

Radiosurgery for Skullbase Meningioma

Dr Christos BOSKOS

Radiosurgeon / Radiation-Oncologist





I have no conflict of interest to disclose

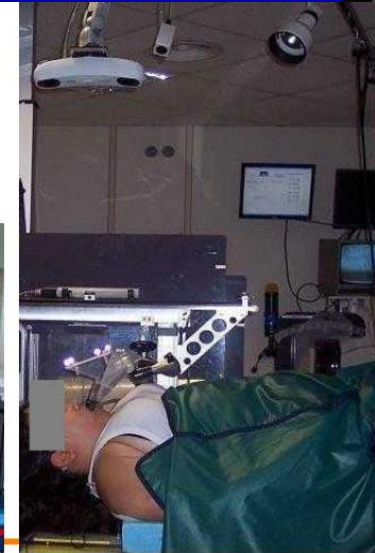
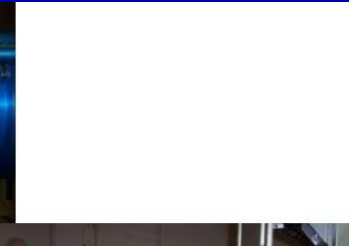
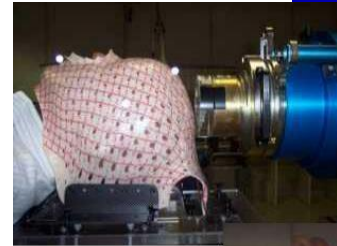
My experience in Charged Particles: **Protontherapy**

Centre Protontherapie Orsay (Paris) 2003 (resident)

Centre Protontherapie Orsay (Paris) 2006-2008

Le Centre de Protonthérapie d'Orsay

- Synchro-cyclotron (IPN/CNRS 1975)
- Dédié exclusivement au médical depuis 1991
- Traitements par protons de 200 Mev (intracrânien) ou 73 Mev (ophtalmologiques)
- 2800 patients traités depuis 1991



CENTRE DE PROTONTHERAPIE D'ORSAY

CPO



My experience in **Stereotactic Radiosurgery**

Saint Savvas Hospital (Athens) Elekta 2001-2005 (resident)

Hopital Pitie-Salpetriere (Paris) Varian BrainLab 2006-2009

UCLA (Los Angeles) Novalis BrainLab 2007

Iatropolis Radiosurgery Center (Athens) CyberKnife Accuray 2019

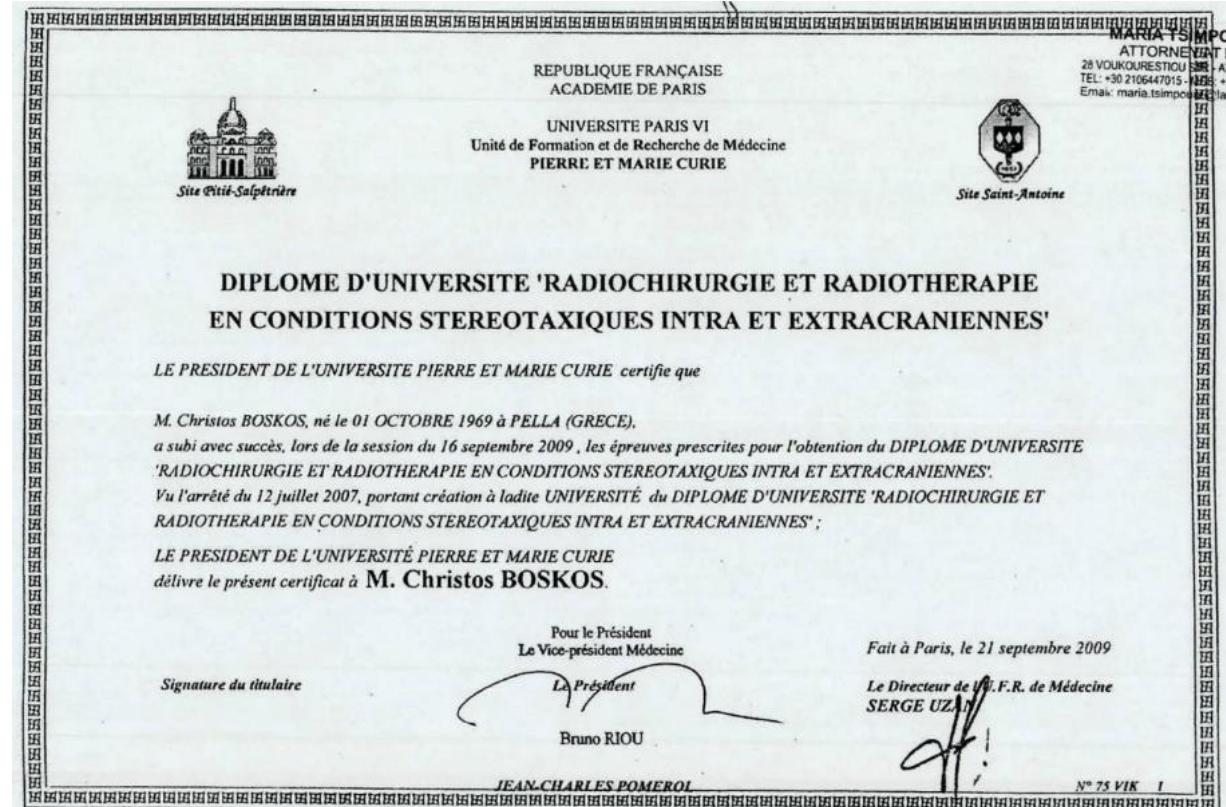
Metropolitan Clinic (Athens) EDGE Varian 2021

Saint Lukas Clinic (Thessaloniki) EDGE Varian 2024



Diplome Universite in Stereotactic Radiosurgery and Protontherapy

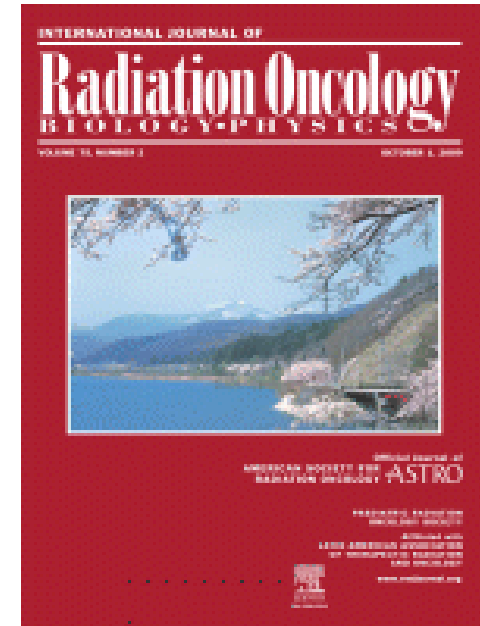
Universite Pierre and Marie Curie (Paris VI)
2010



Research in Protontherapy: Skullbase Meningioma

“Combined proton and photon conformal radiotherapy for intracranial atypical and malignant meningioma”,

[Christos Boskos](#)¹, [Loic Feuvret](#), [Georges Noel](#), [Jean-Louis Habrand](#), [Pascal Pommier](#), [Claire Alapetite](#), [Hamid Mammar](#), [Regis Ferrand](#), [Gilbert Boisserie](#), [Jean-Jacques Mazon](#)



“Focal” is better...

- Radiosurgery
- Protontherapy



But “Focal + radiobiology advantage” is...

- Carbon ions

far better



Challenges for Skullbase Meningiomas

- Meningioma most **common** benign CNS tumor
- **Increasing incidence** as population grow older
- **Imaging quality progression** (easier define **residual** and **relapse**)
- **Factors** affecting management decision (age, volume, location, Simpson grade)
- Skullbase is a high risk area for **surgical complications**

Skull base meningiomas



- **Female 74% and white race 79%**
- **Benign**
- **Atypical** meningiomas are not considered either benign or malignant. They can become **malignant**
- **Grade II and III** almost **7%** National Database (SEER)

skull base meningiomas had **better LC** when compared to those located at the **convexity** of the brain has been reported previously

Pou P., Biau J., Verrelle P., Lemaire J.J., El Ouadih Y., Chassin V., Magnier F., Dedieu V., Lapeyre M., Dupic G., et al. Long-Term Outcomes After Linac Radiosurgery for Benign Meningiomas. *Clin. Oncol.* **2020**

skull base meningiomas

Lower probability of high-grade histology when compared to those located at the rest of the brain

Maclean J., Fersht N., Short S. Controversies in Radiotherapy for Meningioma. *Clin. Oncol.* **2014**

Slower growth of skull base meningiomas

Hashimoto N., Rabo C.S., Okita Y., Kinoshita M., Kagawa N., Fujimoto Y., Morii E., Kishima H., Maruno M., Kato A., et al. Slower growth of skull base meningiomas compared with non-skull base meningiomas based on volumetric and biological studies: Clinical article. *J. Neurosurg.* **2012**

Lower MIB1 proliferation index in skull base meningiomas

McGovern S.L., Aldape K.D., Munsell M.F., Mahajan A., DeMonte F., Woo S.Y. A comparison of World Health Organization tumor grades at recurrence in patients with non-skull base and skull base meningiomas: Clinical article. *J. Neurosurg.* **2010**



skull base meningiomas **originate**

originate from a **variety of different structures** including, but are not limited to:

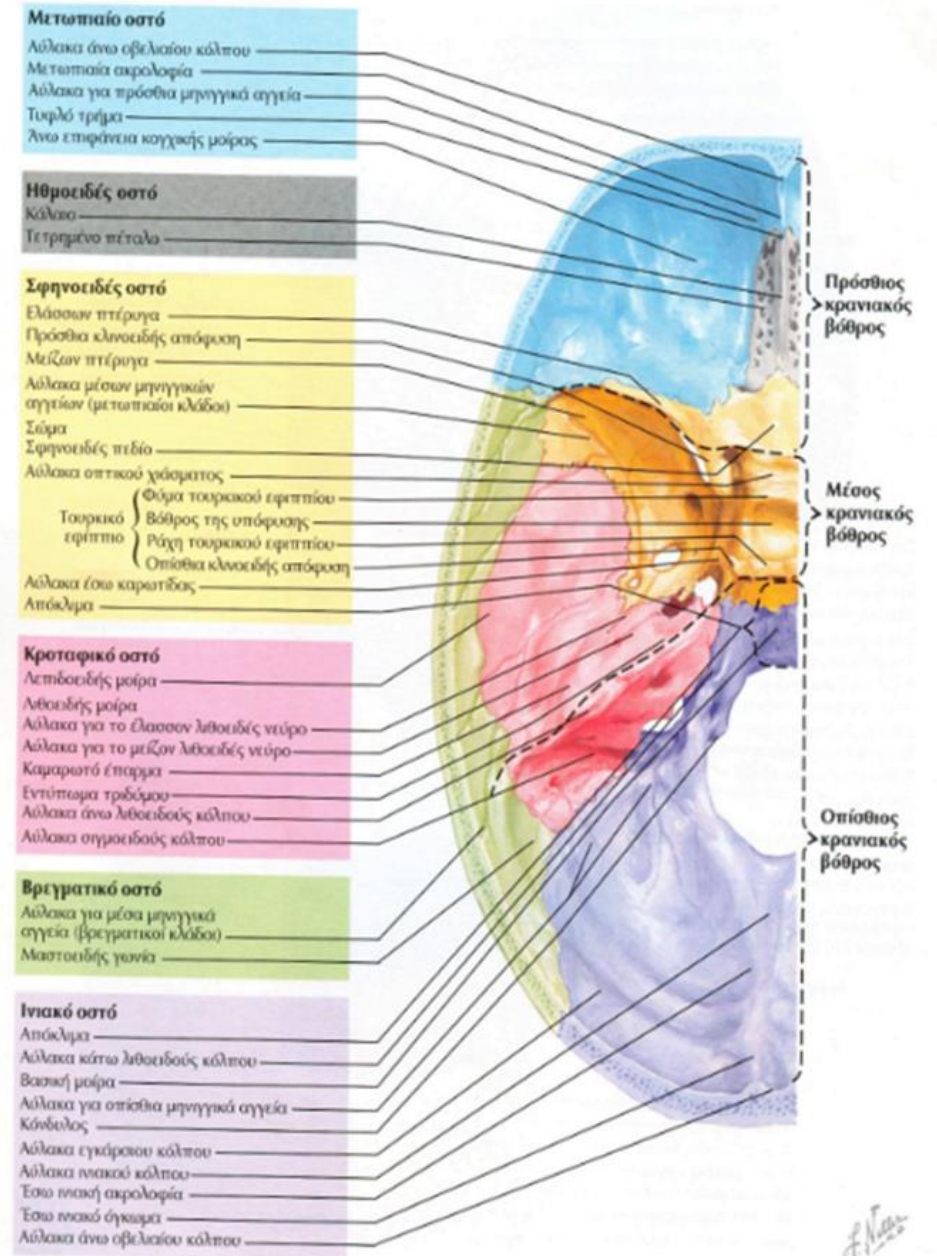
- clinoid processes,
- tuberculum sellae,
- dorsum sellae,
- sphenoid wing,
- petrous/petroclival area,
- falcotentorial region,
- cerebellopontine angle,
- foramen magnum.



Skull base locations

- Frontal fossa
- Median fossa
- Posterior fossa

- Cavernous sinus (CS)
- CerebroPontal Angle (CPA)
- Petro Clival (PC)



3754 skullbase meningiomas for SRS

- Cerebellum Pontine Angle 432
- Petro Clival 468
- Cavernous sinus 1272

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RESEARCH—HUMAN—CLINICAL STUDIES

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Long-term Tumor Control of Benign Intracranial Meningiomas After Radiosurgery in a Series of 4565 Patients

Santacrose, Antonio MD[†]; Walier, Maja Dipl Math[‡]; Régis, Jean MD, PhD[§]; Liščák, Roman MD, PhD[¶]; Motti, Enrico MD[¶]; Lindquist, Christer MD, PhD[‡]; Kemeny, Andras MD^{**}; Kitz, Klaus MD^{‡‡}; Lippitz, Bodo MD^{§§}; Álvarez, Roberto Martínez MD, PhD[¶]; Pedersen, Paal-Henning MD, PhD^{¶¶}; Yomo, Shoji MD[§]; Lupidi, Francesco MD^{¶¶}; Dominikus, Karlheinz PhD^{***}; Blackburn, Philip MD^{‡‡‡}; Mindermann, Thomas MD^{§§§}; Bundschuh, Otto MD^{¶¶}; van Eck, A.T.C.J. MD^{¶¶¶}; Fimmers, Rolf PhD[‡]; Horstmann, Gerhard A. MD^{¶¶¶}

Santacrose et al. Long-term Tumor Control of Benign Intracranial Meningiomas After Radiosurgery in a Series of 4565 Patients 2011

Skullbase meningiomas irradiation

- Main therapy (**single treatment**)
- Post-operative (**residual**)
- Post-operative (**relapse**)

❖ **VMAT**

❖ **Radisurgery**

❖ **Particles Radiotherapy**



ISRS guidelines for Cavernous Sinus meningioma SRS



TABLE 4. Recommendations for Management of CS Meningioma

Evidence level

Level III	SRS/SRT is recommended as a primary/upfront treatment option for an asymptomatic, or mildly symptomatic CS meningioma. The recurrence rate is not appreciably different between primary or adjuvant therapy for a CS meningioma
Level III	Resection should be considered for the treatment of larger and symptomatic CS meningioma in patients both receptive to, and medically eligible, for open surgery
Level III	SRS/SRT delivered to a CS meningioma has a low risk of complications; most cranial nerve functions are preserved or improved due to tumor shrinkage. Carotid artery stenosis after SRS is rare.
Level III	When no residual tumor is observed, or only a small tumor lining on the dura of the CS exists postoperatively, serial neuroimaging studies is not unreasonable. At the time of recurrence or progression of residual tumor, SRS/SRT should be considered
Level III	In patients with a CS meningioma that has rapidly and substantially recurred after prior treatment, a subtotal surgical resection or biopsy may be considered. More aggressive features of the tumor (transformation of the tumor from WHO grade I to a higher grade) should be ruled out. These tumors have a predilection for progression and postoperative SRS/SRT with a higher dose should be strongly considered.
Level III	The technique for SRS or SRT delivery will depend upon the tumor histology, tumor volume and proximity of the tumor to adjacent critical structures (eg, the optic chiasm). SRS using single session marginal doses of 11 to 16 Gy offers a local tumor control rate of 90% or higher at 5 yr post-SRS.

ISRS guidelines

Management of cavernous sinus meningiomas: Consensus statement on behalf of the EANS skull base section (2022)

Stereotactic Radiosurgery SRS (single-dose or fractionated) should be considered in the following cases, insofar as the distance to the **ON is superior to 3 mm** (Level C):

- **Asymptomatic**, > 40 years old patients with a **purely intracavernous** CSMs <2.5 cm showing growth on serial imaging after initial conservative treatment;
- **Asymptomatic** patients with **partly extracavernous** CSMs showing growth on serial imaging after initial conservative treatment;
- **Symptomatic** patients with CSMs <2.5 cm, provided that the symptoms are **not related to ON compression**
- **Symptomatic** patients with partly extracavernous CSMs in whom **surgery is contraindicated**.

fractionated SRS or RT should be considered in cases that warrant treatment (see above) if the distance to the **ON is less than 3 mm** and the ipsilateral visual function is good (Level C).

Brain and Spine 2 (2022) 100864



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Management of cavernous sinus meningiomas: Consensus statement on behalf of the EANS skull base section



Marco V. Corniola^{a,b,c,d}, Pierre-Hugues Roche^e, Michaël Bruneau^f, Luigi M. Cavallo^g, Roy T. Daniel^h, Mahmoud Messerer^h, Sebastien Froelichⁱ, Paul A. Gardner^j, Fred Gentili^{k,†}, Takeshi Kawase^l, Dimitrios Paraskevopoulos^m, Jean Régisⁿ, Henry W.S. Schroeder^o, Theodore H. Schwartz^p, Marc Sindou^{q,r,s}, Jan F. Cornelius^l, Marcos Tatagiba^u, Torstein R. Meling^{d,v,*}

^a Department of Neurosurgery, Centre Hospitalier Universitaire de Rennes/Pontchaillou, Rennes, France

^b Faculty of Medicine, University of Rennes, Rennes, France

^c MedCIS Research Group, INSERM URI, UMR 1099 LTSI, France

^d Faculty of Medicine, University of Geneva, Geneva, Switzerland

^e Service de Neurochirurgie, Aix-Marseille Université, Assistance Publique-Hôpitaux de Marseille, Hôpital Nord, Marseille, France

^f Department of Neurosurgery, Universitair Ziekenhuis Brussel, Vrije Universiteit Brussel, Laarbeeklaan, Brussels, Belgium

^g Department of Neurological Sciences, Division of Neurosurgery, Università Degli Studi di Napoli Federico II, Naples, Italy

^h Department of Neurosurgery, Centre Hospitalier Universitaire Vaudois, Lausanne, Switzerland

ⁱ Service de Neurochirurgie, Hôpital Lariboisière, Assistance Publique-Hôpitaux de Paris, APHP Nord, Paris, France

^j Department of Neurological Surgery, University of Pittsburgh, Pittsburgh, PA, USA

^k Department of Neurosurgery, Toronto Western Hospital, University Health Network, Toronto, Ontario, Canada

^l Department of Neurosurgery, School of Medicine, Keio University, Tokyo, Japan

^m Department of Neurosurgery, Barts Health NHS Trust, St. Bartholomew's and the Royal London Hospital, London, United Kingdom

ⁿ Department of Functional and Stereotactic Neurosurgery and Radiosurgery, Timone University Hospital, Marseille, France

^o Department of Neurosurgery, Ernst Moritz Arndt University, Greifswald, Germany

^p Department of Neurosurgery, Weill Cornell Medicine, New York Presbyterian Hospital, New York, USA

^q University of Lyon, France

^r IRR Flavigny, UGECAM Nord-Est, Nancy, France

^s Pediatric Neurosurgery Department, Centre Hospitalier Régional de Nancy, Nancy, France

^t Department of Neurosurgery, Heinrich Heine Universität, Düsseldorf, Germany

^u Department of Neurosurgery, Uniklinik Tübingen, Tübingen, Germany

<https://www.ncbi.nlm.nih.gov/pmc/articles/>

EANS guidelines

Management of cavernous sinus meningiomas: Consensus statement on behalf of the EANS skull base section

Marco V. Corniola^{a,b,c,d}, Pierre-Hugues Roche^e, Michaël Bruneau^f, Luigi M. Cavallo^g, Roy T. Daniel^h, Mahmoud Messererⁱ, Sebastian Froelich^j, Paul A. Gardner^k, Fred Gentili^{l,m}, Takeshi Kawaseⁿ, Dimitrios Paraskevopoulos^o, Jean Régis^p, Henry W.S. Schroeder^q, Theodore H. Schwartz^r, Marc Sindou^{s,t,u}, Jan F. Cornelius^v, Marcos Tatagiba^w, Torstein R. Meling^{x,y,z}

^a Department of Neurosurgery, Centre Hospitalier Universitaire de Rennes/Spinechirurgie, Rennes, France

^b Faculty of Medicine, University of Rennes, Rennes, France

^c Medical Research Group, INSERM UMR, UMR 1099 I2U, France

^d Faculty of Medicine, University of Geneva, Geneva, Switzerland

^e Service de Neurochirurgie, Aix Marseille Université, Assistance Publique Hôpital de Marseille, Hôpital Nord, Marseille, France

^f Department of Neurosurgery, Universitair Ziekenhuis Brussel, Vrije Universiteit Brussel, Laarbeeklaan, Brussels, Belgium

^g Department of Neurological Sciences, Division of Neurosurgery, Università Degli Studi di Napoli Federico II, Naples, Italy

^h Department of Neurosurgery, Centre Hospitalier Universitaire Vaudois, Lausanne, Switzerland

ⁱ Service de Neurochirurgie, Hôpital Lariboisière, Assistance Publique Hôpitaux de Paris, APHP Nord, Paris, France

^j Department of Neurological Surgery, University of Pittsburgh, Pittsburgh, PA, USA

^k Department of Neurosurgery, Toronto Western Hospital, University Health Network, Toronto, Ontario, Canada

^l Department of Neurosurgery, School of Medicine, Keio University, Tokyo, Japan

^m Department of Neurosurgery, Barns Health NHS Trust, St. Barnabas's and the Royal London Hospital, London, United Kingdom

ⁿ Department of Functional and Stereotactic Neurosurgery and Radiosurgery, Thomsen University Hospital, Marseille, France

^o Department of Neurosurgery, Ernst Moritz Arndt University, Greifswald, Germany

^p Department of Neurosurgery, Weill Cornell Medicine, New York Presbyterian Hospital, New York, USA

^q University of Ljubljana, Ljubljana, Slovenia

^r IIR Hologry, UGICAM Nord Est, Nancy, France

^s Pediatric Neurosurgery Department, Centre Hospitalier Régional de Nancy, Nancy, France

^t Department of Neurosurgery, Heinrich Heine Universität, Düsseldorf, Germany

^u Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

^v Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

^w Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

^x Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

^y Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

^z Department of Neurosurgery, Friedrich-Wilhelms-Universität, Bonn, Germany

Conclusions

- **SRS** should be advocated as first line treatment in **small/asymptomatic** lesions/in **elderly** patients.
- Offer **excellent tumour control** with **low rates of oculomotor/visual complications**.
- **Endoscopic Endonasal Approach (EEA)**, a safe strategy of **bony skull base decompression** and **limited tumour removal** in the **exophytic component** of the tumour, outside the cavernous sinus. **Combined with SRS** can be for **symptoms relief** and **tumour control**

Radiosurgery for Skullbase meningioma

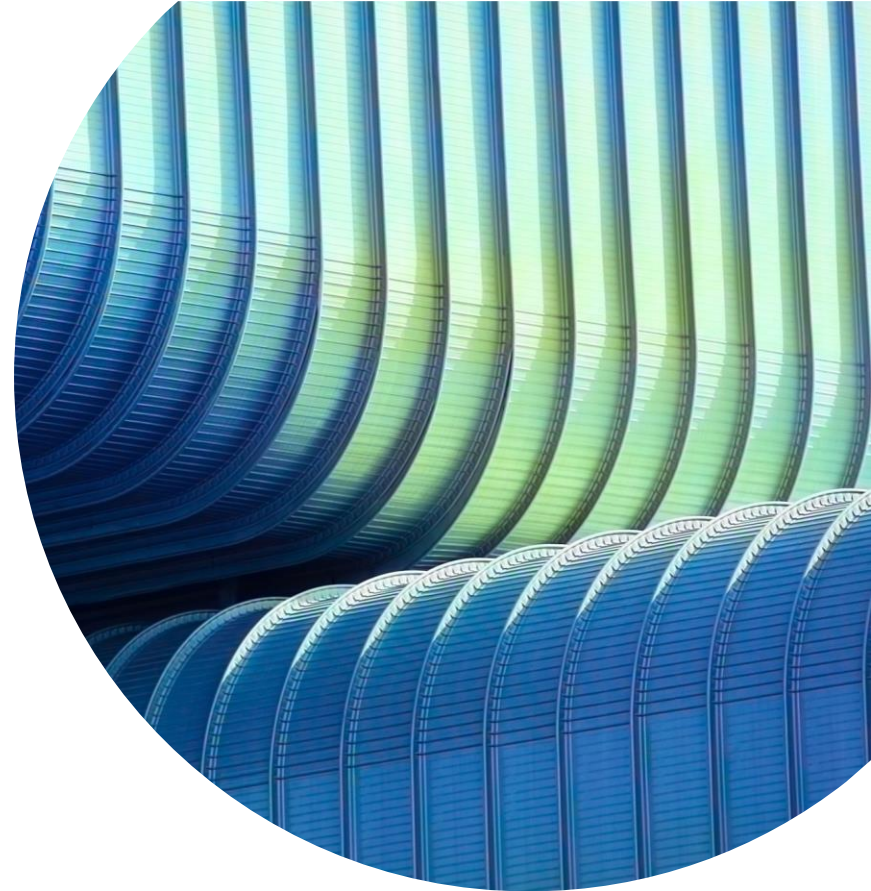


Radiosurgery - Definition

High dose in 1 fraction

High accuracy <1mm

High gradient of dose (rapid fall off dose) →
normal tissue preservation





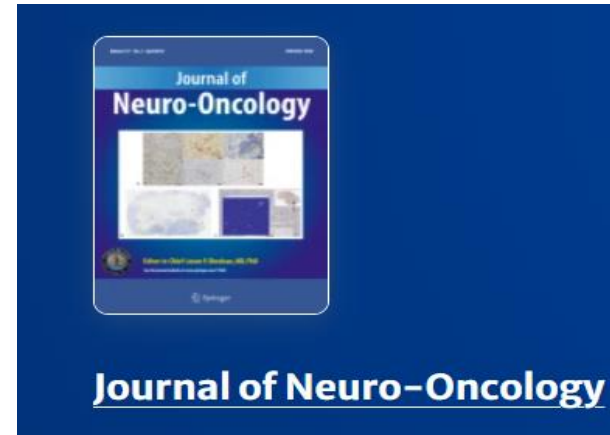
SRS Radiobiology

RADIOSURGERY:

- creates more **double-strand breaks** in DNA,
- results in **less DNA damage repair**,
- has **anti-vascular effects**,
- Has ***in situ* vaccine effects** and **abscopal effect**
(immuno-stimulation and immuno-upregulation)

Factors for SRS treatment decision

- Size
- Location
- Rate of growth (aggressiveness)
- Age
- Performance Status
- Patients goals for treatment (choice)

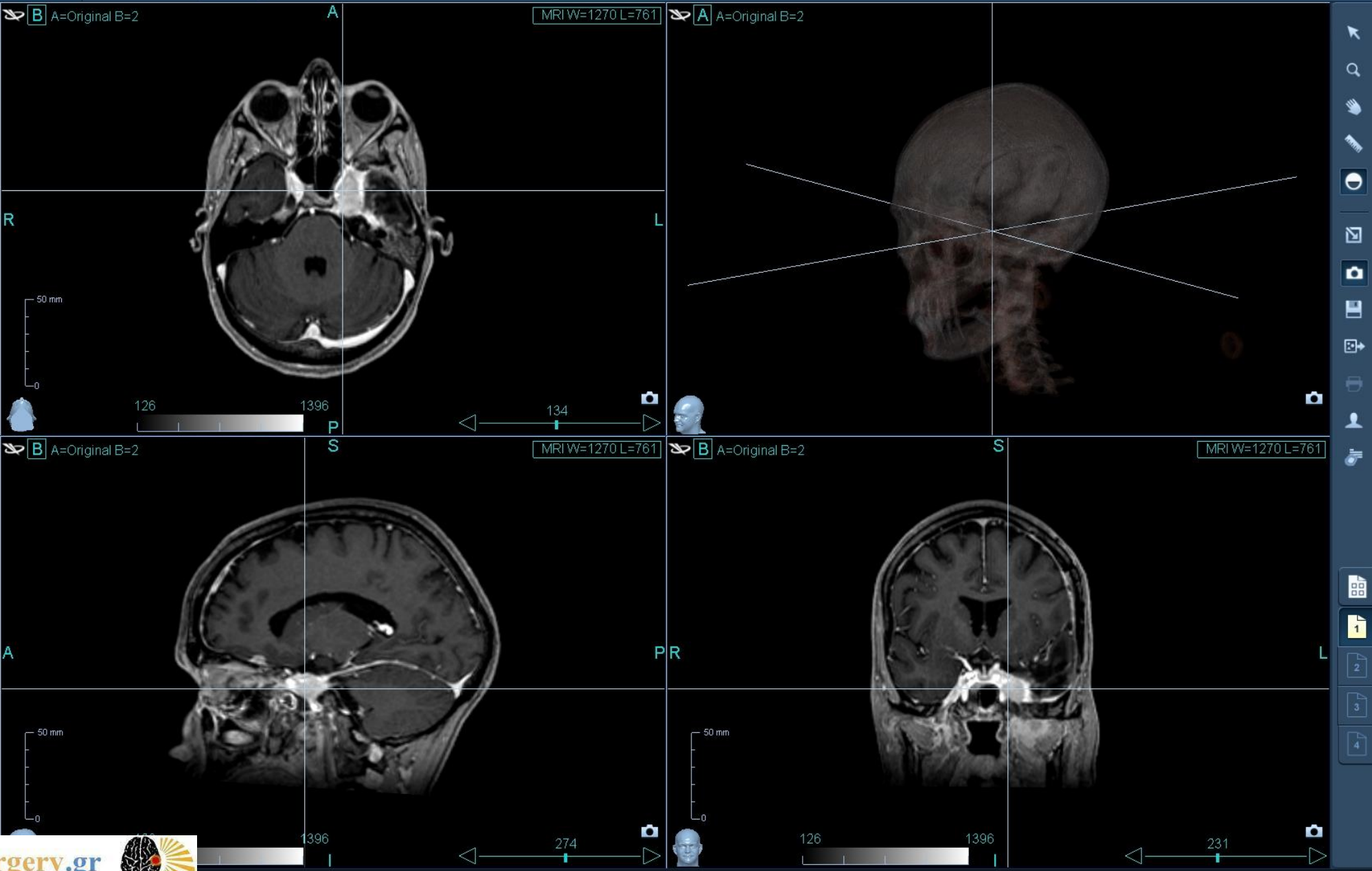


Alfredo Quinones Hinojosa
J Neurooncology 2020

Tools VOIs Display

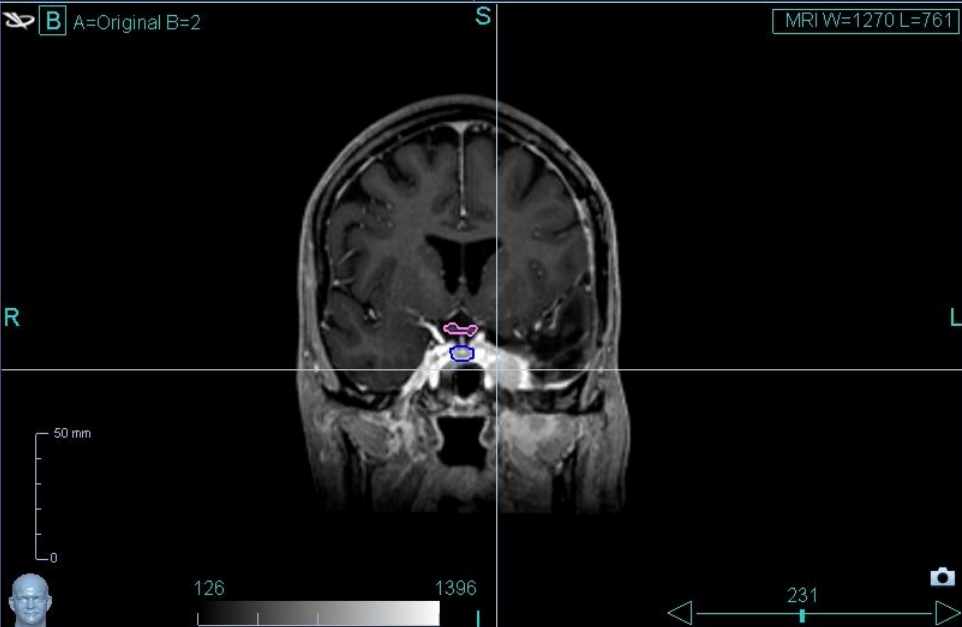
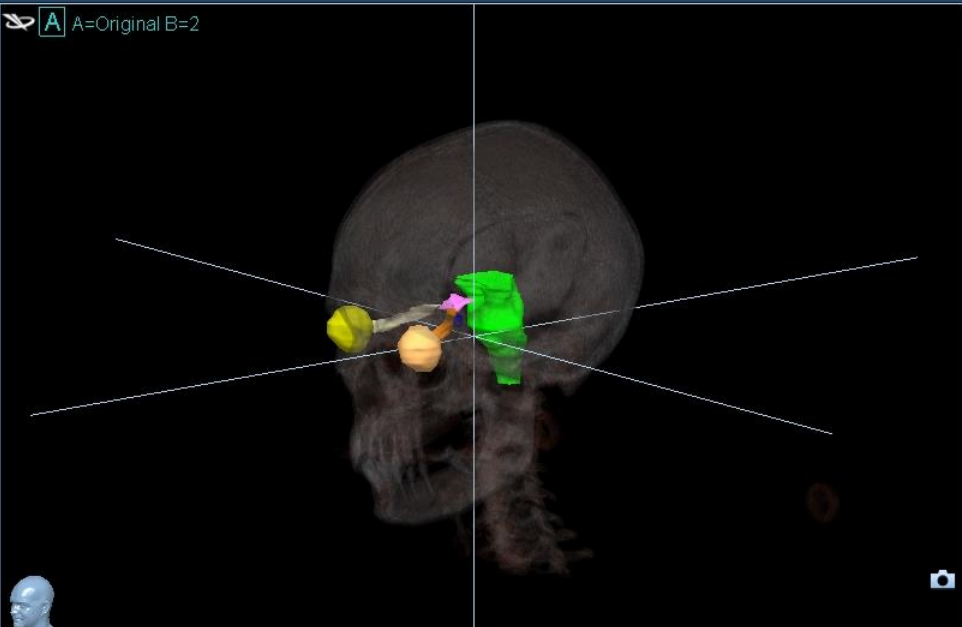
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<input type="checkbox"/>	Right Eye
<input type="checkbox"/>	Right Optic Nerve
<input type="checkbox"/>	Right Lens
<input type="checkbox"/>	Left Eye
<input type="checkbox"/>	Left Optic Nerve
<input type="checkbox"/>	Left Lens
<input type="checkbox"/>	Brainstem
<input type="checkbox"/>	Pituitary Gland

Display VOI as overlay



Tools VOIs Display

	Name
<input type="checkbox"/>	GTV
<input checked="" type="checkbox"/>	Optic Chiasm
<input checked="" type="checkbox"/>	Right Eye
<input checked="" type="checkbox"/>	Right Optic Nerve
<input checked="" type="checkbox"/>	Right Lens
<input checked="" type="checkbox"/>	Left Eye
<input checked="" type="checkbox"/>	Left Optic Nerve
<input checked="" type="checkbox"/>	Left Lens
<input checked="" type="checkbox"/>	Brainstem
<input checked="" type="checkbox"/>	Pituitary Gland



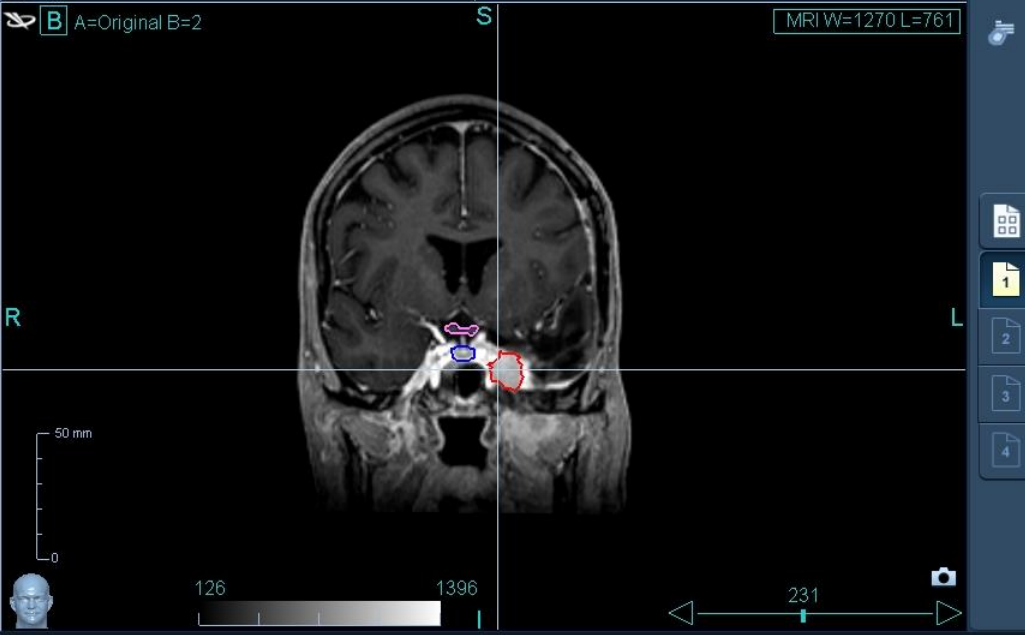
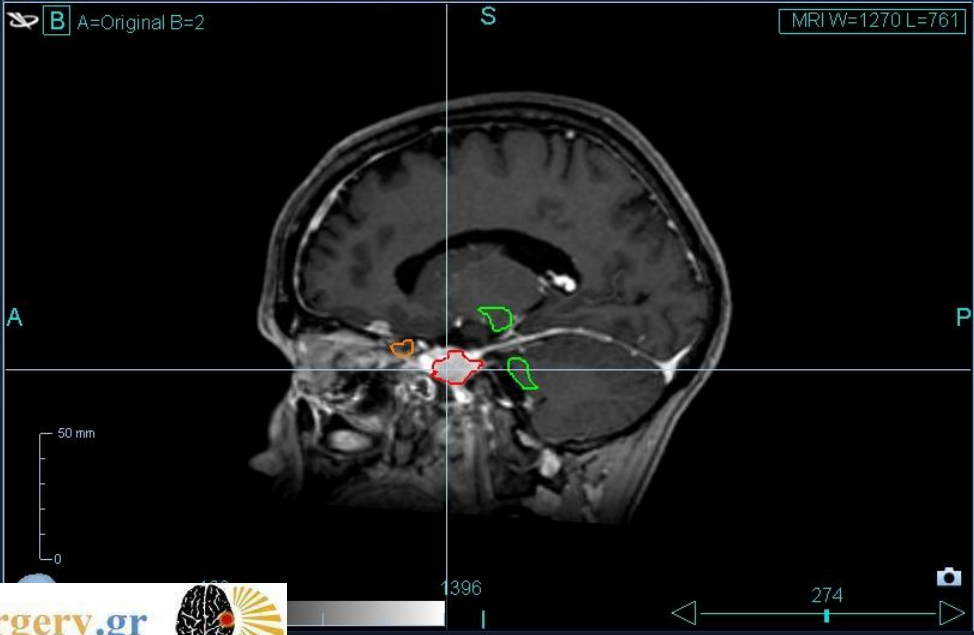
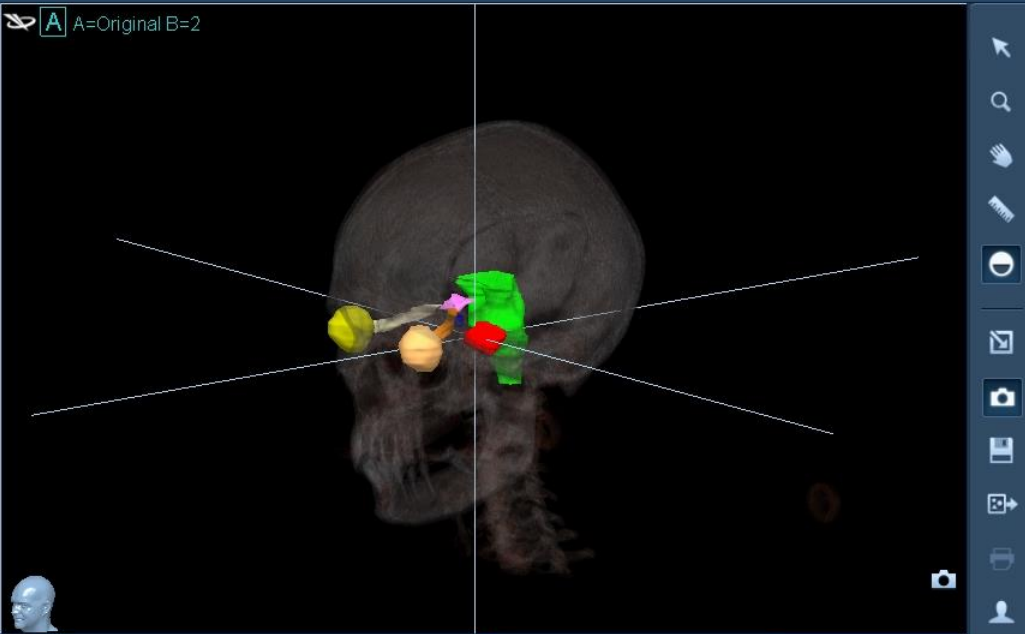
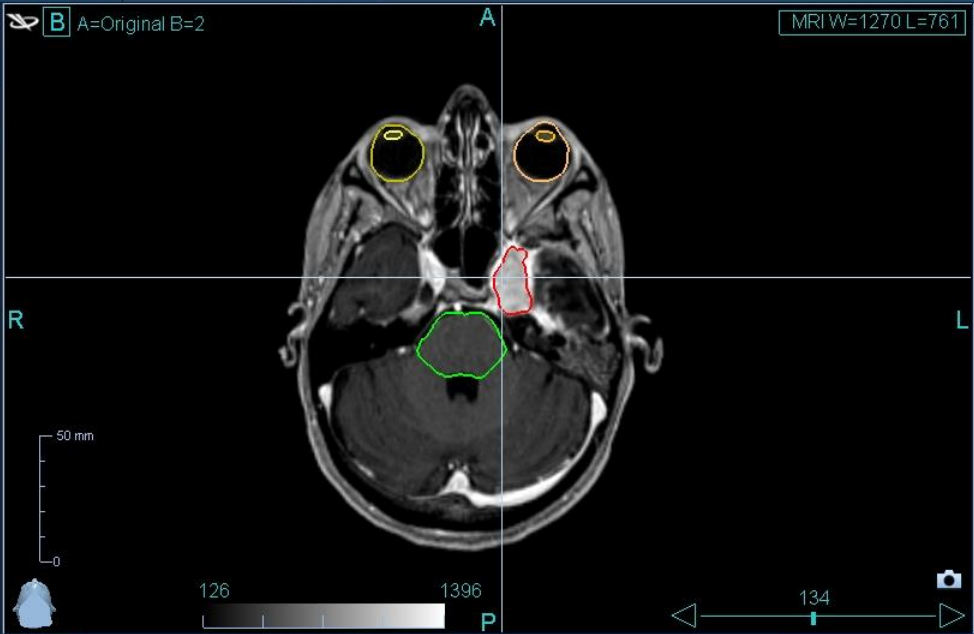
Display VOI as overlay



Tools VOIs Display

<input type="checkbox"/>	Name
<input checked="" type="checkbox"/>	T GTV
<input checked="" type="checkbox"/>	C Optic Chiasm
<input checked="" type="checkbox"/>	C Right Eye
<input checked="" type="checkbox"/>	C Right Optic Nerve
<input checked="" type="checkbox"/>	C Right Lens
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<input checked="" type="checkbox"/>	C Left Optic Nerve
<input checked="" type="checkbox"/>	C Left Lens
<input checked="" type="checkbox"/>	C Brainstem
<input checked="" type="checkbox"/>	C Pituitary Gland

Display VOI as overlay



Navigation and tool icons including zoom, pan, and window management.

Prescription Dose

Single fraction: 12-13Gy

Dose (crute) (Local Control)
Dose conformality (Toxicity)
Dose fall off (Toxicity)

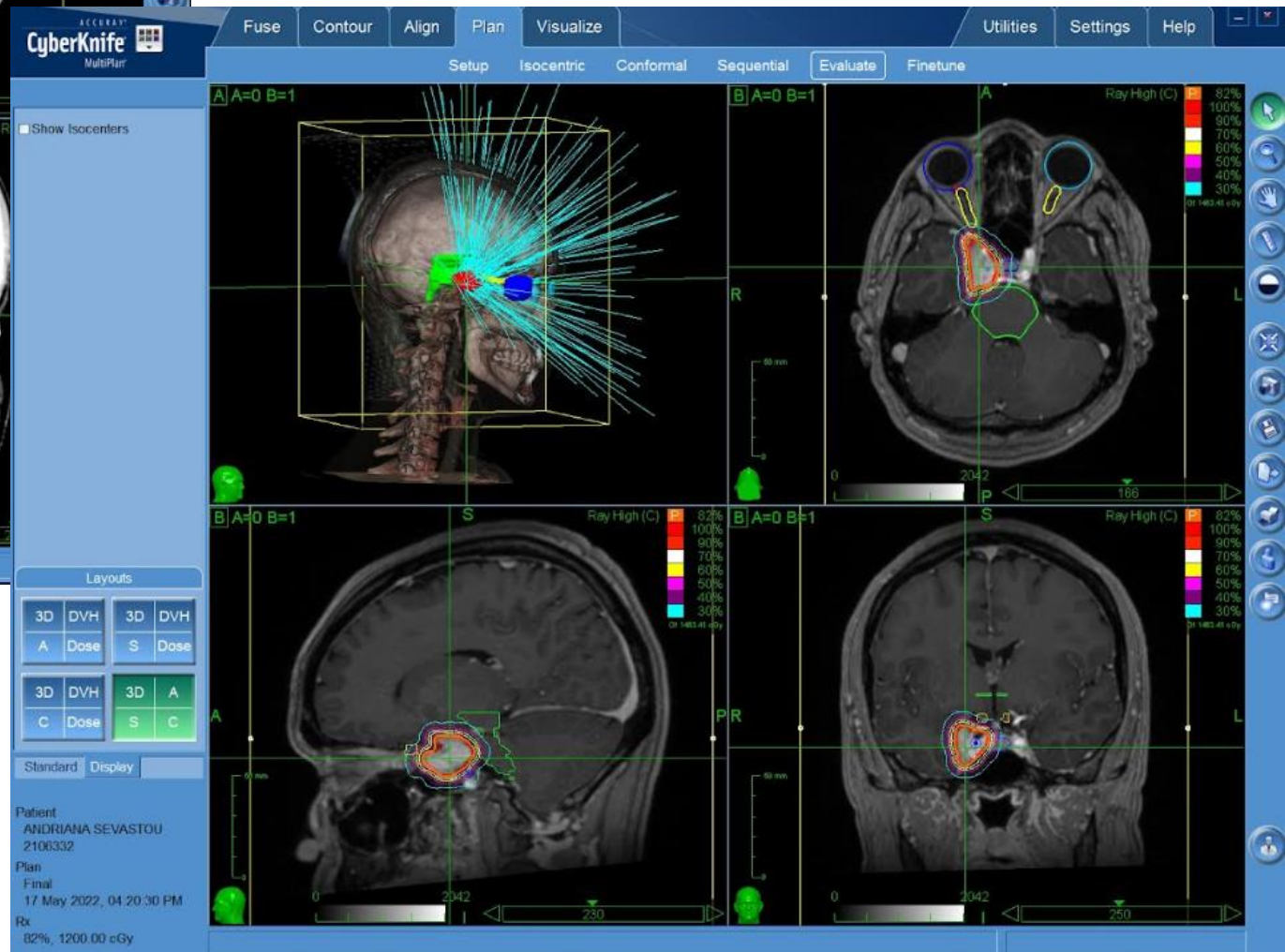
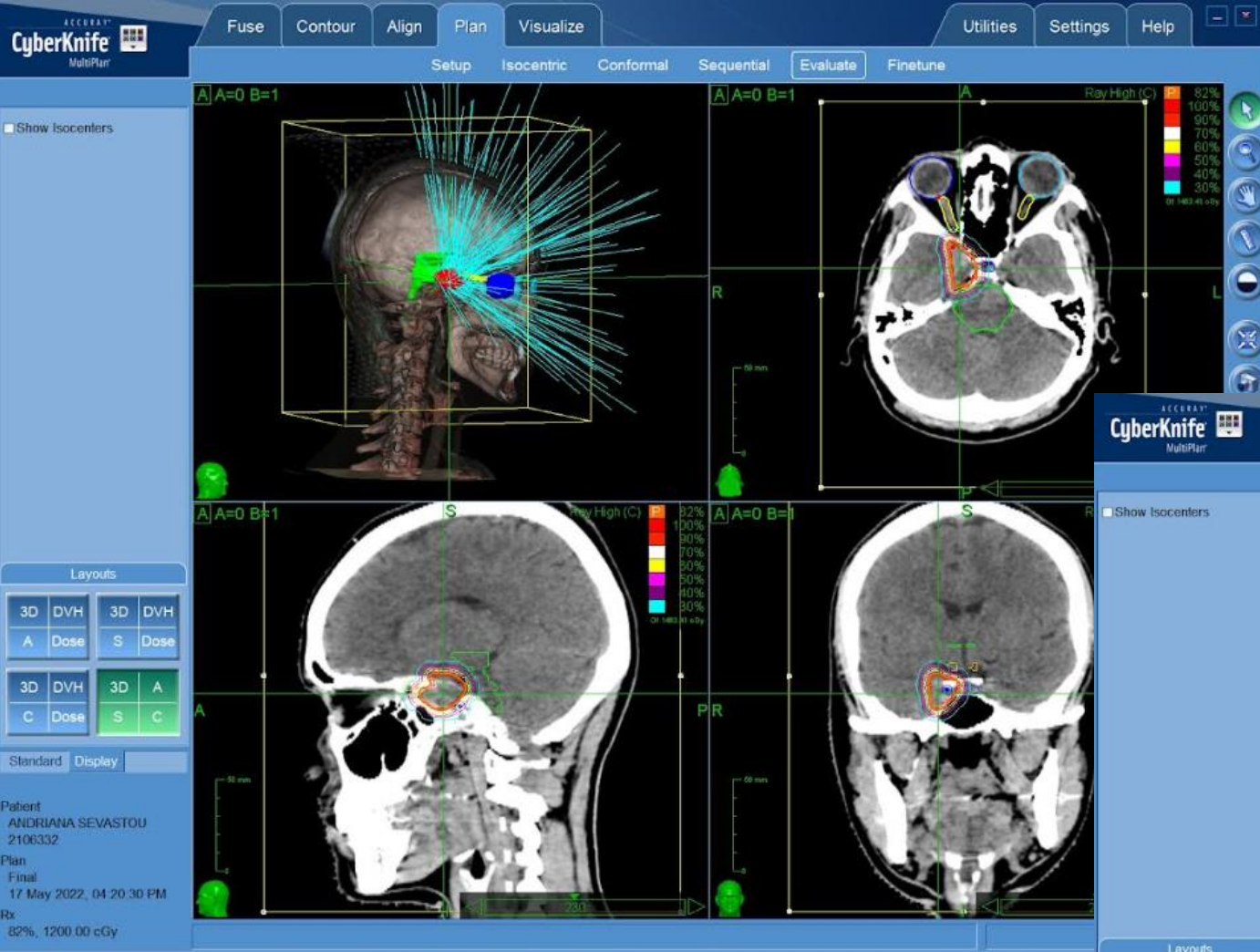
Dose and Local Control

11Gy possible working

12Gy – 13Gy standard working

14Gy+ working but...





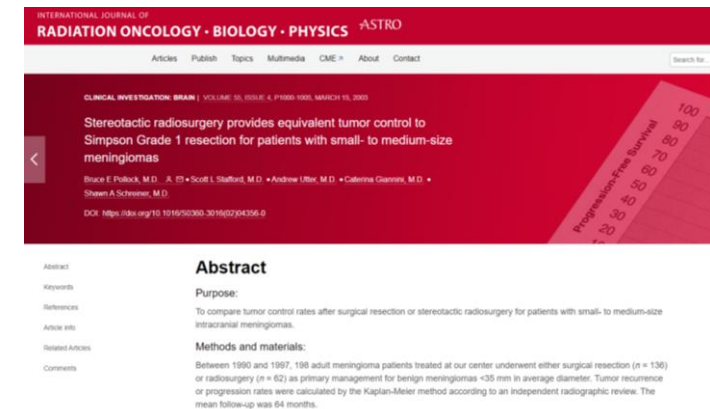
Radiosurgery outcomes



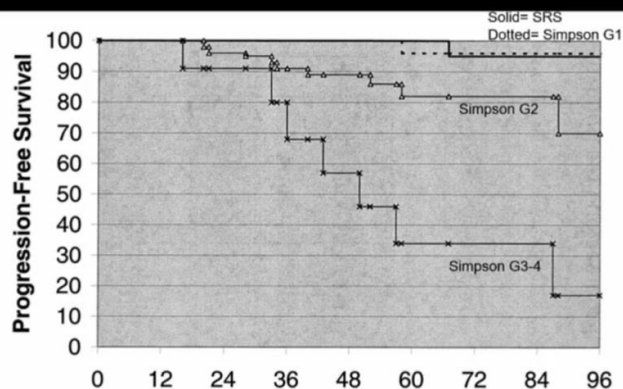
SRS equal long-term control to Simpsons Grade 1

No statistically significant difference was detected in the 3- and 7-year actuarial progression-free survival (PFS) rate between patients with Simpson Grade 1 resections (100% and 96%) and patients who underwent radiosurgery (100% and 95%, $p = 0.94$).

198 patients , surgical resection (n = 136) or radiosurgery (n = 62)
as primary management for benign meningioma



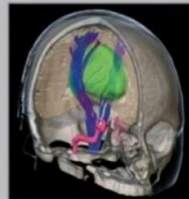
SRS PROVIDES EQUAL LONG-TERM CONTROL TO SIMPSON GRADE 1



Controversies in Neuro-Oncology

Best Evidence Medicine for Brain Tumor Surgery

Alfredo Quilones-Hinojosa
Shaan M. Raza



Chapter 28: The role of SRS in newly diagnosed meningiomas

<https://pubmed.ncbi.nlm.nih.gov/121570360/>

Pollock IJROBP 2003



ISRS Guidelines for Cavernous Sinus Meningiomas

REVIEW

Cheng-Chia Lee, MD, PhD^{*555}
Daniel M. Trifiletti, MD[†]
Arjun Sahgal, MD[‡]
Antonio DeSalles, MD[§]
Laura Fariselli, MD^{||}
Motohiro Hayashi, MD[¶]
Marc Levivier, MD^{**}
Lijun Ma, PhD^{††}
Roberto Martínez Álvarez, MD^{§§}
Ian Paddick, PhD^{¶¶}
Jean Regis, MD^{|||}
Samuel Ryu, MD^{##}
Ben Slotman, MD, PhD^{***}
Jason Sheehan, MD, PhD^{†††}

Stereotactic Radiosurgery for Benign (World Health Organization Grade I) Cavernous Sinus Meningiomas—International Stereotactic Radiosurgery Society (ISRS) Practice Guideline: A Systematic Review

- 5-yr PFS 86% to 99%
- 10-yr PFS 69% to 97%
- 15-yr PFS rates 92%
- 20-yr PFS rates ranging from 87%

5- yr **Local Control** rate of 99%
10-yr **Local Control** rate of 93%



ISRS guidelines **Cavernous Sinus Meningioma**

Skull base meningioma Local Control

	3years	5years	10 years	20years
Cavernus Sinus	94	91	85	72
Petroclival	97	95	94	86
C P Angle	95	90	86	81
Clinoid	95	93	88	82
Sphenoid Wing	96	96	90	90
Olfactory Groove	93	88	83	78
Tentorial	96	94	87	84

Clinical follow up SRS treated Skullbase Meningioma

Neurological improvement

- CS 44.2%
- PC 34%
- CPA 38.5%

Complete symptoms response

CS 23.2%
PC 15%
CPA 11.5%



CS:Cavernous sinus

CPA:CerebroPontal Angle

PC:Petro Clival

Santacrose et al. Long-term Tumor Control of Benign Intracranial Meningiomas After Radiosurgery in a Series of 4565 Patients 2011

Cranial nerve outcomes in patients who underwent SRS for CS meningiomas

with or without prior microsurgery

Improvement rates specific to Cranial Nerve Deficits after SRS :

- 20% at 1 yr,
- 34% at 2 yr,
- 36% at 3 yr, and
- 39% at 5 yr.

Fractionated SRS



Fractionated SRS for Skullbase Meningioma

- Large Tumor **Volume**
- Proximity to **Optic Pathway**

- Lesion >30mm
- Lesion >15cc
- Lesion <2-3mm optic nerve distance

- Local Control ?
- Toxicity ?

Multi session SRS for meningiomas

Phase II Prospective study
Dose: **25Gy** in 5 fractions
N=178

5-year tumor control: 97%
Overall Toxicity:12,7%

RADIATION ONCOLOGY • BIOLOGY • PHYSICS ASTRO

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CLINICAL INVESTIGATION | VOLUME 115, ISSUE 1, P153-163, JANUARY 01, 2023

Hypofractionated Radiosurgery for Large or in Critical-Site Intracranial Meningioma: Results of a Phase 2 Prospective Study

Valentina Pinzi, MD [✉](#) • [Marcello Marchetti, MD](#) • [Anna Viola, MD](#) • ... [Irene Cane, BSc](#) • [Cecilia Iezzi, PsyD](#) • [Laura Fariselli, MD](#) • [Show all authors](#)

[Open Access](#) • Published: September 05, 2022 • DOI: <https://doi.org/10.1016/j.ijrobp.2022.08.064>

Image Guided Multisession Radiosurgery of Skullbase meningiomas

Retrospective Analysis
Dose: **25Gy** in 5 fractions
N=156

5-year tumor control: 90%

- **Progression-free survival at 2-, 5-, and 10- years was 95%, 90%, and 80.8%, respectively.**
- **There were no new visual or motor deficits, nor cranial nerves impairments, excluding trigeminal neuralgia, which was reported by 5.7% of patients.**



Article

Image-Guided Multisession Radiosurgery of Skull Base Meningiomas

Alfredo Conti ^{1,2,*}, Antonio Pontoriero ³, Giuseppe Iati ³, Salvatore M. Cardali ⁴, Anna Brogna ³, Filippo Friso ², Vittoria Rosetti ², Matteo Zoli ^{1,2}, Silvana Parisi ³, Alberto Cacciola ³, Sara Lillo ³, Stefano Pergolizzi ³ and Diego Mazzatenta ^{1,2}

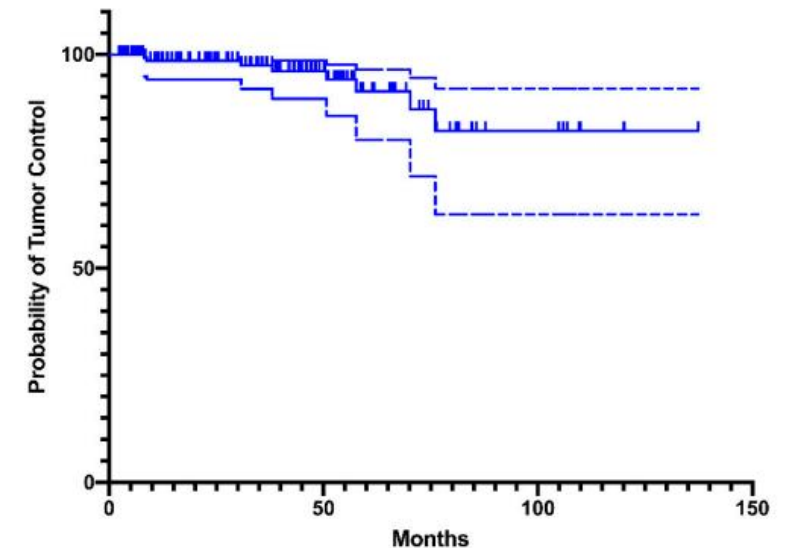


Figure 4. Progression-free survival (PFS) ± 95% confidence interval.



Radiobiological advantage in fractionation?

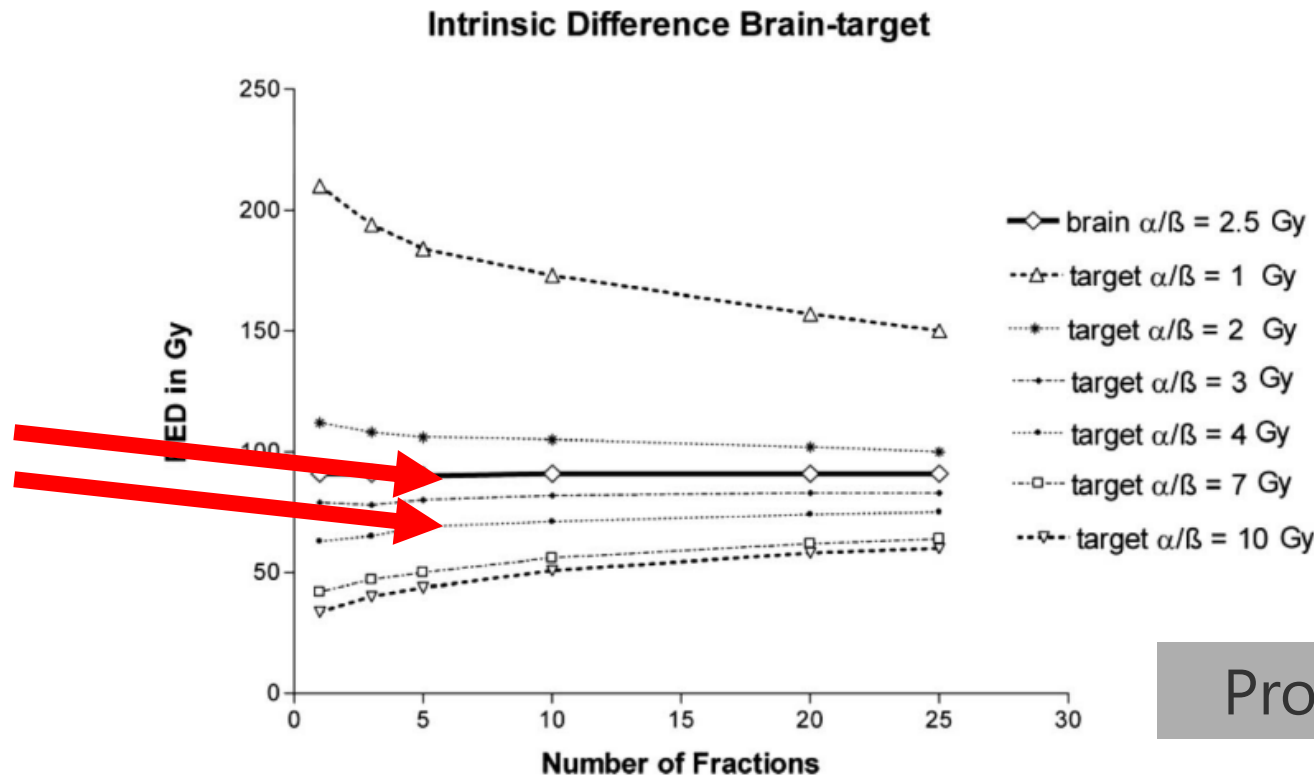
Assessment of the α/β ratios for meningiomas

Assessment of the α/β ratios for arteriovenous malformations, meningiomas, acoustic neuromas, and the optic chiasma

<https://www.tandfonline.com/doi/full/10.3109/09553001003667982>

Assessment of the α/β ratios for arteriovenous malformations, meningiomas, acoustic neuromas, and the optic chiasma

Frederik J. A. I. Vernimmen & Jacobus P. Slabbert
Pages 486-498 | Received 23 Jan 2009, Accepted 29 Jan 2010, Published online: 15 May 2010
Cite this article | <https://doi.org/10.3109/09553001003667982>



Brain normal tissue alpha/beta 2.5
meningiomas alpha/beta 3.3 – 3.7

Probably **not so much** as we think

Figure 6. BED variations for a number of target α/β values in relation to the brain. The brain maintains a constant effect (constant BED value) across a number of isoeffective schedules of increasing number of fractions. While the physical dose to the target and the brain being the same for each fractionation schedule, a changing BED for the target can be seen across the spectrum of fractionations.

SRS vs. SRT CS Meningioma: tumor shrinkage ???



CLINICAL STUDIES: RADIOSURGERY: RADIOTHERAPY: TUMOR

Evaluation of Fractionated Radiotherapy and Gamma Knife Radiosurgery in Cavernous Sinus Meningiomas: Treatment Strategy

Metellus, Philippe M.D.; Regis, Jean M.D.; Muracciole, Xavier M.D.; Fuentes, Stephane M.D.; Dufour, Henry M.D.; Nanni, Isabelle M.D.; Chinot, Oliver M.D.; Martin, Pierre-Marie M.D., Ph.D.; Grisoli, Francois M.D.

Author Information

Neurosurgery 57(5):p 873-886, November 2005. | DOI: 10.1227/01.NEU.0000179924.76551.cd

However, radiologically 29% of patients who underwent SRT, and 53% of patients who underwent **SRS**, showed **tumor shrinkage (P < .04)**

The result implied that **SRS** offered a **higher rate of tumor shrinkage**, but no significance in clinical improvement.

Metellus P, Regis J, Muracciole X, et al. Evaluation of fractionated radiotherapy and gamma knife radiosurgery in cavernous sinus meningiomas: treatment strategy. Neurosurgery. 2005

High Grade skullbase meningiomas



Grade II and Grade III meningioma criteria

Pathology grading

Grade II

- Brain invasion
- Few or more mitoses (<20 mitoses)
- Three of the following:
 - Increased cellularity
 - Prominent nucleoli
 - Loss of architecture
 - Necrosis
 - Small cell change

Grade III

- Overt anaplasia
- >20 mitoses



Grade II: immediate or waiting RT after GTR ?

- EORTC 22042-26042

Radiotherapy & Oncology

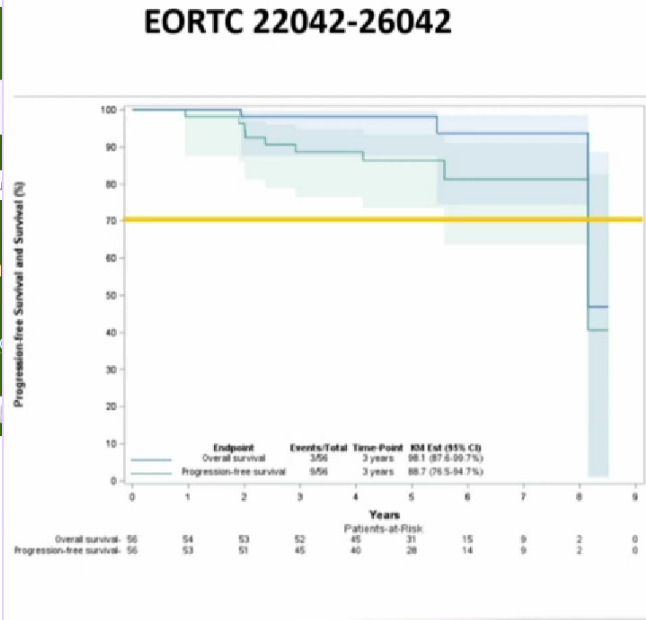
Articles Publish Topics About Contact

PHASE II TRIAL | VOLUME 128, ISSUE 2, P260-265, AUGUST 2018 [Download Full Text](#)

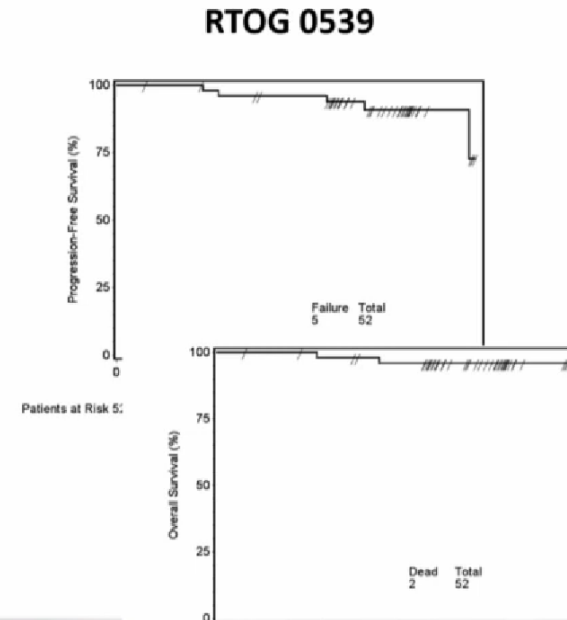
Adjuvant postoperative high-dose radiotherapy for malignant meningioma: A phase-II parallel non-randomized controlled trial versus observation study (EORTC 22042-26042)

Damien C. Weber • Carmen Ares • Salvador Villa • ... Yan Liu • Laurence ...
[Show all authors](#)

Published: June 27, 2018 • DOI: <https://doi.org/10.1016/j.radonc.2018.06.018>



- RTOG 0539



BIOLOGY • PHYSICS

Articles Topics Multimedia CME About Contact

VOLUME 128, ISSUE 4, P790-799, MARCH 15, 2020

Initial Outcomes From NRG 0539

Michael A. Vogelbaum, MD, PhD • ...
 M. M. Amos, MD • Michael A. Vogelbaum, MD, PhD • ...
 PhD • Mitesh P. Mehta, MD • [Show all authors](#)

DOI: <https://doi.org/10.1016/j.ijrobp.2019.11.028> [Check for updates](#)

Excellent results in Favor of post-operative RT

EORTC 22042-26042 and RTOG 0539

Using **modern RT technics** (like IMRT)

- we can deliver **higher dose** in the target, protecting the normal brain
- very low rate of **high grade late toxicity**

paradox

SRS :

- **higher conformity** in the target
- **increased dose gradient** outside the target (rapid fall off the dose)

Adjuvant SRS improves Post-surgical long term outcomes (regardless of the extent of resection)

7486 patients,
6788 with atypical meningiomas
698 with malignant meningiomas

[Home](#) > [Journal of Neuro-Oncology](#) > [Article](#)

Adjuvant radiation for WHO grade II and III intracranial meningiomas: insights on survival and practice patterns from a National Cancer Registry

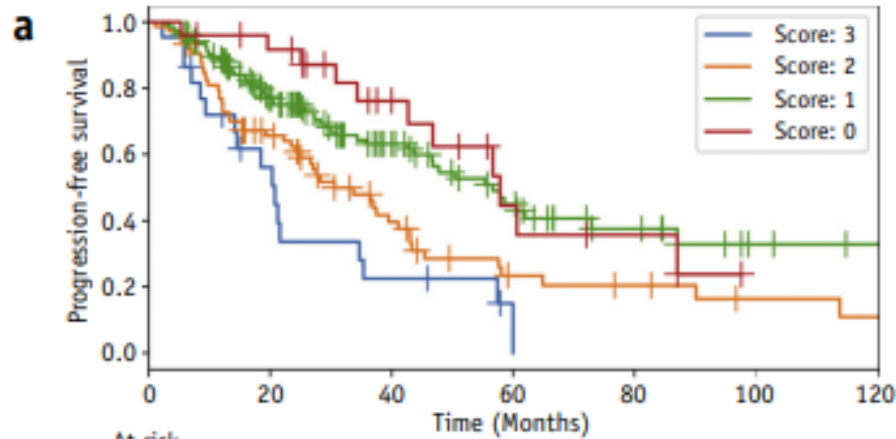
Clinical Study | Published: 28 August 2020

Volume 149, pages 293–303, (2020) [Cite this article](#)

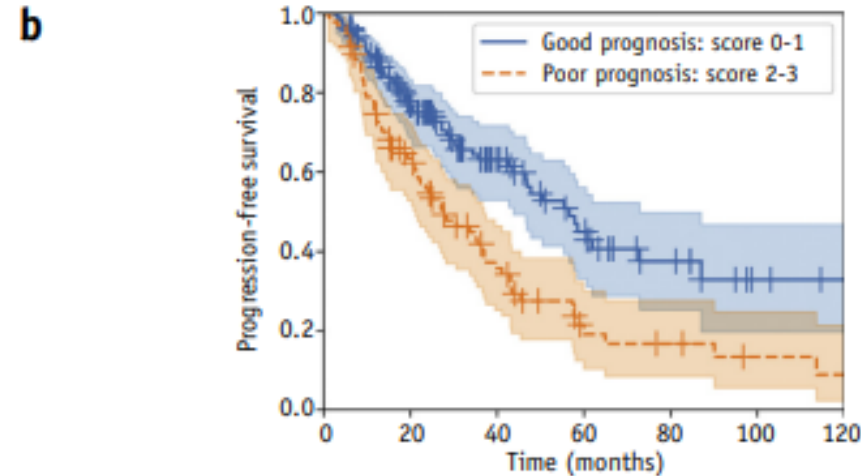
- Overall **5-year survival** was **76.9%** and **43.3%** among patients with WHO grades **II and III** meningiomas, respectively.
- **Adjuvant RT** correlated with **improved survival** in a multivariable model in patients with **grade II** tumors ($p = 0.029$)

Selection of pts Grade II meningioma for SRS (adjuvant or definitive)

Stratifications of the cohort



At risk	0	20	40	60	80	100	120
Score: 3	22	10	4	1	0	0	0
Score: 2	74	41	19	8	6	3	2
Score: 1	137	82	42	23	11	3	1
Score: 0	25	21	12	5	3	0	0



At risk	0	20	40	60	80	100	120
Good prognosis: score 0-1	137	82	42	23	11	3	1
Poor prognosis: score 2-3	96	51	23	9	6	3	2

SRS is a good choice for patients age<50, up to 1 previous resection, no previous RT (+ Volume<11.5cc)

Clinical Investigation

Treatment of WHO Grade 2 Meningiomas With Stereotactic Radiosurgery: Identification of an Optimal Group for SRS Using RPA

Roman O. Kowalchuk, MD,* Matthew J. Shepard, MD,[†]
Kimball Sheehan,[†] Darrah Sheehan,[†] Andrew Faramand, MD, MSc,[‡]
Ajay Niranjana, MD, MBA,[‡] Hideyuki Kano, MD, PhD,[§]
Jason Gurewitz, BA,[§] Kenneth Bernstein, MS,[¶] Roman Liscak, MD,[¶]
Khumar Guseynova, MD,[¶] Inga S. Grills, MD,[¶] Jacob S. Parzen, MD,[¶]
Christopher P. Cifarelli, MD, PhD,** Azeem A. Rehman, MD,**
Ahmet Atik, MD,^{††} Joshua Bakhsheshian, MD,^{††} Gabriel Zada, MD,^{††}
Eric Chang, MD,^{§§} Steven Giannotta, MD,^{‡‡} Herwin Speckter, MSc,^{‡‡}
Hsiu-mei Wu, MD,^{¶¶} Douglas Kondziolka, MD,[§]
David Mathieu, MD,** Cheng-chia Lee, MD, PhD,^{##,†††}
Ronald E. Warnick, MD,^{†††} L. Dade Lunsford, MD,[‡]
Daniel M. Trifiletti, MD,^{§§§} and Jason P. Sheehan, MD, PhD[†]



Factors affecting SRS outcomes



factors associated with improved SRS local control

Statistically significant ($P < .05$) factors associated with improved SRS local control outcomes were:

- higher marginal **dose**,
- small- to medium-**sized** tumors,
- WHO **grade I**,
- **upfront SRS** (irradiated tumor without surgical resection),
- **early SRS** (cranial deficits < 1 yr),
- **female sex**,
- younger **age**,
- **less conformal** plans



ISRS guidelines

Multivariate analysis of **factors** associated with **improved local control** after **SRS** (stable and reduced volume)

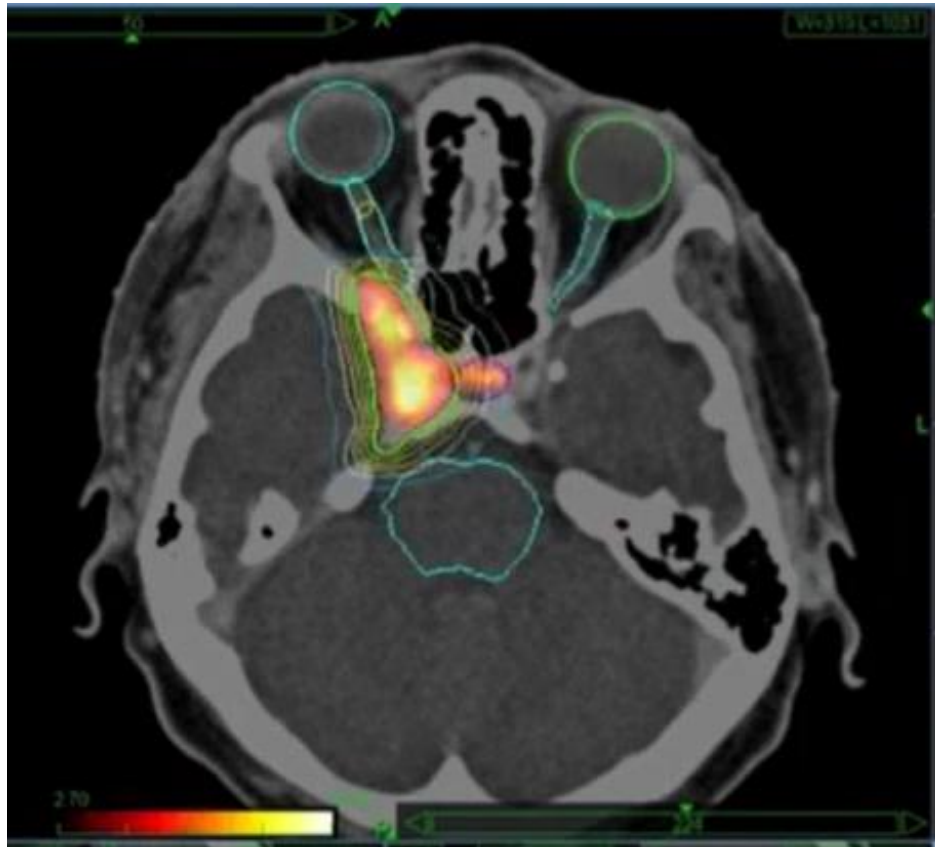
- Center experience (**CS**)
- Female (**CS,CP**)
- meningiomatosis vs sporadic (**CS, PC**)
- Prescription dose (**PC**)
- Maximum dose (**CP**)

- Cavernous sinus
- CerebroPontal Angle
- Petro clival



Santacrose et al. Long-term Tumor Control of Benign Intracranial Meningiomas After Radiosurgery in a Series of 4565 Patients **2011**

IMAGING: impact of ^{68}Ga -DOTATOC PET to SRS on target volume delineation of meningiomas



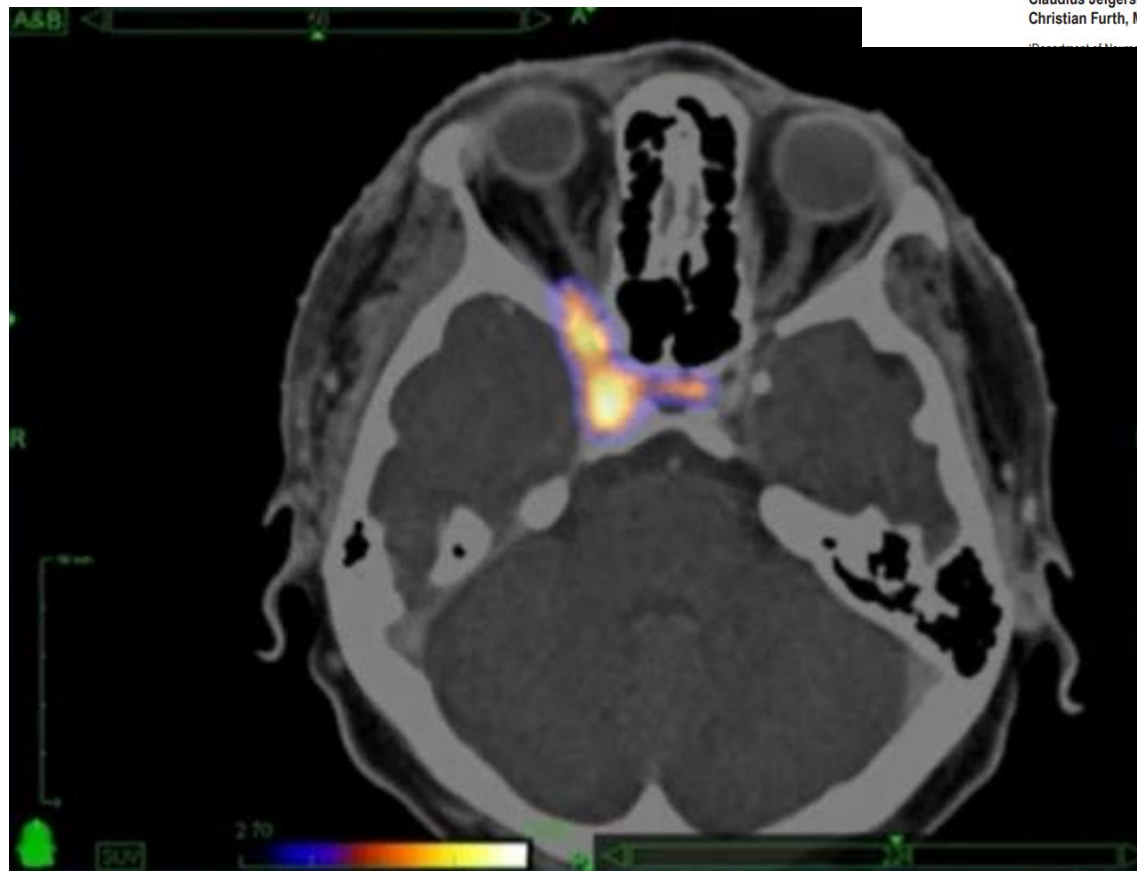
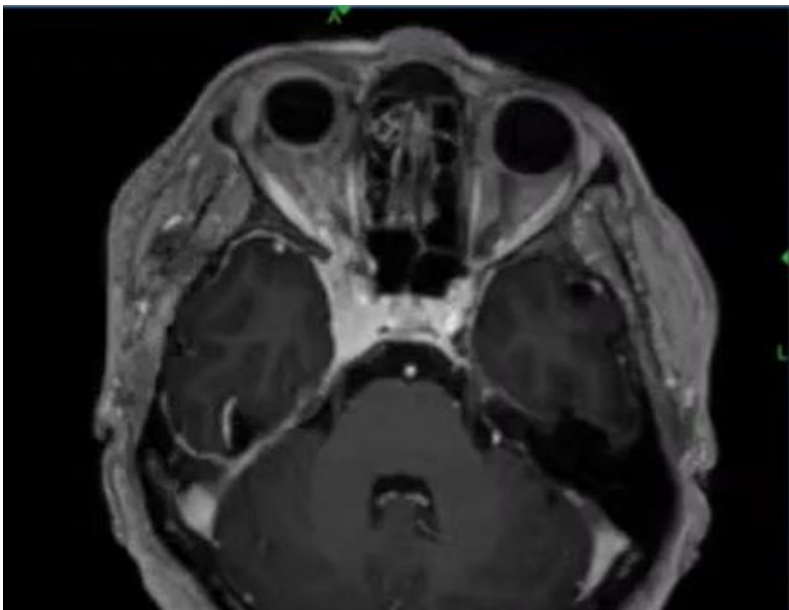
Impact of ^{68}Ga -DOTATOC PET/MRI on robotic radiosurgery treatment planning in meningioma patients: first experiences in a single institution

¹Gülüz Acker, MD,^{1,2} Anne Kluge, PhD,^{3,4} Mathias Lukas, MSc,⁵ Alfredo Conti, MD, PhD, FEBNS,^{1,3,6} Diana Pasemann, MSc,^{3,4} Franziska Meinert,^{1,3} Phuong Thuy Anh Nguyen,^{1,3} Claudius Jelgersma,^{1,3} Franziska Loebel, MD,^{1,3} Volker Budach, MD,^{3,4} Peter Vajkoczy, MD,^{1,3} Christian Furth, MD,⁵ Alexander D. J. Baur, MD,⁵ and Carolin Senger, MD^{3,4}

- Easier to define target
- planning volumes showed significantly smaller per physician
- preference for PET/MRI by radiosurgeons (particularly in proximity to critical structures)

Impact of ⁶⁸Ga-DOTATOC PET/MRI on robotic radiosurgery treatment planning in meningioma patients: first experiences in a single institution

*Güliz Acker, MD,^{1,3} Anne Kluge, PhD,^{3,4} Mathias Lukas, MSc,⁵ Alfredo Conti, MD, PhD, FEBNS,^{1,3,6} Diana Pasemann, MSc,^{3,4} Franziska Meinert,^{1,3} Phuong Thuy Anh Nguyen,^{1,3} Claudius Jelgersma,^{1,3} Franziska Loebel, MD,^{1,3} Volker Budach, MD,^{3,4} Peter Vajkoczy, MD,^{1,3} Christian Furth, MD,⁵ Alexander D. J. Baur, MD,⁵ and Carolin Senger, MD^{3,4}



Skullbase meningioma Radiosurgery Complications



Complications SRS for Skullbase Meningiomas

- Neuropathy
- Optic pathway toxicity
- Facial Nerve toxicity
- Radiation Necrosis Brain
- Oedema
- Pituitary gland hormone deficit
- Headache

neurological deterioration



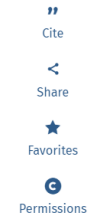
- Incidence of **neurological deterioration**, or development of **new neurological deficits** in those series with long-term follow-up, has been relatively **low**.
- Approximately **80%** to **100%** of patients **preserve neurological functions**

ISRS guidelines

complications

Neurological deterioration

- CS 10.8%
- PC 15%
- CPA 14.2%



RESEARCH—HUMAN—CLINICAL STUDIES

Long-term Tumor Control of Benign Intracranial Meningiomas After Radiosurgery in a Series of 4565 Patients

Santacrose, Antonio MD¹; Walier, Maja Dipl Math²; Régis, Jean MD, PhD³; Liščák, Roman MD, PhD⁴; Motti, Enrico MD⁵; Lindquist, Christer MD, PhD⁶; Kemeny, Andras MD^{7*}; Kitz, Klaus MD^{2,†}; Lippitz, Bodo MD^{8,§}; Álvarez, Roberto Martínez MD, PhD¹; Pedersen, Paal-Henning MD, PhD^{9,†}; Yomo, Shoji MD⁵; Lupidi, Francesco MD^{10,¶}; Dominikus, Karlheinz PhD^{11,||}; Blackburn, Philip MD^{12,||}; Mindermann, Thomas MD^{13,§§}; Bundschuh, Otto MD¹⁴; van Eck, A.T.C.J. MD^{15,††}; Fimmers, Rolf PhD²; Horstmann, Gerhard A. MD^{16,†††}

Permanent morbidity rates

CS 5.9%

PC 8.4%

CPA 8.3%

Post-SRS Edema

- **Skullbase** meningiomas **lower risk** of post-SRS edema
- Edema is related with **Dose** and **Volume**
- Edema is NOT DIRECTLY RELATED with Dose and Volume

Multivariate mechanism with unclear relationship



Risk of radiation-associated intracranial malignancy after stereotactic radiosurgery: a retrospective, multicentre, cohort study

THE LANCET
Oncology

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ARTICLES | VOLUME 20, ISSUE 1, P159-164, JANUARY 2019

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Risk of radiation-associated intracranial malignancy after stereotactic radiosurgery: a retrospective, multicentre, cohort study

Amparo Wolf, PhD • Kyla Naylor, PhD • Moses Tam, MD • Akram Habibi, BSc • Josef Novotny, PhD • Prof Roman Liščák, MD • et al. [Show all authors](#)

6.87 per 100 000 patient-years for malignant transformation

2.26 per 100 000 patient-years for radiosurgery-associated intracranial malignancy

- estimated risk for **intracranial secondary malignancy** or **malignant transformation** of a benign tumour in patients treated with **stereotactic radiosurgery** remains low at long-term follow-up
- **similar to the risk** of the general population to have a **primary CNS tumour**

<https://sci-hub.se/10.1016/>

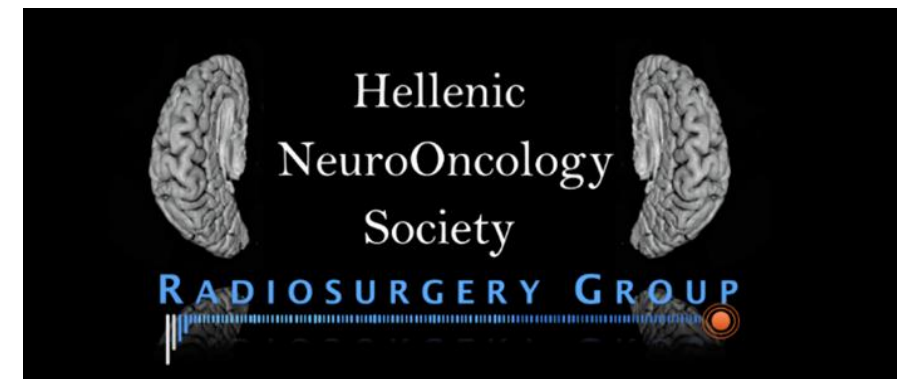
Douglas Kondjiolka et al,
Lancet 2018

“Stereotactic Radiosurgery”



“Stereotactic Radiosurgery”

- **Dedicated** in SRS
- International **SRS Training and Practice** 2006
- International **SRS Certification** 2010
- Establish “**SRS Group**” of Hellenic Neuro-Oncology Society 2017
- Establish Greek **Radiosurgery Guidelines** 2019
- **Affiliated ISRS members** of 2020
- **Linkedin** “Stereotactic Radiosurgery” (8500 members) 2021
- Represented in **Radiosurgery Board** 2021
 - Greek Ministry of Health
 - Greek Public Insurance (EOPYY)




CERTIFICATE OF AFFILIATED MEMBERSHIP

This is to certify that

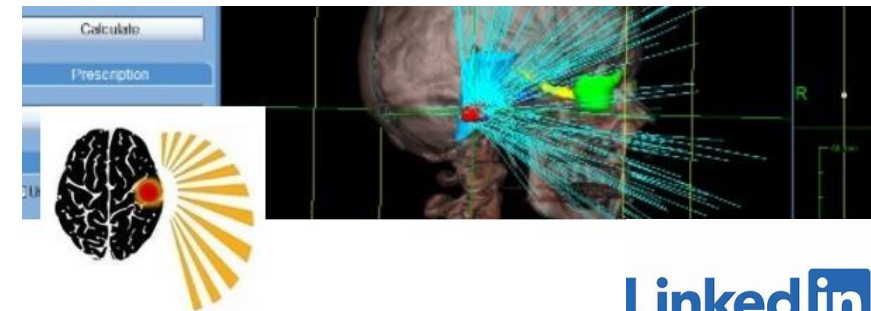
Hellenic Neuro-Oncology Society

is Officially Affiliated to the
International Stereotactic Radiosurgery Society (ISRS)
for 2020


Marc Levivier
ISRS Vice-president


Laura Fariselli
ISRS President


Arjun Sahgal
ISRS Secretary



LinkedIn

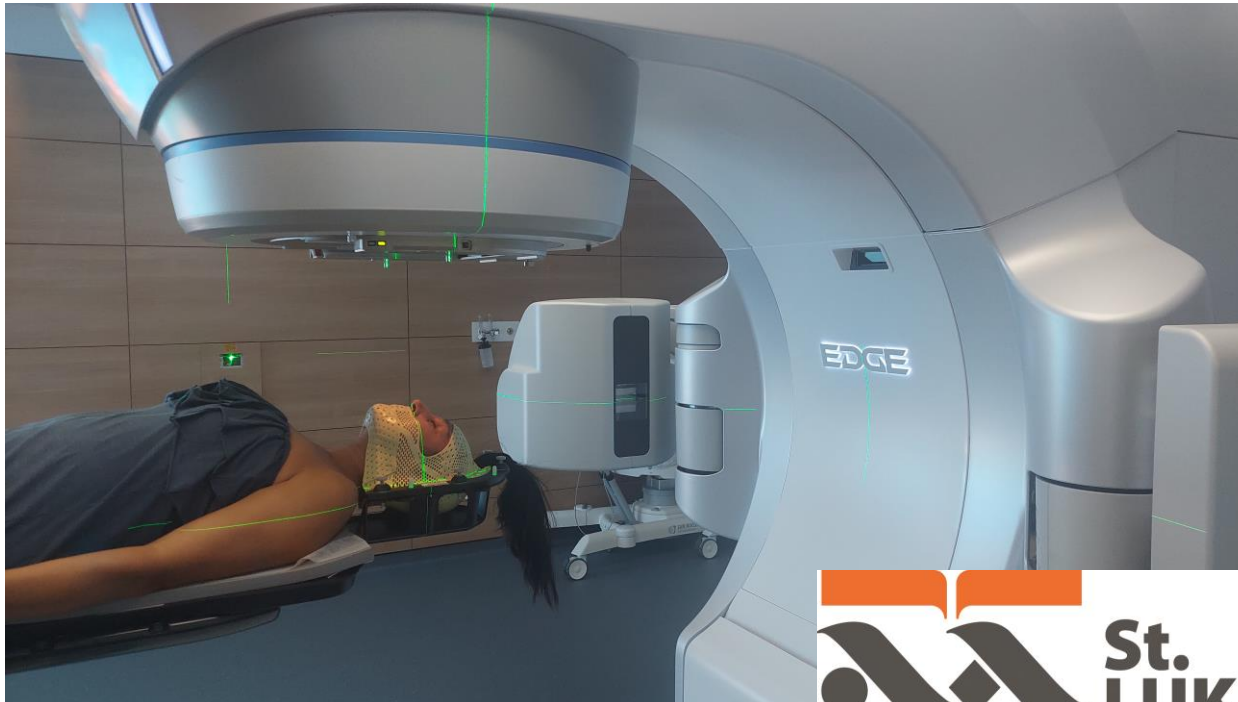
Stereotactic Radiosurgery

"...About Radiosurgery around the World..."

Hospitals and Health Care · Chalandri, Attica · 8K followers · 2-10 employees

Operating systems

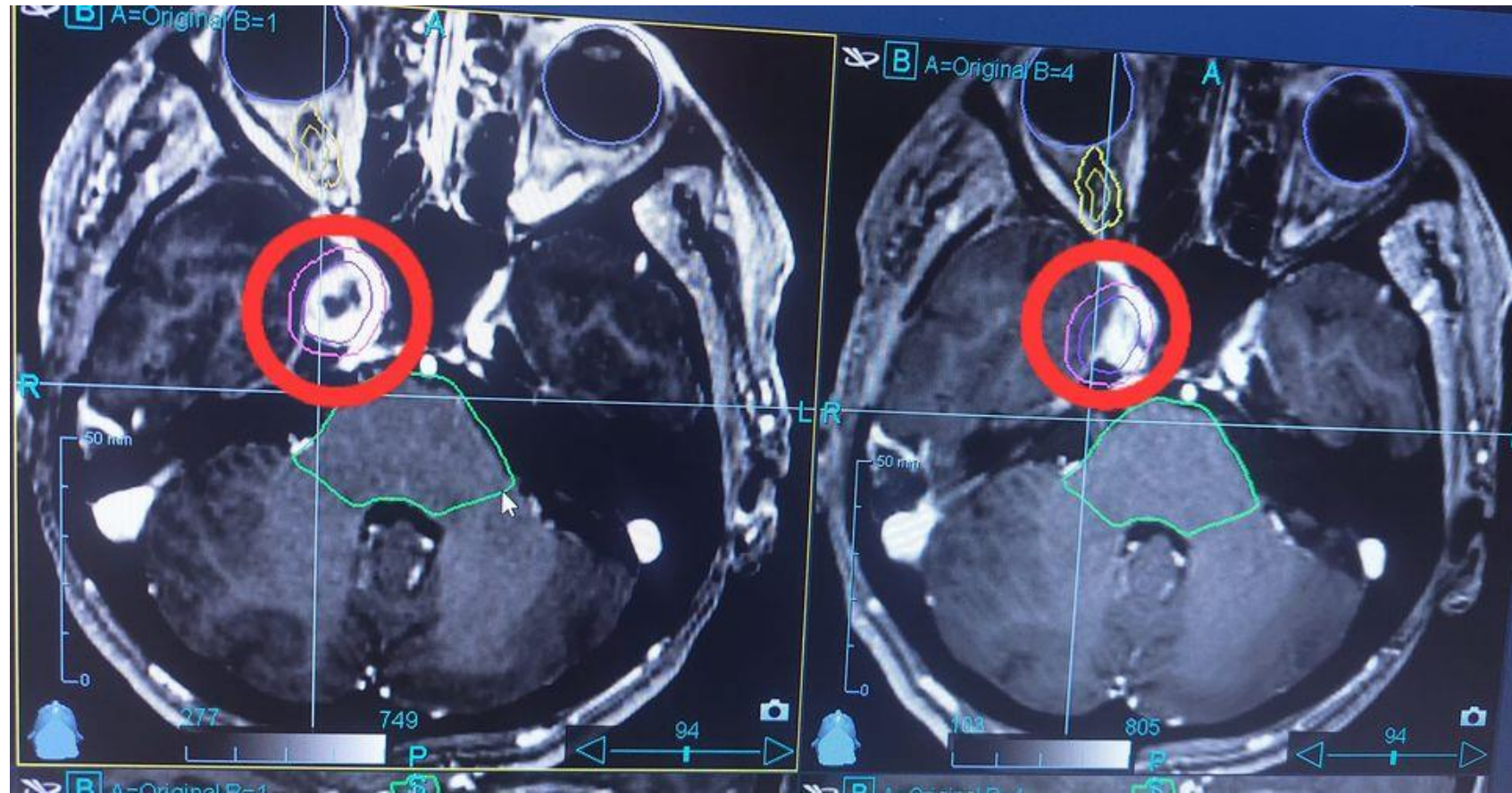
- Radiosurgery (isocentric) THESSALONIKI
- Robotic Radiosurgery (non isocentric) ATHENS



Case 1

Cavernous Sinus Meningioma treated with SRS

- Response after 3 years
- **Tumor Necrosis and Shrinkage**



Case 2

Meningioma treated with SRS

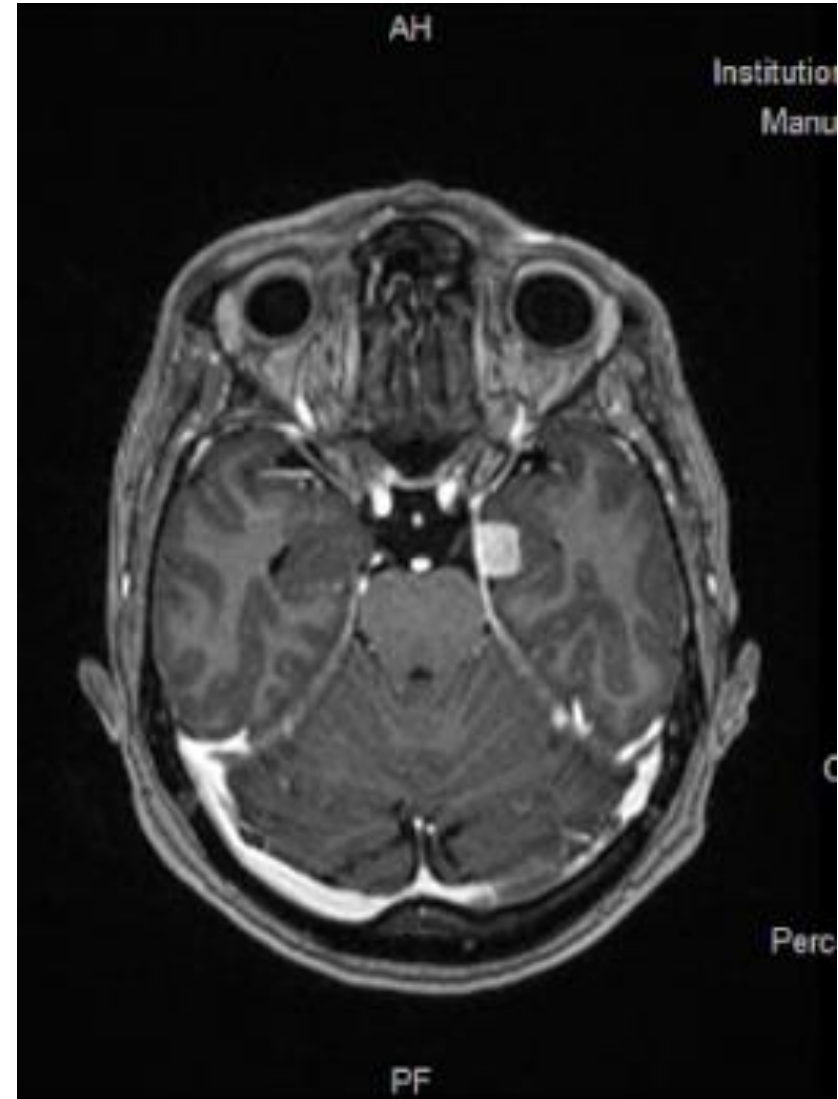
Female 52 yrs

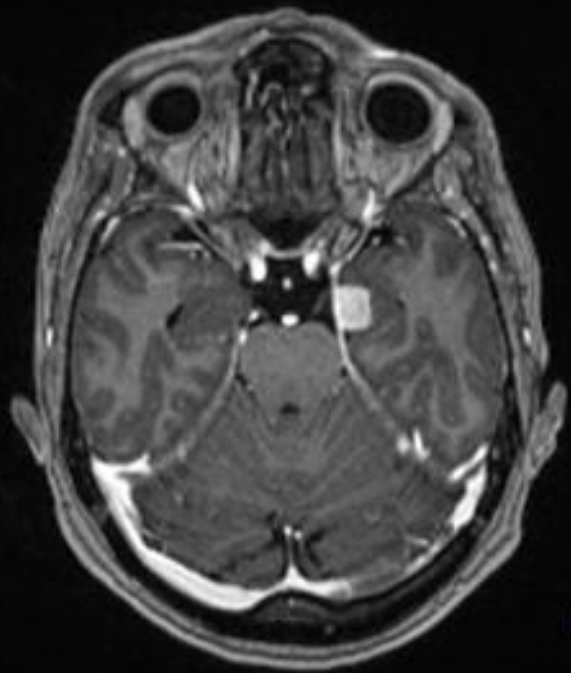
Meningioma

**Initial enlargement and Delay
Response after SRS**

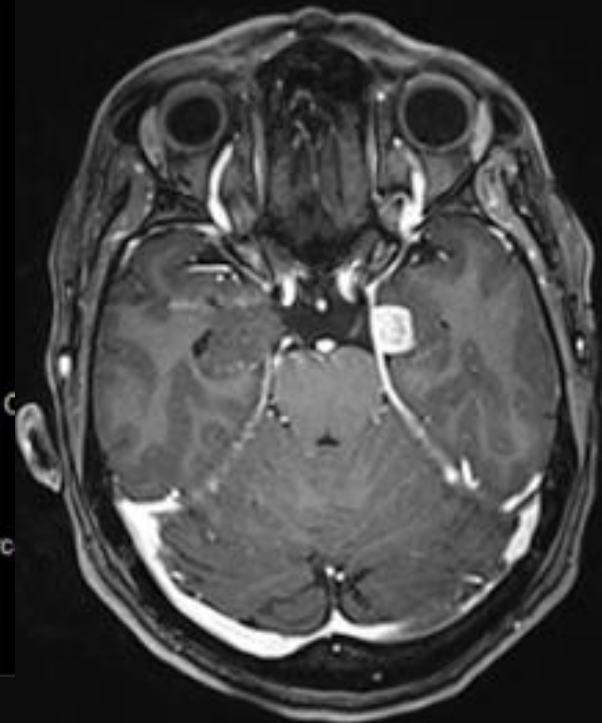
6months: enlargement

12 months Shrinkage

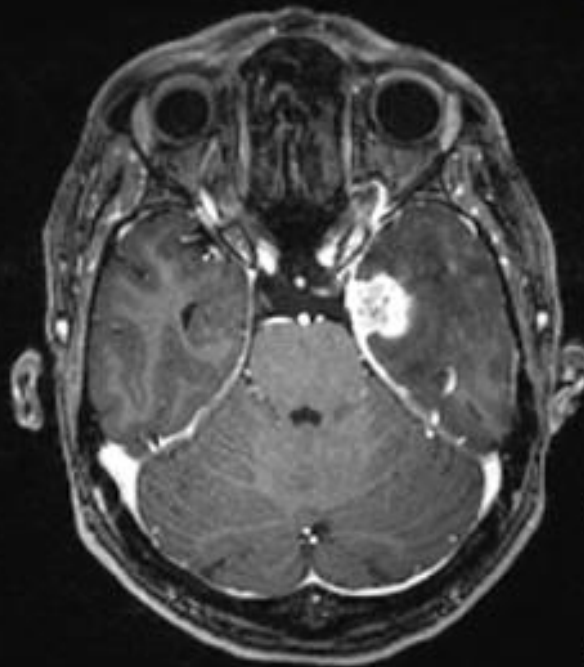




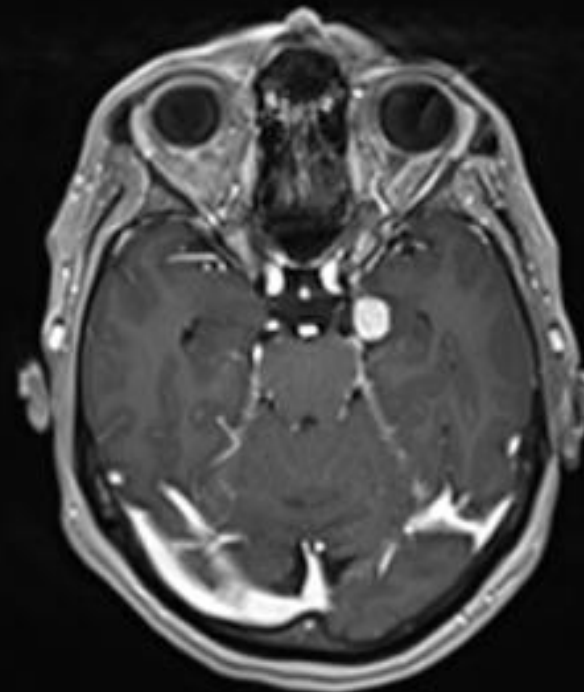
Sep 2022 SRS



Nov 2022, 2 Months



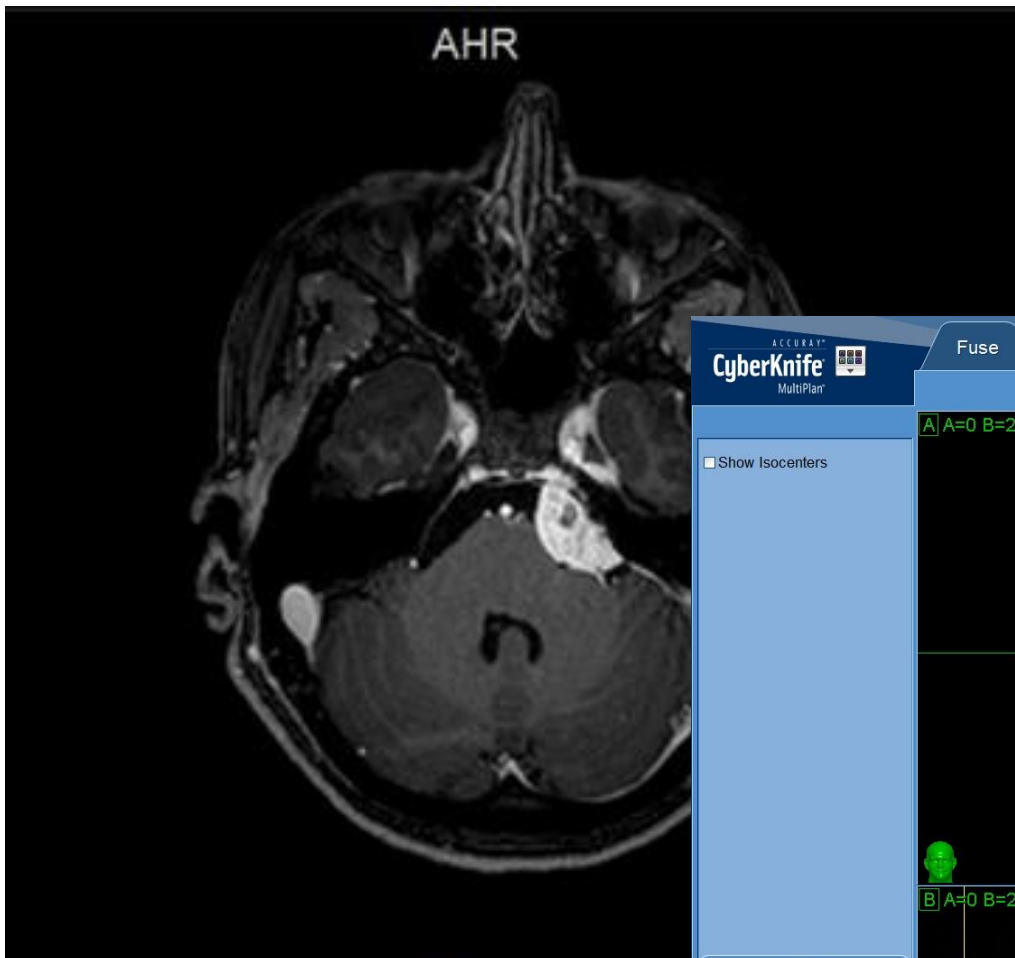
Apr 2023, 6 Months



Sep 2023, 12 Months

Case 3

Jan 2021: SRS



CyberKnife MultiPlan

Fuse Contour Align Plan Visualize Utilities Settings Help

Setup Isocentric Conformal Sequential Evaluate Finetune

■ Show Isocenters

Layouts

3D A	DVH Dose	3D S	DVH Dose
3D C	DVH Dose	3D S	A C

Standard Display

Patient
EVAGGELIA KLOTSOTIRA
1808217

Plan
Final_plan
21 Jan 2021, 11:58:05 AM

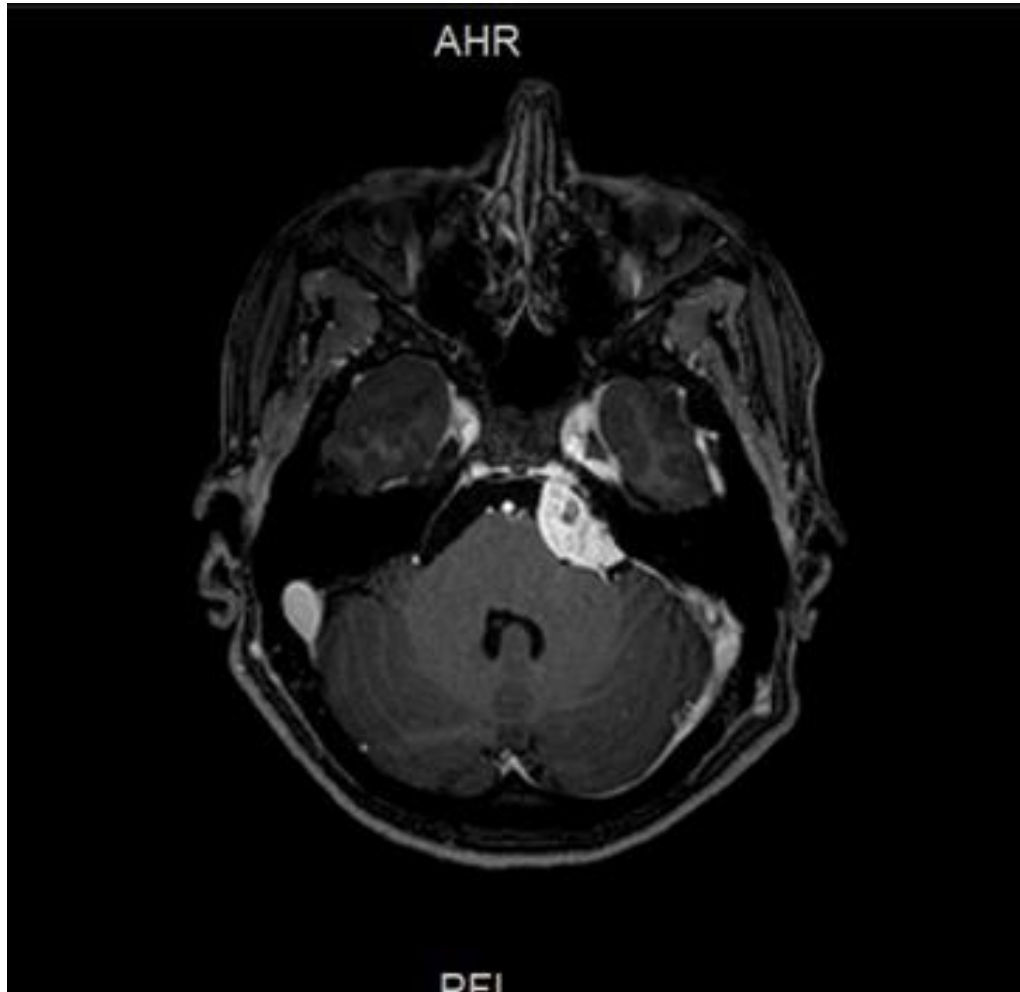
Rx
71%, 1400.00 cGy

Ray High (C) 71% 60% 50% 40% 30% 25% Of 1071.83 cGy

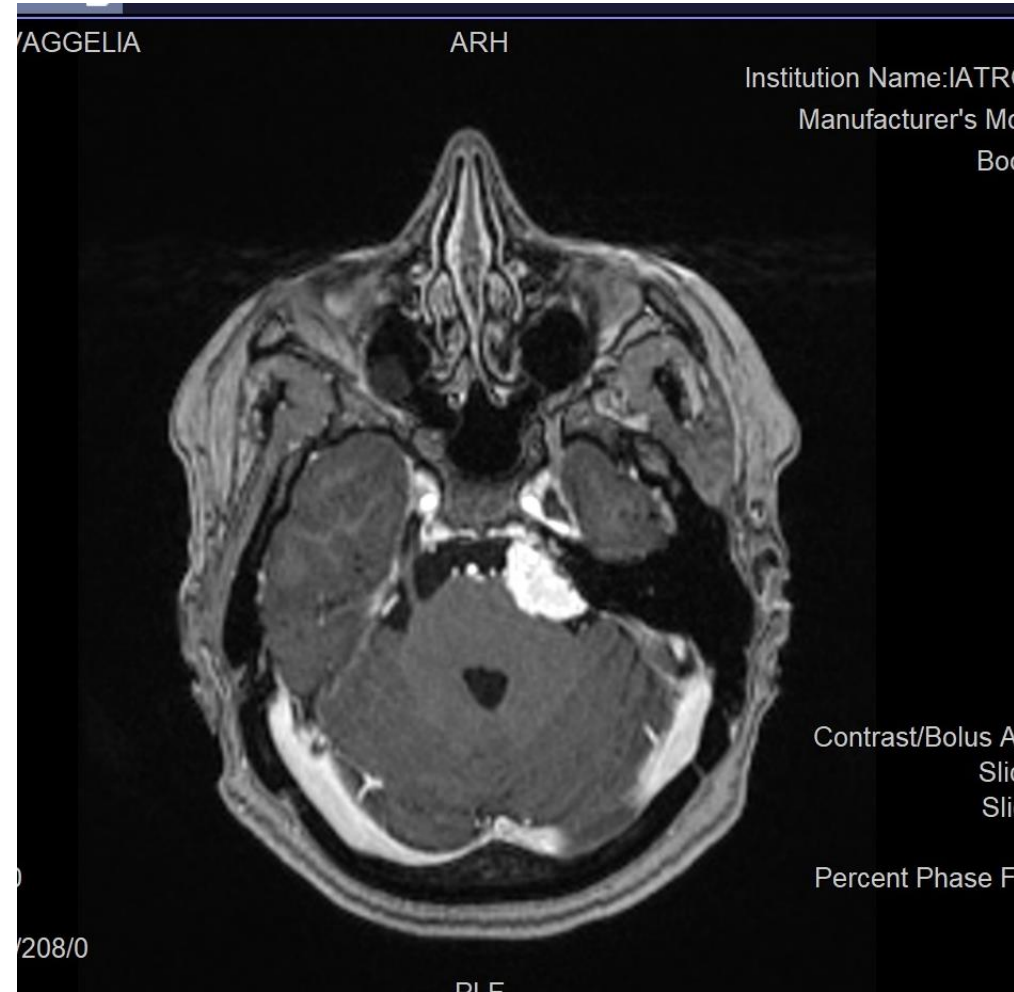
X:260 Y:256 Z:140 Value:1199

Female, 44 yrs
mild deficit ocular motor

Jan 2021: SRS



Feb 2023: Response



Relapse - Feb 2023 (2nd SRS)

23
1
56
nts:

100.00
:1960.00
ix:0/256/208/0
R

PLF

1808217
Plan
Final_plan2023
16 Feb 2023, 07:12:48 AM
Rx
74%, 1200.00 cGy

100%
90%
70%
60%
50%
40%
30%

100%
90%
70%
60%
50%
40%
30%

100%
90%
70%
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30%

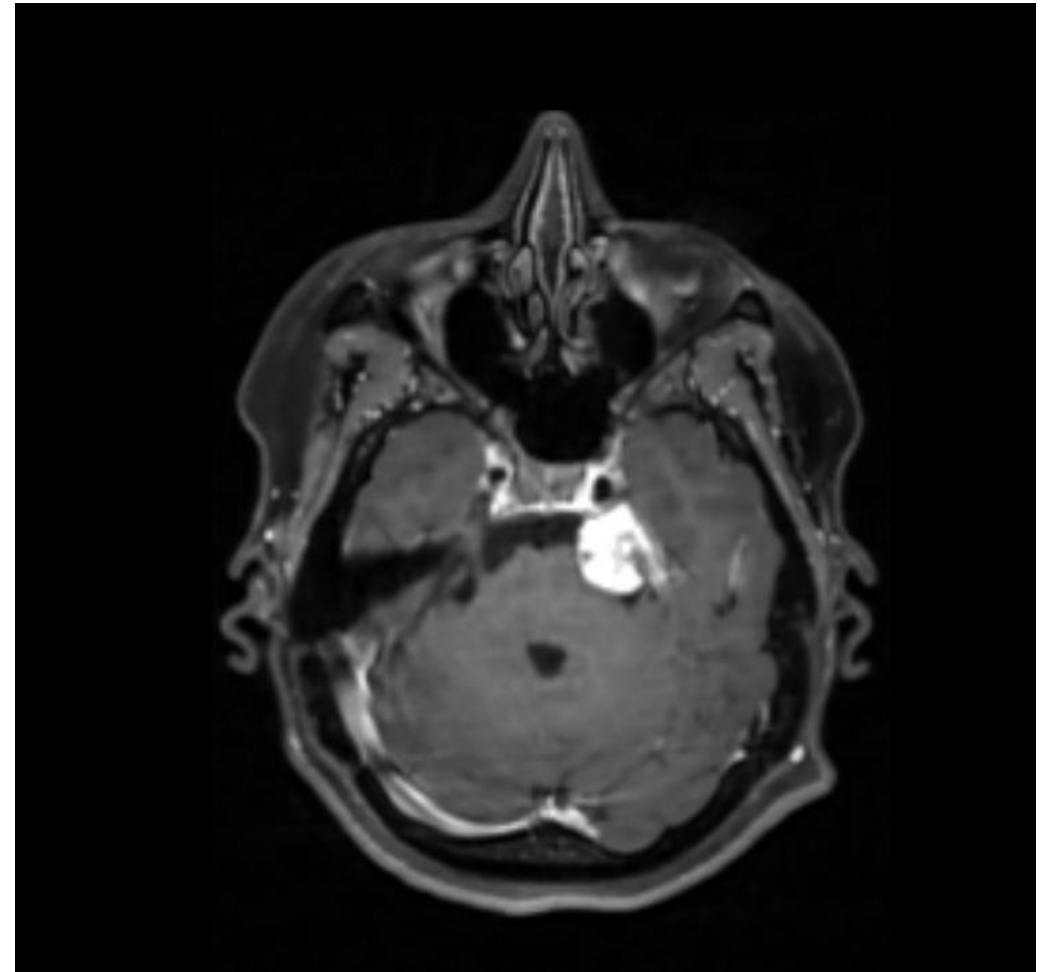
X:283 Y:227 Z:140 Value:1049



Relapse - Feb 2023



Local Control Aug 2024



Conclusions

- Radiosurgery is a **safe** and **efficient** therapy for skullbase meningioma
- high rate of **tumor response** and **neurological improvement**
- Low rates of **complications**
- **Molecular** and **Genetic profile** of the tumor is a challenge for better outcomes
- **Modern imaging** is a precious tool
- Selection of the proper **Dose** is crucial



THANKS!

OUR TEAM

Stereotactic Radiosurgery

"All about radiosurgery"



StereotacticRadiosurgery.gr
"All about Radiosurgery"

www.radiosurgery.gr



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