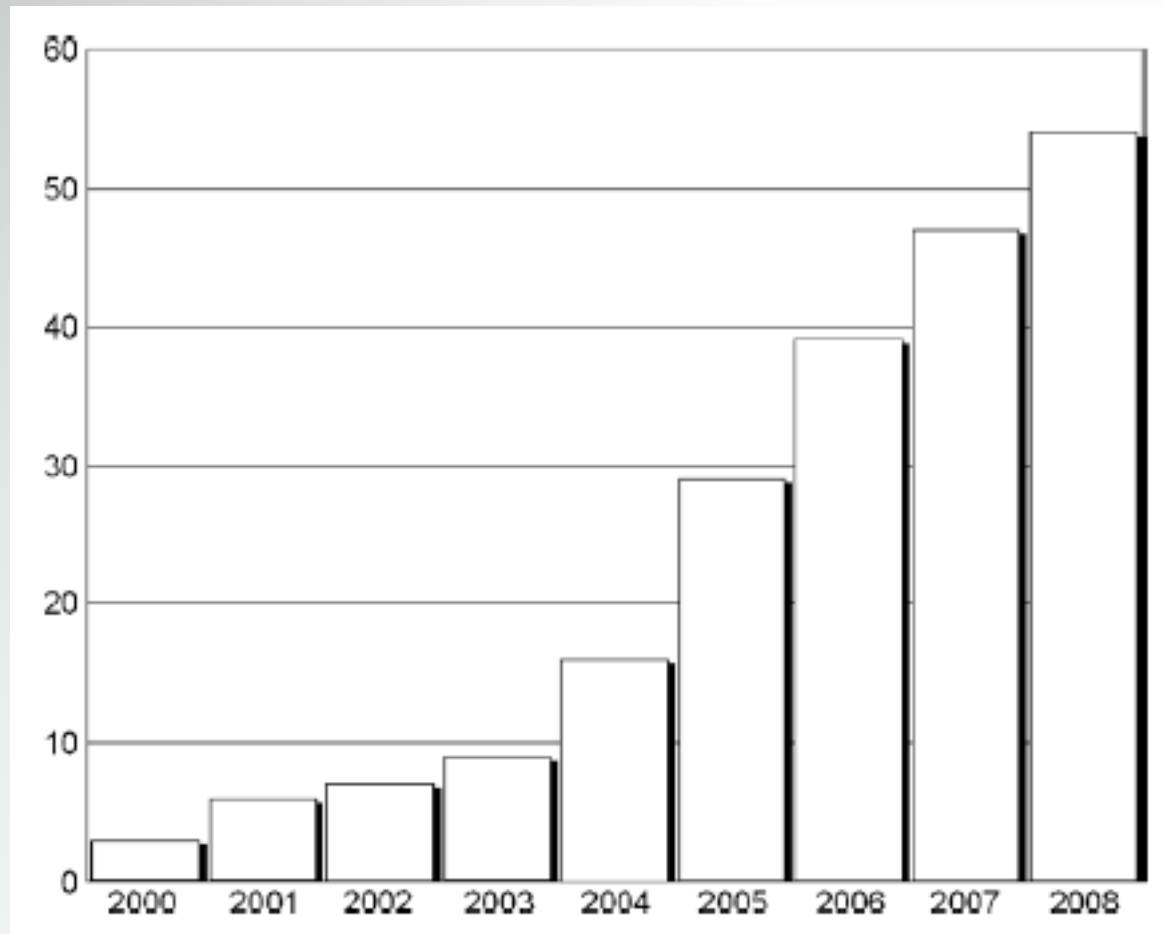
**Division of Neurotology and Skull Base Surgery, Department of Otolaryngology-Head and Neck Surgery;
and †Department of Biomedical Engineering, University of California, Irvine, California, U.S.A.*

This article discusses the results of a cross-sectional survey of members of the American Neurotology Society (ANS) that was conducted to ascertain practitioner attitudes and behaviors pertaining to radiation as a treatment modality for VS.

Verona, Sept. 17th, 2015



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Cumulative number of neurotology performing SRS between 2000 and 2008

The maximum size recommended for radiation by Members of the American Neurotology Society who perform SRS for VSs was 2.5 cm



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VERONA



GAMMA KNIFE IN VSs

INDICATIONS



Radiosurgery Practice Guideline Initiative

Stereotactic Radiosurgery for Patients with

Vestibular Schwannomas

Radiosurgery Practice Guideline Report #4-06

ORIGINAL GUIDELINE: May 2006

MOST RECENT LITERATURE SEARCH: March 2006

This practice guideline, together with a report on “Vestibular Schwannoma Management” is an original guideline approved by the IRSA® (International RadioSurgery Association) Board of Directors and issued in May 2006.

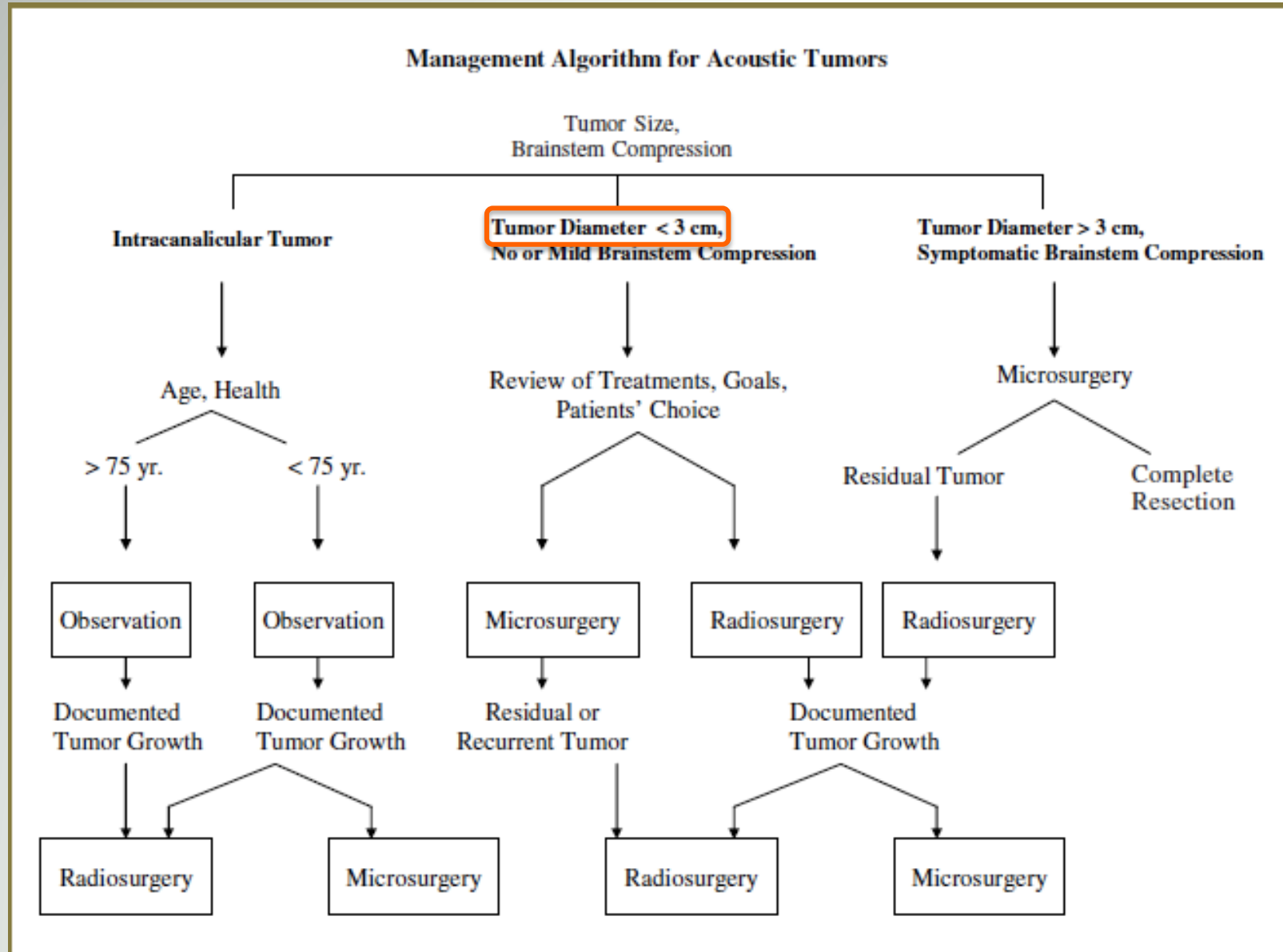
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Management Algorithm for Acoustic Tumors





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VERONA



GAMMA KNIFE IN VSs

OUR EXPERIENCE

February 1993 – August 2015

TREATED PATIENTS:	9,100
VSs:	1,013

Verona, Sept. 17th, 2015



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GAMMA KNIFE IN VSs

OUR EXPERIENCE

Reprint from

J. Kanzaki, M. Tos, M. Sanna, D.A. Moffat, T. Kunihiro, Y. Inoue (Eds.)

Acoustic Neuroma: Consensus on Systems for Reporting Results

Gamma Knife Radiosurgery in Vestibular Schwannomas: Clinical and Radiological Impact on the Tumor Course

MASSIMO GEROSA, ANTONIO NICOLATO, ROBERTO FORONI, and
ALBINO BRICOLO

2002

Verona, Sept. 17th, 2015



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VERONA



GAMMA KNIFE IN VSs

OUR EXPERIENCE

J Neurosurg 113:128–135, 2010

Gamma Knife surgery in vestibular schwannomas: impact on the anterior and posterior labyrinth

Clinical article

MASSIMO GEROSA, M.D.,¹ NAZARENA MESIANO, M.D.,² MICHELE LONGHI, M.D.,¹
ANTONIO DE SIMONE, PH.D.,¹ ROBERTO FORONI, PH.D.,¹ ANGELA VERLICCHI, M.D.,¹
BRUNO ZANOTTI, M.D.,³ AND ANTONIO NICOLATO, M.D.¹

	%
• HEARING PRESERV. (G-R 1/2)	66.0
• TGC	93.0
• FACIAL IMPAIRMENT	2.7

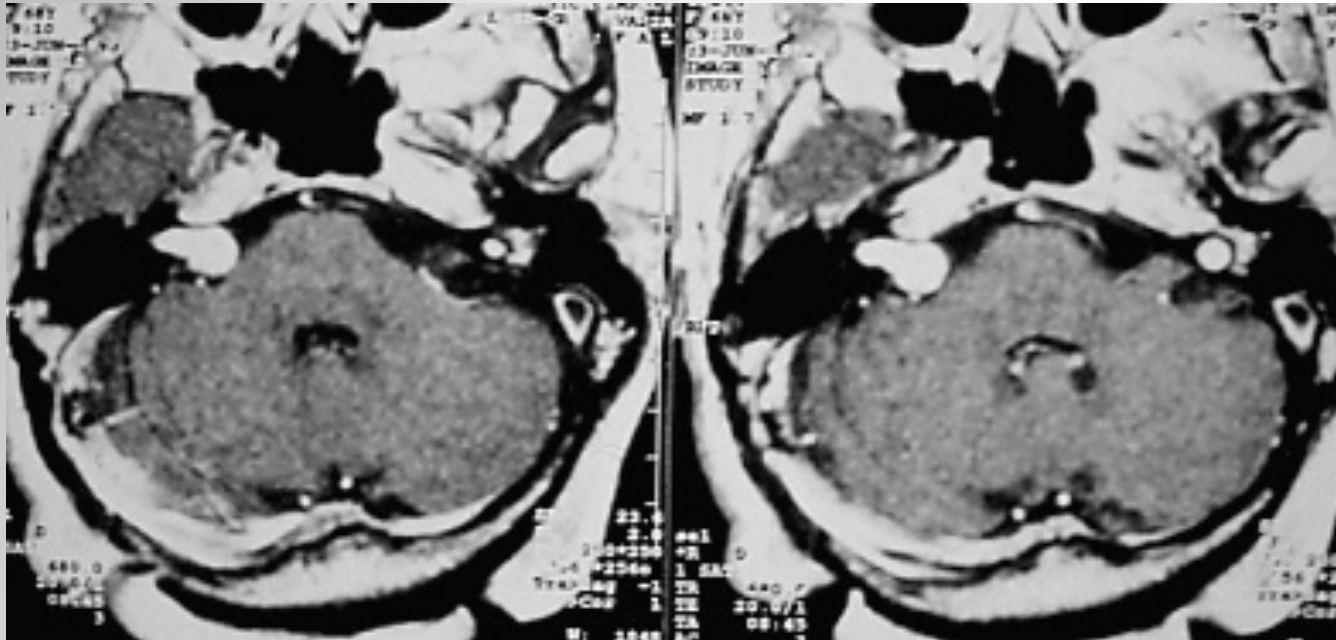
Verona, Sept. 17th, 2015



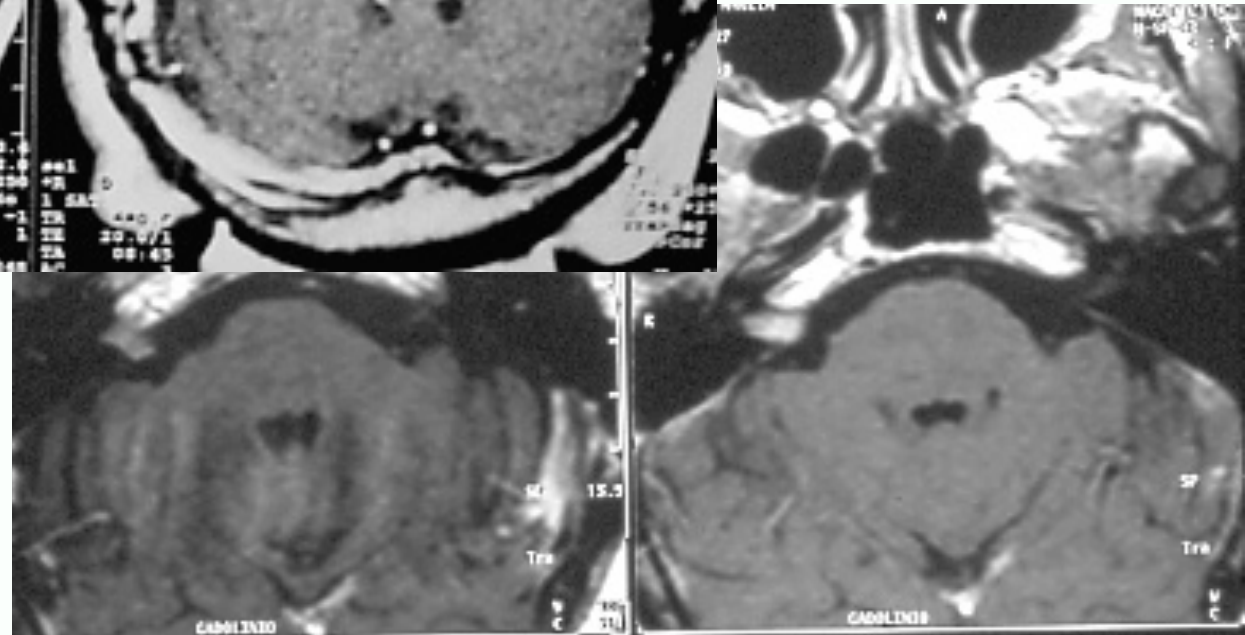
GAMMA KNIFE IN VSs

OUR EXPERIENCE

PRE-GK MRI



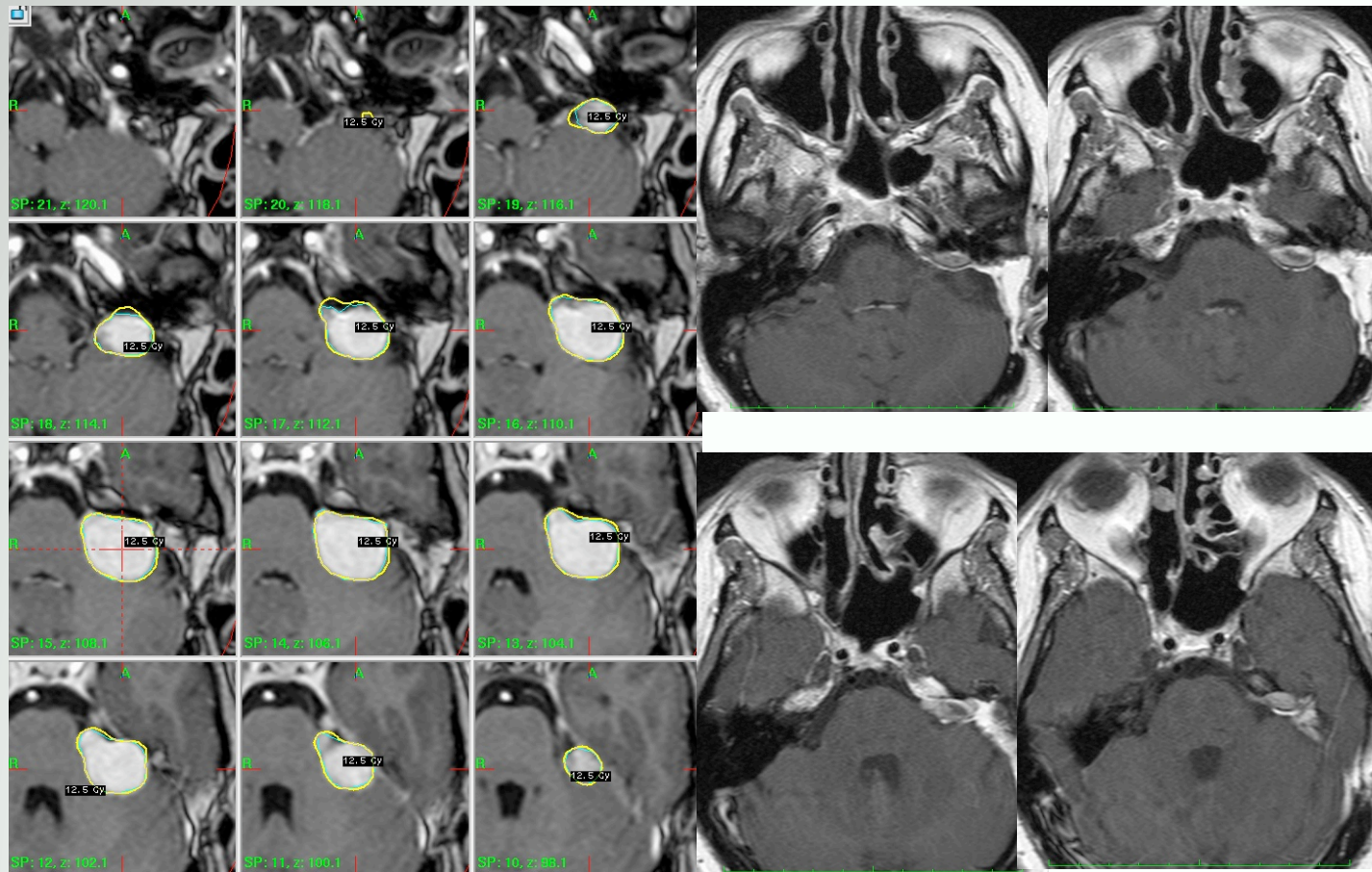
**POST-GK MRI
AT 50 mos.**





GAMMA KNIFE IN VSs

OUR EXPERIENCE



**MR on
GK day**

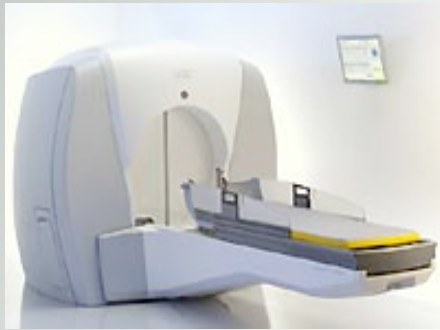
**MR FU
AT 37
mos.**

Verona, Sept. 17th, 2015



GAMMA KNIFE IN VSs

SUMMARY OF RECENT SRS LITERATURE



Author	N° Pts	Device	FU period	LTC %	Hear. Preser.	Facial N. Preserv.	Malign. Transfor.
Boari 2014	379	GK	Med.69.5m	97.1	49%	98,9%	0.0
Hasegawa 2013	440	GK	Med.12.5y.	93.0 (92% \geq 10y.)	43% at 5y.	99.7	Annual incidence: 0,02



COMPARISON MS vs. SRS

Author	Type of study	TC %		Hear. %		Facial N %		Preser. %	Cost-effectiveness
		MS	SRS	MS	SRS	MS	SRS		
Sarmiento 2013 (Ø ≤ 3 cm)	Focused re: • 3 prospec, • 6 retrosp, • Variuos case series	94-100	90.5-100	0-5	40-68	66-83	98-100	MS: • over 2 times higher than SRS. • 0.5% postop. mortality	
Wolbers 2013 (Ø ≤ 3 cm)	Focused re: • 6 cohort studies selected	—	—	—	Better Hear. preser.	—	Better Facial N. outcome	SRS: • Better QOL • No mortality • No surg/ anaes. compl.	
Myrseth 2009 (Ø ≤ 2.5cm)	Prospective open, non-randomized study	Compa rable	Compara ble	0.0	68.0 at 2 y.	53.6	98.3	SRS: Better QOL at 2y (Glasgow Benefit Inventory questionnaire)	

MS: microsurgery; SRS: stereotactic radiosurgery



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VERONA



COMPARISON MS vs. SRS

CONCLUSIONS

Sarmiento JM, et al. IMPROVING OUTCOME IN PATIENTS WITH VESTIBULAR SCHWANNOMAS. MICROSURGERY *versus* RADIOSURGERY (*focused review*).

J NEUROSURG SCI 2013;57:23-44

SRS can be considered as the the primary modality of choice for treatment of most vestibular schwannomas that are <3cm

Verona, Sept. 17th, 2015



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COMPARISON MS vs. SRS

CONCLUSIONS

Open Access

Research



What intervention is best practice for vestibular schwannomas? A systematic review of controlled studies

John G Wolbers,¹ Alof HG Dallenga,¹ Alejandra Mendez Romero,² Anne van Linge³

BMJ Open 2013;**3**:e001345. doi:10.1136/bmjopen-2012-001345
(*focused review*).

The available evidence indicates radiosurgery to be the best practice for solitary vestibular schwannomas up to 30 mm in cisternal diameter.

Verona, Sept. 17th, 2015



COMPARISON MS vs. SRS

CONCLUSIONS

VESTIBULAR SCHWANNOMA: SURGERY OR GAMMA KNIFE RADIOSURGERY? A PROSPECTIVE, NONRANDOMIZED STUDY

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Received, May 16, 2008.

Accepted, October 23, 2008.

OBJECTIVE: To conduct a prospective, open, nonrandomized study of treatment-associated morbidity in patients undergoing microsurgery or gamma knife radiosurgery (GKRS) for vestibular schwannomas.

METHODS: Ninety-one patients with vestibular schwannomas with a maximum tumor diameter of 25 mm in the cerebellopontine angle were treated according to a prospective protocol either by GKRS (63 patients) or open microsurgery (28 patients) using the suboccipital approach. Primary end points included hearing function, according to the Gardner-Robertson scale, and facial nerve function, according to the House-Brackmann scale at 2 years. Clinical data included a balance platform test, score for tinnitus and vertigo using a visual analog scale, and working ability. Patients responded to the quality-of-life questionnaires Short-Form 36 and Glasgow Benefit Inventory.

RESULTS: Three elderly GKRS patients withdrew; all remaining patients were followed for 2 years. Both primary end points were highly significant in favor of GKRS ($P < 0.001$). Evidence of reduced facial nerve function (House-Brackmann grade 2 or poorer) at 2 years was found in 13 of 28 open microsurgery patients and 1 of 60 GKRS patients. Thirteen of 28 patients who underwent surgery had serviceable hearing (Gardner-Robertson grade A or B) preoperatively, but none had serviceable hearing postoperatively. Twenty-five of 60 GKRS patients had serviceable hearing before treatment, and 17 (68%) of them had serviceable hearing 2 years after treatment. The tinnitus and vertigo visual analog scale score, as well as balance platform tests, did not change significantly after treatment, and working status did not differ between the groups at 2 years. Quality of life was significantly better in the GKRS group at 2 years, based on the Glasgow Benefit Inventory questionnaire. One GKRS patient required operative treatment within the 2-year study period.

CONCLUSION: This is the second prospective study to demonstrate better facial nerve and hearing outcomes from GKRS than from open surgery for small- and medium-sized vestibular schwannomas.

KEY WORDS: Facial nerve function, Gamma knife radiosurgery, Hearing preservation, Microsurgery, Quality of life, Vestibular schwannoma

This prospective study demonstrates better facial nerve and hearing outcomes from GKRS than from open surgery for small- and medium-sized (maximum tumor $\varnothing \leq 25$ mm) vestibular schwannomas.

Neurosurgery 2009

Prospective open, non-randomized study

Verona, Sept. 17th, 2015



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VERONA



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE

February 1993 – August 2015

TREATED PATIENTS:	9,100
VSs:	1,013
IAC-VSs:	107

IAC: internal acoustic canal

Verona, Sept. 17th, 2015



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VERONA



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE

February 1993 – August 2015
107 TREATED PATIENTS

M/F	54/53
MEAN AGE (Range)	56.1 yrs. (12–80)
SURGICAL RESIDUAL	13/107 pts. (12.1%)
MEAN VOLUME (RANGE)	0.19 mL (0.04–0.70)
Mean PI/ PD/ MD /#ISOCENTERS	52.1% / 12.1Gy / 23.3Gy / 3.7

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE

February 1993 – August 2012

66 TREATED PATIENTS WITH AT LEAST 36-MONTH-F-U

MEDIAN SURVIVAL TIME	52.6 Mos
ALIVE	66/66 (100%) Pts.
TGC	65/66 (98,5%)
HEARING PRESERVATION	22/47 (46.8%) Pts. (47 Pts. with GR I-III pre-GK)
FACIAL PRESERVATION	65/66 (98,5%)*
OTHER WORSENING (tinnitus, ataxia)	3/66 (4.5%)
MALIGNANT TRANSFORMATION	0/66

***mild spasm of facial nerve**

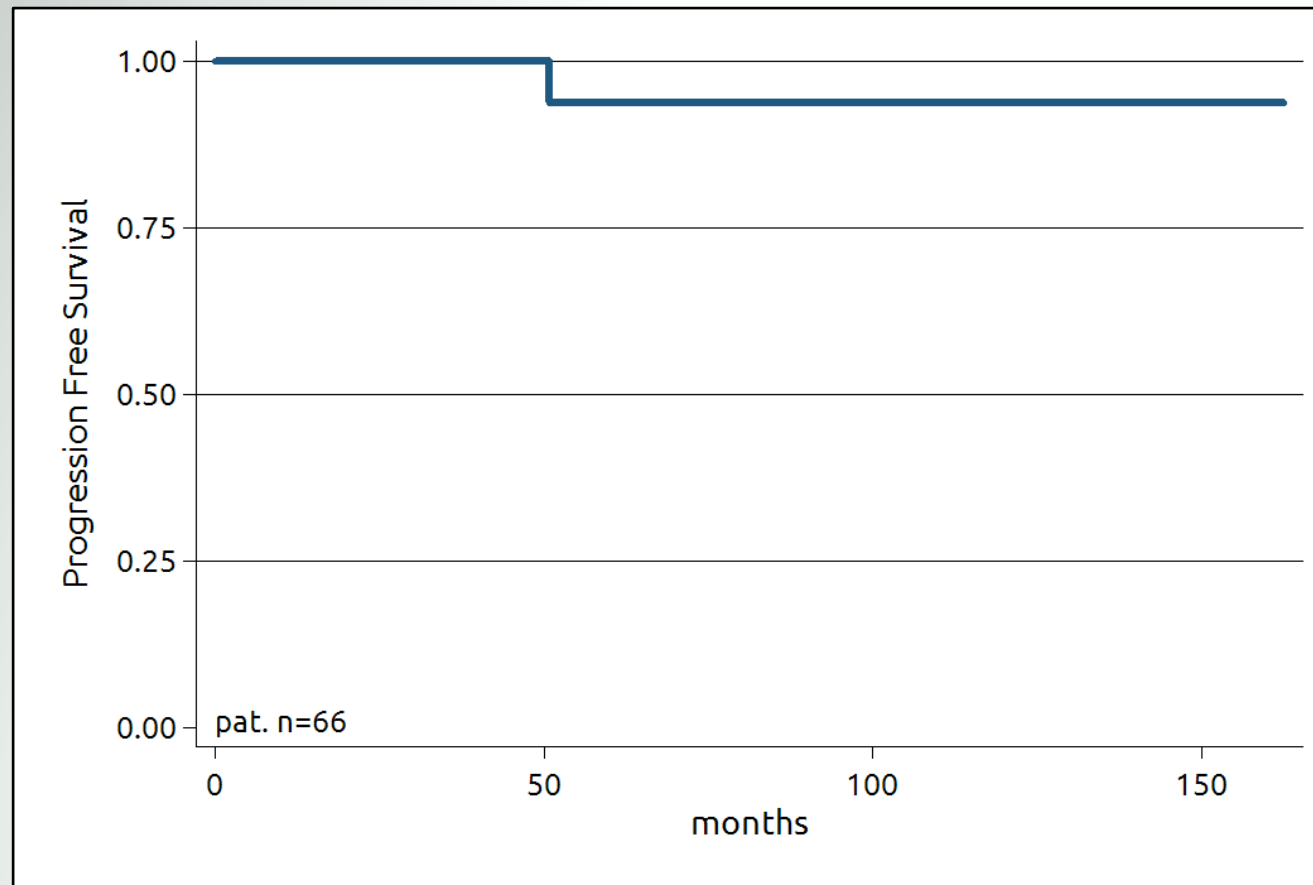


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GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE



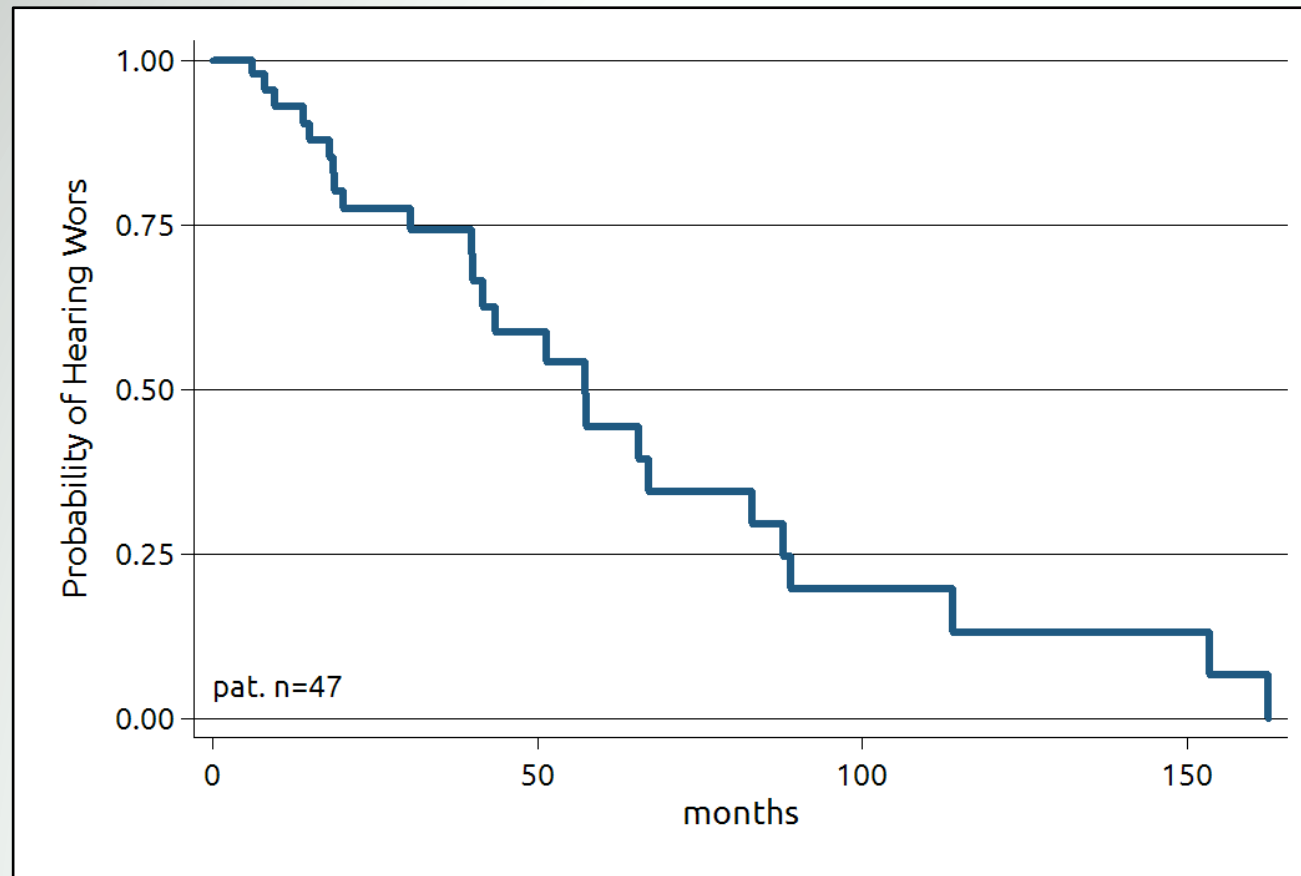
ACTUARIAL PFS RATE:

96% at 10 Y.



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE



ACTUARIAL HEARING PRESERVATION RATE: 43% at 5 Y.



GAMMA KNIFE IN IAC-VSs

STATISTICAL ANALYSIS: prognostic factors influencing hear.wors.

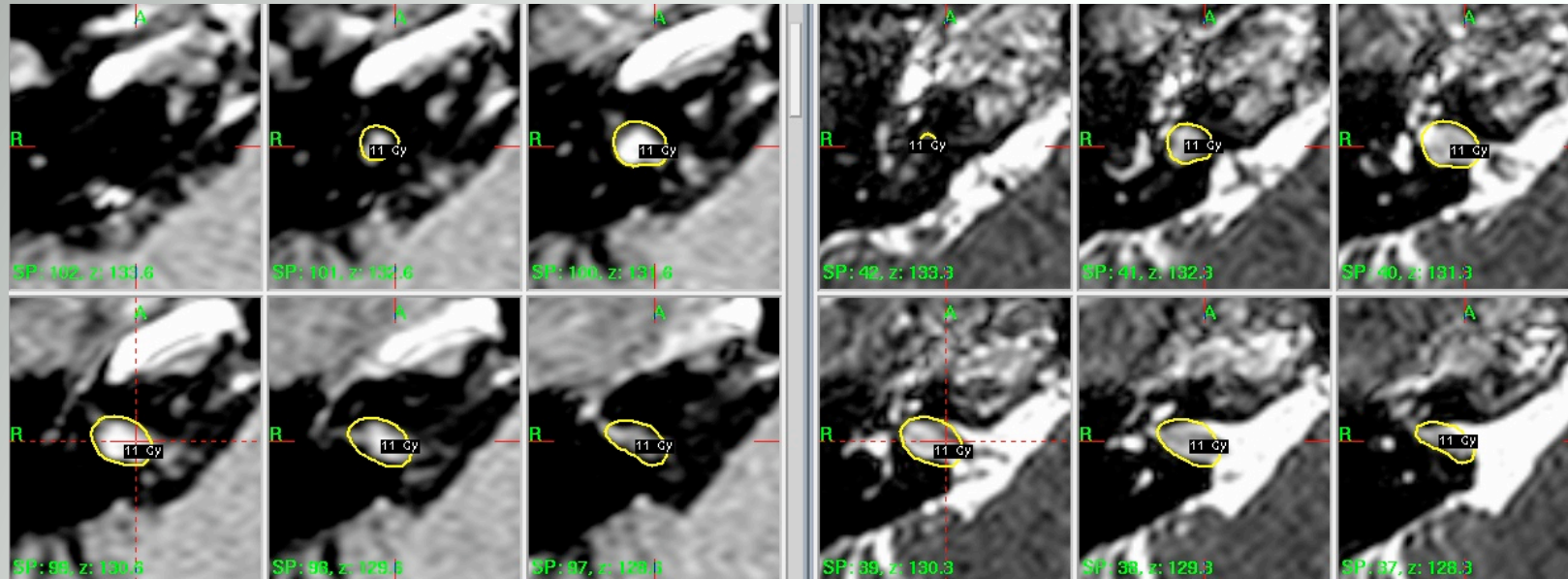
Independent variables	Odds Ratio	p-value	# Observ.	Statistically significant
Pre-GK VS increased volume const	1.05 1.428571	0.946 0.469	32	N
Post-GK intra-VS necrosis time const	1.011856 .5319617	0.049 0.228	38	Y
Average dose to the cochlea const	.8609878 1.862945	0.509 0.497	23	N
VS volume (mL) const	.3367242 1.285514	0.679 0.665	45	N
Prescription dose (Gy) const	.6008056 561.6	0.031 0.029	47	Y



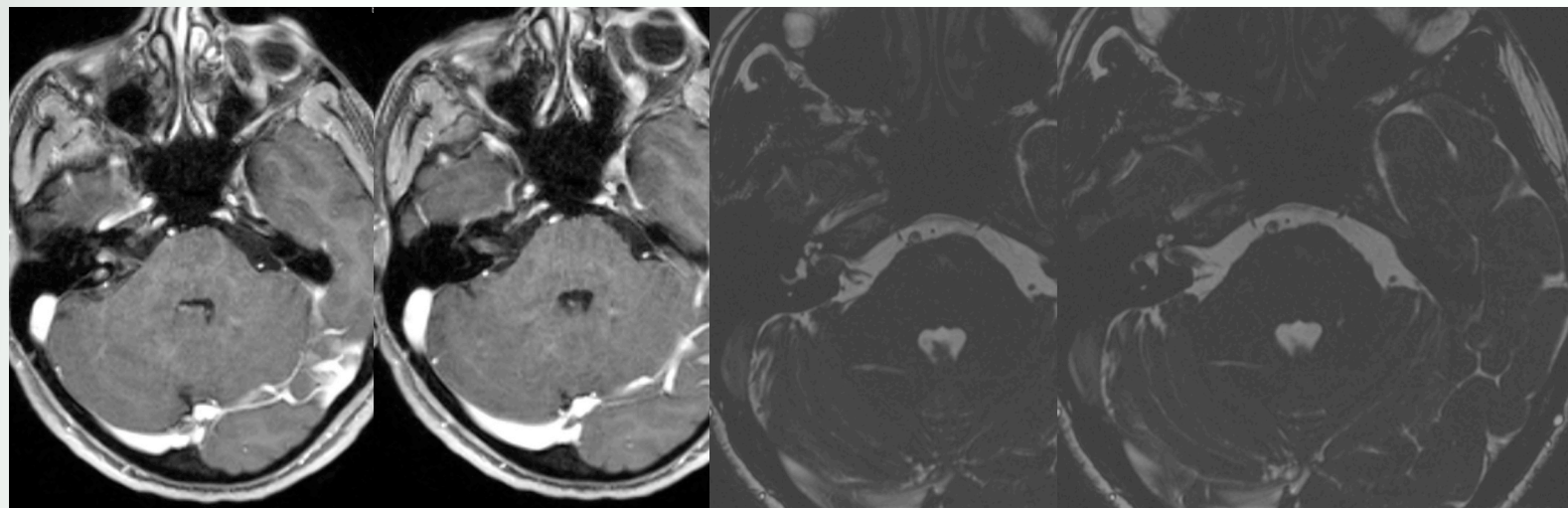
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**MR on
GK day,
Jan 2012**



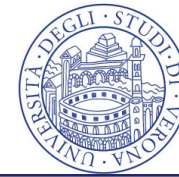
**MR FU AT
22 mos.**



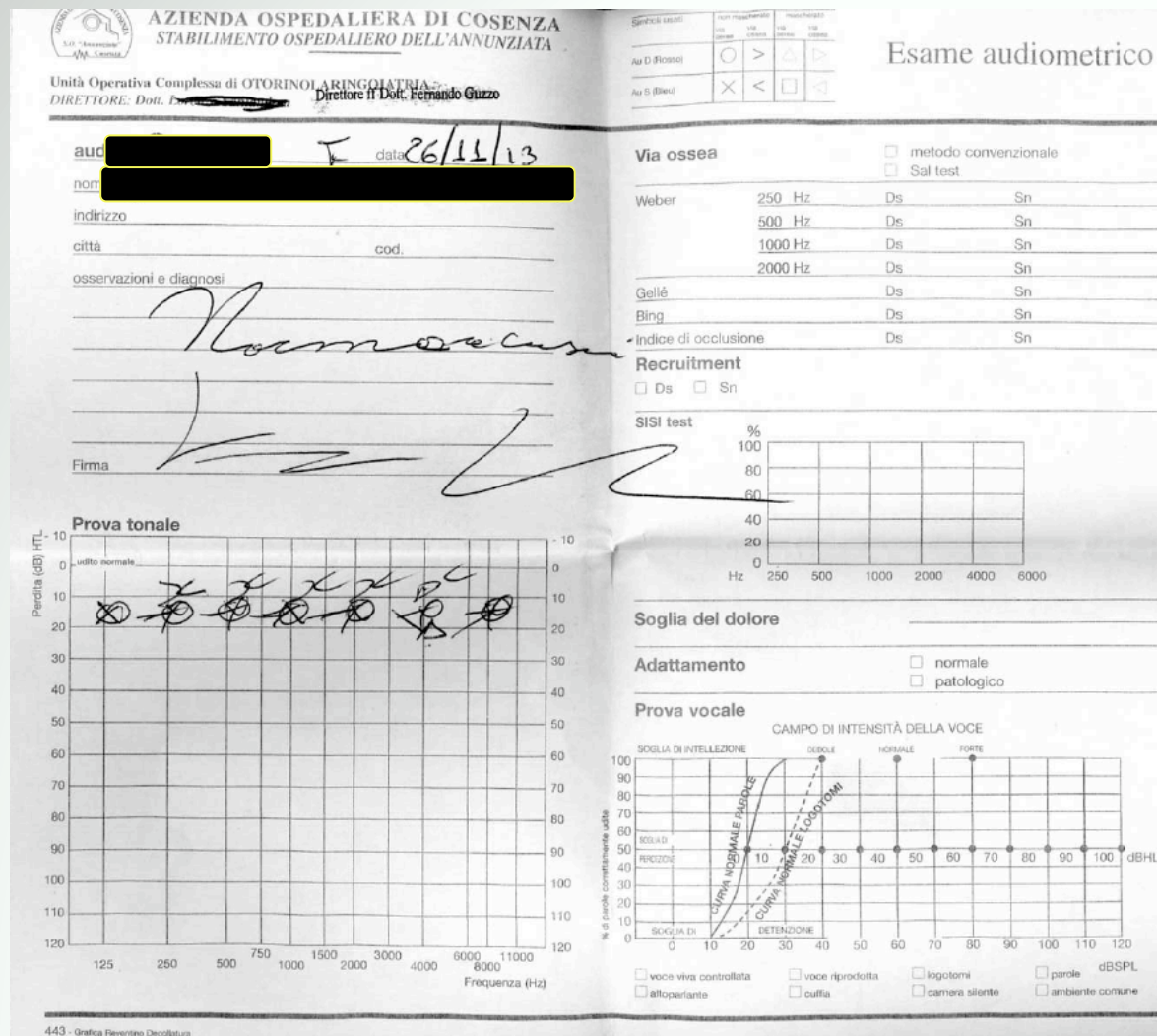
Verona, Sept. 17th, 2015



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GAMMA KNIFE IN IAC-VSs



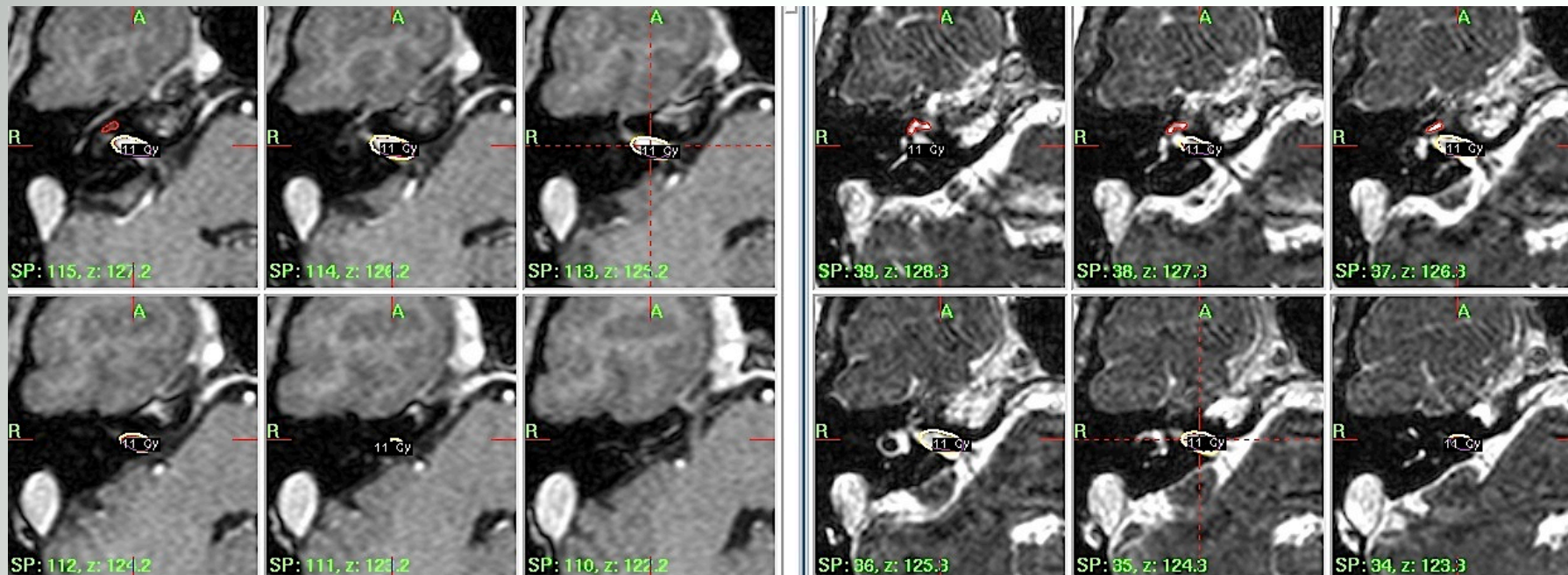
**AUDIOGRAM
on Nov, 2013**

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE



MR on GK day, October 4th, 2012

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

OUR EXPERIENCE

AUDIOGRAM
on Dec. 2nd,
2013

REGIONE VENETO
ULSS10 VERONA ORIENTALE
AZIENDA ULSS N. 10 "VENETO ORIENTALE"
PRESIDIO OSPEDALIERO DI PORTOGRUARO
UNITÀ OPERATIVA SEMPLICE DI OTORINOLARINGOIATRIA

AUDIOGRAMMA N. [redacted] data 2/12/13

Webber

	non mascherato		mascherato	
	via aerea	via ossea	via aerea	via ossea
Au. D. (rosso)	○	>	△	▷
Au. S. (blu)	×	<	□	◁

250 Hz Ds Sn
500 Hz Ds Sn
1000 Hz Ds Sn
2000 Hz Ds Sn

PROVA TONALE *Neurifonia dx*

SONDA IN AU DS

TIMPANOGRAMMA

SONDA IN AU SIN.

C: cc P: daPa C: cc P: daPa

RIFLESSO STAPEDIALE

	500 Hz	1000 Hz	2000 Hz	4000 Hz	Rumore bianco
	Contra				
I p s i					

osservazioni e diagnosi
L'ascolto Neurosensoriale Biotale.
Max discriminazione vocale e 40 dB.

firmo *[Signature]*
AZIENDA ULSS N. 10 "VENETO ORIENTALE"
Presidio Ospedaliero di Portogruaro
Ambulatorio di Otorinolaringoiatria
Dr. MARIANO COLLETTA
PR 1867 CN

Med ST-085061 - 11 Bezzolo N.B. CONSERVARE E RIPRESENTARE AGLI ESAMI SUCCESSIVI

N.B. CONSERVARE E RIPRESENTARE AGLI ESAMI SUCCESSIVI

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

Author	N° Pts	Device	FU period (Median)	LTC %	Hear. Preser.	Facial N Preser.	Prog.Fact. for hear. Deterior.
Kim 2013	60	GK	62.0 m.	100 clinical TC	57.0 PTA \leq 50 dB SDS \geq 50%	—	• TVE \geq 20%
Present series	66 47 p. GR I-III	GK	52.6 m.	98.5 (96%at10y)	46.8% (43% at5y.)	99.7	• Necrosis time • PD (Gy)

PTA: pure tone average
SDS: speech discrimination score
TVE: transient volume expansion



GAMMA KNIFE IN IAC-VSs

Original Article 157

Facial Nerve Outcome after Vestibular Schwannoma Resection: A Comparative Meta-Analysis of Endoscopic versus Open Retrosigmoid Approach

Abdullah Alobaid¹ Mohammed Aref¹ Michael Ross Bennardo¹ Forough Farrokhyar² Kesava Reddy¹

¹Division of Neurosurgery, McMaster University, Hamilton, Ontario, Canada

²Department of Surgery, McMaster University, Hamilton, Ontario, Canada

Address for correspondence Abdullah Alobaid, MD, FRCSC, 96 St. Patrick Street, Apt. 1210, Toronto, Ontario M5T1V2, Canada (e-mail: abdullah.alobaid@medportal.ca).

J Neurol Surg B 2015;76:157–162.

Verona, Sept. 17th, 2015



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Table 1 Final studies included in the analysis

	Study	No. of patients included in final analysis
Open approach	Samii et al ⁷⁻¹⁰	962
	Colletti and Fiorino ¹²	103
	Moffat et al ¹³	50
	Jung et al ¹⁴	30
	Mazzoni et al ¹⁵	150
	Tonn et al ¹⁶	508
	Strauss ¹⁷	22
	Lassaletta et al ¹⁸	65
	Maw et al ¹⁹	40
	Yamakami et al ²⁰	50
	Darwish et al ²¹	97
	Zhang et al ²²	105
	Samii et al ²³	200
	Sinha and Sharma ²⁴	58
	Veronezi et al ²⁵	20
	Yang et al ¹¹	110
	Chen et al ²⁶	103
	Chen et al ²⁷	145
	Zhao et al ²⁸	89
	Di Maio et al ²⁹	47
Gerganov et al ³⁰	53	
Endoscopic approach	Göksu et al ³¹	32
	Magnan et al ³²	119
	Kabil and Shahinian ³³	112
	Shahinian and Ra ⁵	527

META-ANALYSIS REVIEW FOR ALL ARTICLES DESCRIBING BOTH APPROACHES FOR VS FROM 1996 TO 2011:

- **21 OPEN APPROACH**
 - **4 ENDOSCOPIC APPROACH**
- SELECTED ARTICLES.**

Verona, Sept. 17th, 2015



GAMMA KNIFE IN IAC-VSs

Table 2 Summary of meta-analysis comparing open series with minimal access endoscopic series with 95% confidence interval

	Open retrosigmoid, % (95% CI)	Minimal access endoscopic retrosigmoid, % (95% CI)
Good facial nerve outcome (HB 1 or 2)	67.0% (61–73%)	94% (92–95%)
GTR	91% (80–98%)	97% (92–99%)
Meaningful hearing (PTA < 80 dB and speech discrimination > 20%)	22.6% (10.4–37.6%)	46% (38–54%)
CSF leak	8.2% (4.8–12.3%)	4.6% (2.3–7.2%)
Wound infection	1.3% (0.6–2.3%)	2.6% (1.5–4.0%)
Recurrence	5.4% (1.8–10.1%)	2.2% (1.3–3.4%)
Death	0.9% (0.3–2%)	0%

Facial nerve Outcome (HB 1)	98.0%-100.0%
LTC	90.5%-100.0%
Hearing Preservation (GR I-II)	40.0%-68.0%
Other surgical complications	0.0%
Death	0.0%



GAMMA KNIFE IN IAC-VSs

CONCLUSIONS

SRS IN VSs ALLOWS:

- VERY HIGH TGC
- POSSIBILITY OF HEARING PRESERVATION
- VERY HIGH RATE OF FACIAL NERVE PRESERVATION
- NO RISK OF “SURGICAL” COMPLICATIONS
- DAY SURGERY PROCEDURE
- HIGH NET INCOME FOR HEALTH SYSTEM



GAMMA KNIFE IN IAC-VSs

CONCLUSIONS

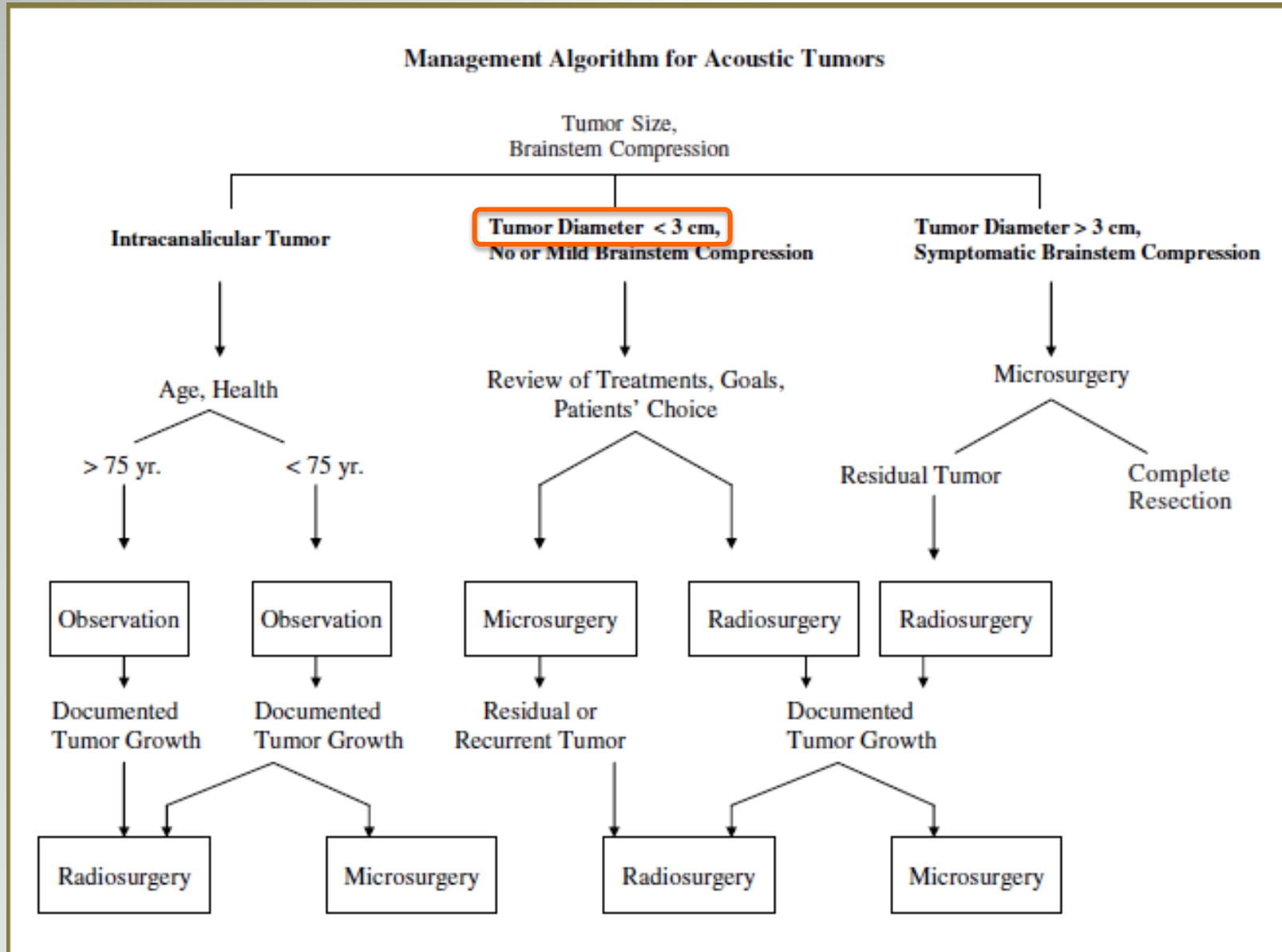
-THEN, NO NEED FOR FURTHER STUDIES ?
- $\emptyset \leq 2$ cm: GK/SRS
- $\emptyset \geq 3$ cm (USA), $\emptyset \geq 2.5$ cm (EC):
 - microsurgical endoscopic approach
- $2 \text{ cm} > \emptyset < 2,5 \text{ cm}$:
 - what is the goal standard for these VSs ?



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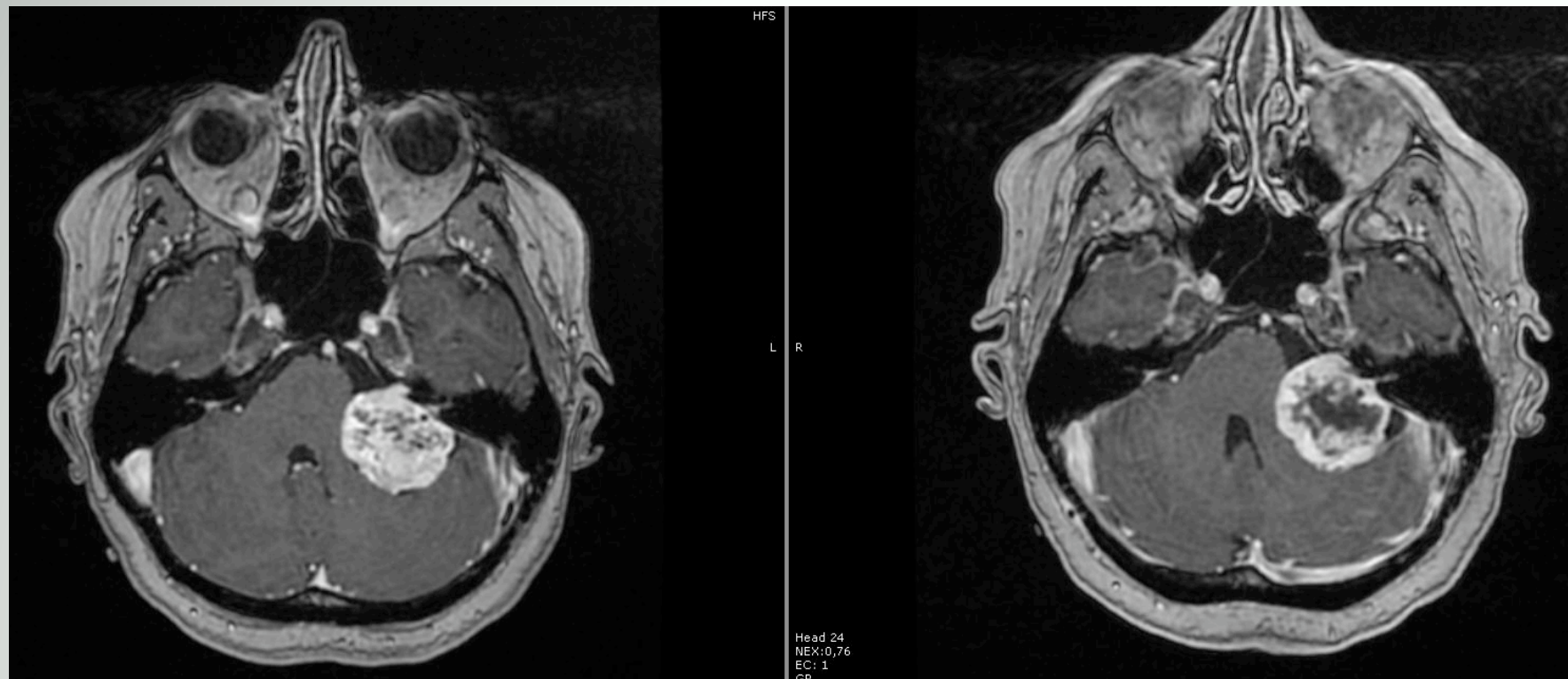
Management Algorithm for Acoustic Tumors





GAMMA KNIFE IN IAC-VSs

CONCLUSIONS



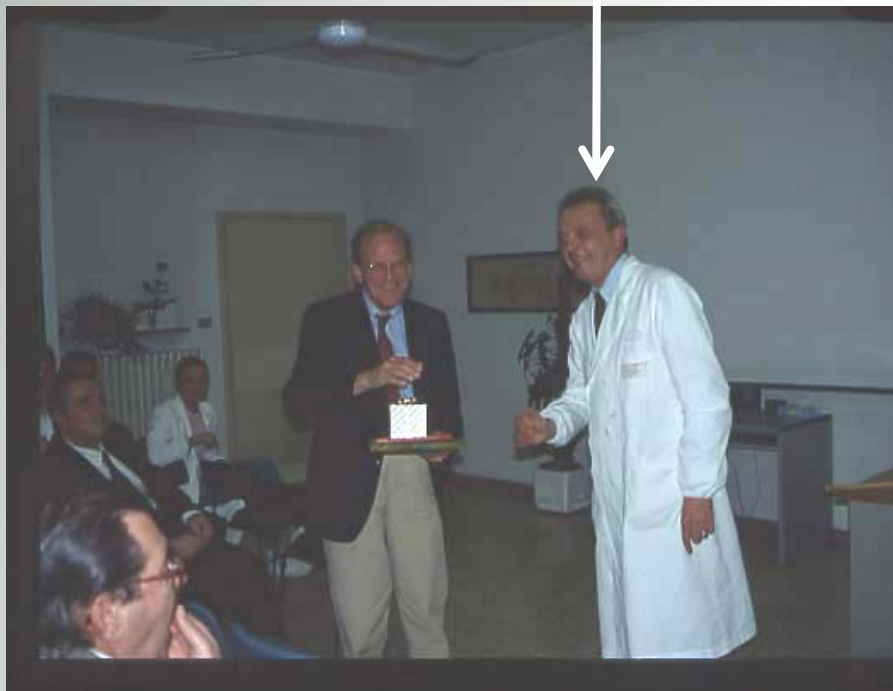
VS WITH $\emptyset > 3$ cm TREATED WITH PROTON BEAM:.....



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GAMMA KNIFE VERONA: 20 YEAR-RESULTS

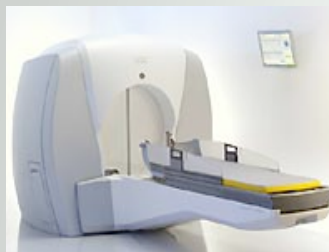


February 1993 – February 2013

Verona, 13 Dicembre 2013



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THANK YOU FOR THE ATTENTION