



Knowledge-Based Planning for SRS: From Quality Control to Full Automation

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UC San Diego
MOORES CANCER CENTER

Disclosure Statement

- 2012 and 2014 patent filings
- Varian Medical Systems
 - Licensing Agreement
 - Master Research Agreement
 - Consulting
 - Honoraria



Outline

- Treatment plan quality control
- What is knowledge-based planning (KBP)?
- Case study: KBP for SRS at UCSD
- The future of treatment planning for SBRT/SRS (and everything else)



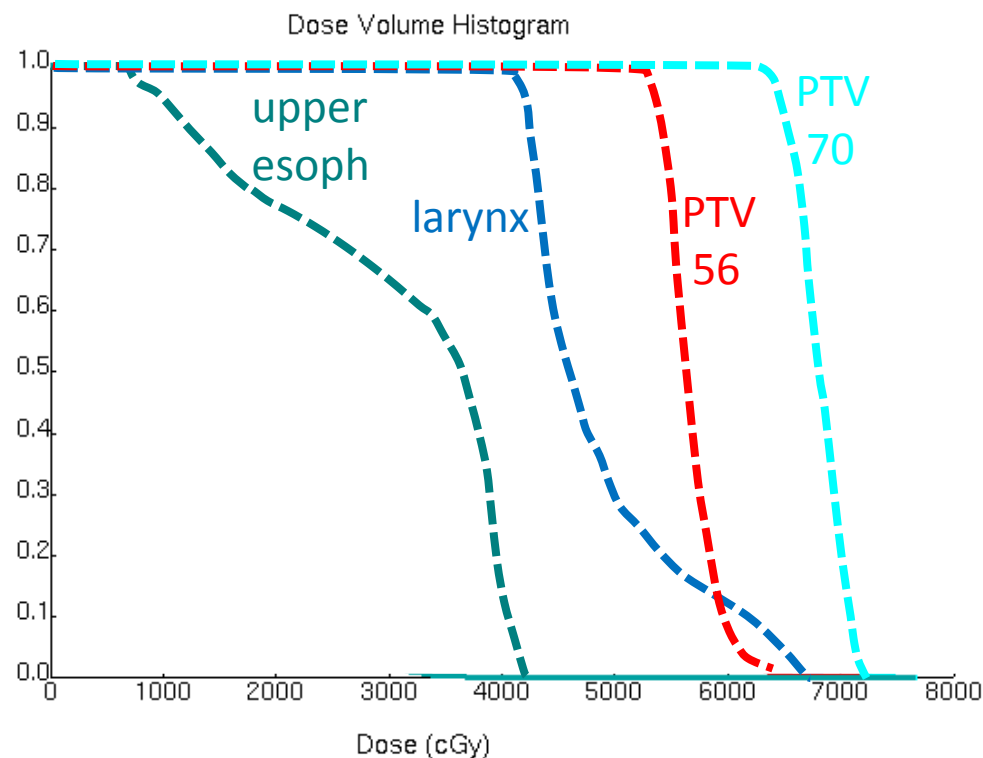
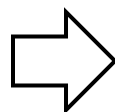
Do IMRT planning goals ensure “safe” plans?

		Bilateral Neck Treatment	Ipsilateral Neck Treatment
H&N	PTV	95% of PTV > 95% of Rx; Max dose < 110% of Rx	95% of PTV > 95% of Rx; Max dose < 110% of Rx
	Spinal Cord	Max dose 40 Gy	Max dose 40 Gy
	Spinal Cord + Margin	Max dose 52 Gy; < 1% (or 1 cc) exceeds 50 Gy	Max dose 52 Gy; < 1% (or 1 cc) exceeds 50 Gy
	Optic Nerves, Optic Chiasm	Max dose 54 Gy	Max dose 54 Gy
	Brainstem	Max dose 54 Gy; < 1% exceeds 60 Gy	Max dose 54 Gy; < 1% exceeds 60 Gy
	Brain	Max dose 60 Gy; < 1% exceeds 65 Gy	Max dose 60 Gy; < 1% exceeds 65 Gy
	Retina	Max dose 50 Gy; < 5% exceeds 45 Gy	Max dose 50 Gy; < 5% exceeds 45 Gy
	Larynx	As low as possible; mean dose < 45 Gy	As low as possible; mean Dose < 25 Gy
	Upper Esophagus	As low as possible; mean dose < 45 Gy	As low as possible; mean dose < 25 Gy
	Parotid	As low as possible; mean dose \leq 26 Gy	As low as possible; mean dose \leq 10 Gy (contralateral)
	Pharyngeal Constrictors	As low as possible; V60 < 60 Gy	As low as possible; V60 < 45 Gy
	Submandibular	As low as possible; mean dose < 39 Gy	As low as possible; mean dose < 24 Gy (contralateral)
	Oral Cavity	As low as possible; mean dose < 35 Gy	As low as possible; mean dose < 20 Gy
	Mandible	Max 70 Gy; < 5% exceeds PTV Rx	Max 70 Gy; < 5% exceeds PTV Rx
	Unspecified Tissue	Less than PTV Rx; < 5% exceeds PTV Rx	Less than PTV Rx; < 5% exceeds PTV Rx



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	Brainstem	Max dose 54 Gy; < 1% exceeds 60 Gy
	Brain	Max dose 60 Gy; < 1% exceeds 65 Gy
	Retina	Max dose 50 Gy; < 5% exceeds 45 Gy
	Larynx	As low as possible; mean dose < 45 Gy
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	Unspecified Tissue	Less than PTV Rx; < 5% exceeds PTV Rx

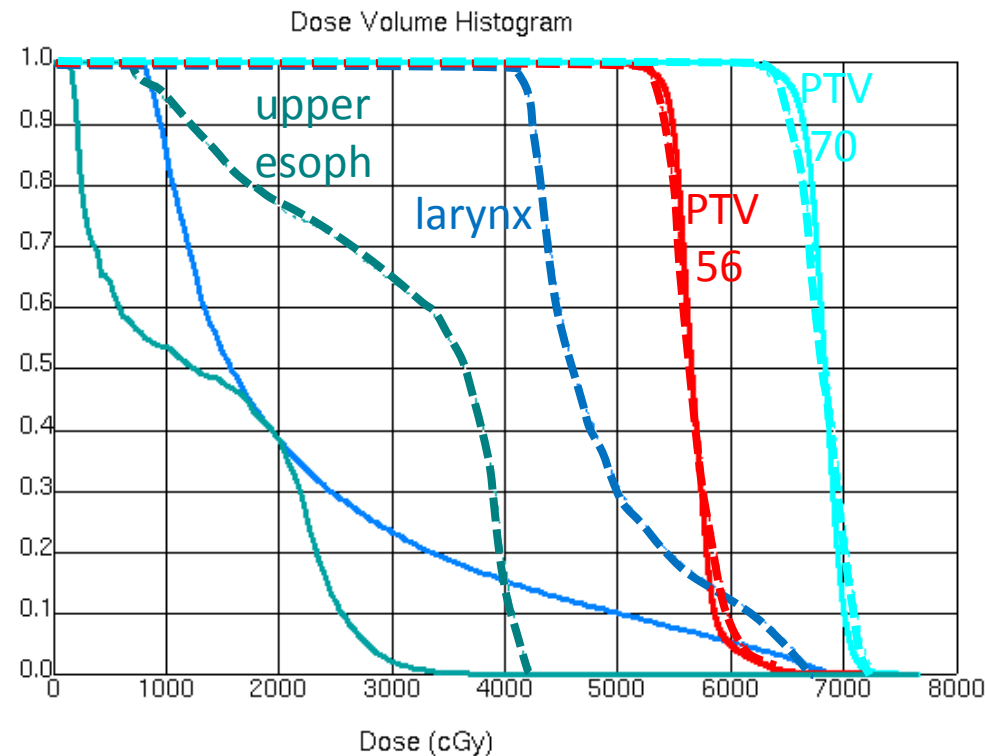
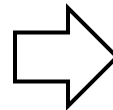


- This plan was QA'd at the treatment machine, passed all standard criteria.



Do IMRT planning goals ensure “safe” plans?

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	Unspecified Tissue	Less than PTV Rx; < 5% exceeds PTV Rx

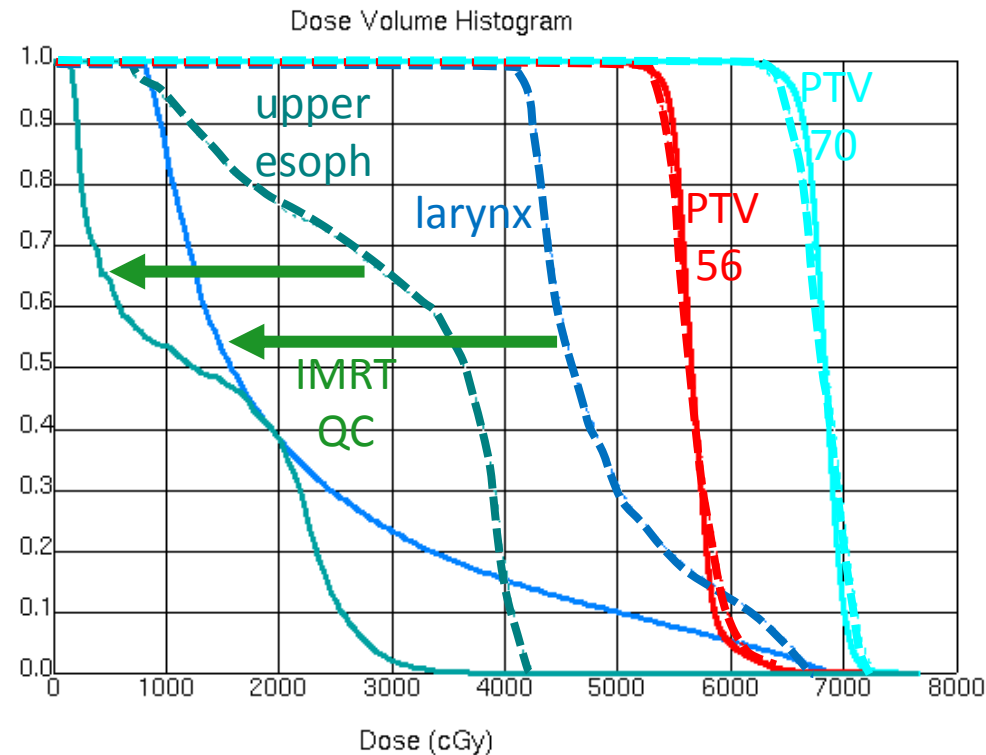
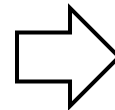


- (Dotted line plan was approved but not treated)
- Treatment plan was *safe* with respect to PTV coverage (TCP), but decidedly *unsafe* with respect to critical OARs (NTCP)



Do IMRT planning goals ensure “safe” plans?

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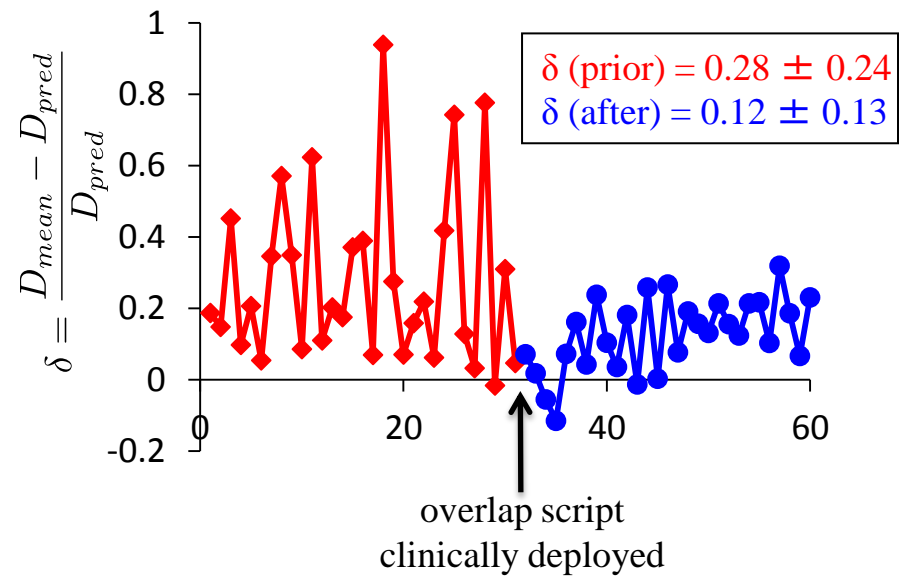
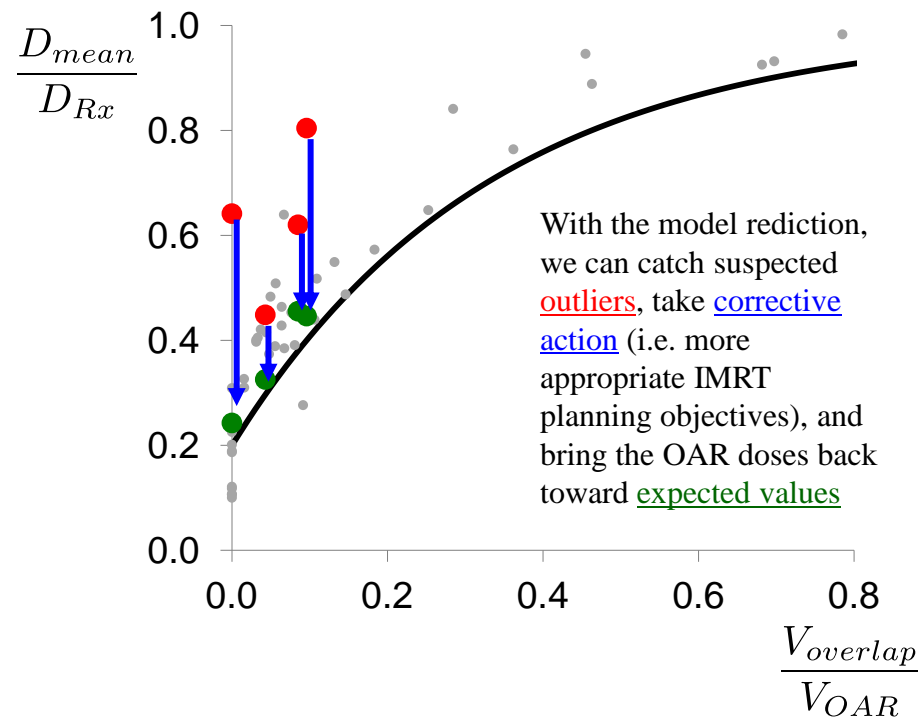


- Unless planning systems make trade-offs explicit, ignorance of what's possible can result in unsafe plan
- IMRT QC can addresses this problem on both input and output



The need for IMRT quality control

Goal is a system that can identify sub-optimal plans (most typically manifested as insufficient OAR sparing)



Salvageable parotids: 3 mos. before QC vs 3 mos. after

clinical plans
prior to feedback

D _{mean} (Gy)	D _{pred} (Gy)
18.9	11.6
16.1	12.3
16.9	13.3
14.9	15.2
24.7	18.0
26.6	18.8
26.4	19.6
36.6	21.0
27.4	23.6
46.8	24.2
43.4	27.7
40.5	29.1
52.3	29.4

Avg D _{mean}	Avg D _{pred}
33.6 Gy	22.4 Gy

clinical plans
after feedback

D _{mean} (Gy)	D _{pred} (Gy)
12.8	10.8
11.8	11.5
14.6	11.6
15.6	11.8
14.7	12.3
16.3	14.2
17.1	14.4
15.0	15.0
17.3	15.4
27.3	23.6
25.2	24.7
30.4	29.1
26.1	29.5

Avg D _{mean}	Avg D _{pred}
20.3 Gy	18.8 Gy

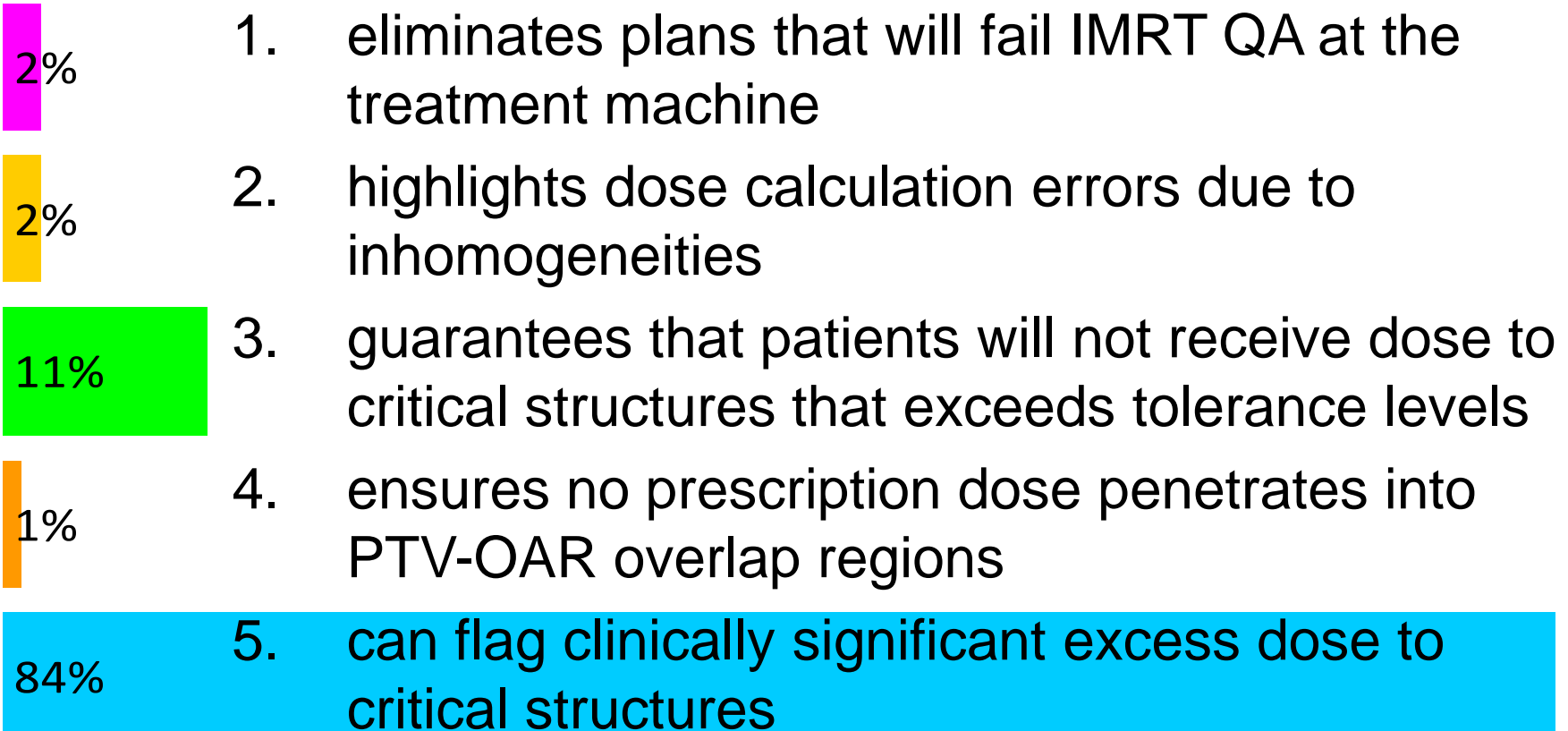


The need for treatment plan quality control

1. Need system that can identify sub-optimal plans (most typically manifested as insufficient OAR sparing)
2. Requirement is quantitative knowledge of what trade-offs must be made on the Pareto optimal frontier.
3. Absence of such a “system” will inevitably rely on subjective quality assessments and user experience/alertness... classic safety hazard!



Treatment Plan Quality Control:

- 
- A horizontal bar chart with five bars of different colors (magenta, yellow, green, orange, and blue) representing the percentage of treatment plans failing IMRT QA at different stages of quality control. The bars are arranged vertically, with the percentage value displayed to the left of each bar. The corresponding list item is to the right of each bar.
- | Percentage | Item |
|------------|--------------------------------------------------------------------------------------------------------|
| 2% | 1. eliminates plans that will fail IMRT QA at the treatment machine |
| 2% | 2. highlights dose calculation errors due to inhomogeneities |
| 11% | 3. guarantees that patients will not receive dose to critical structures that exceeds tolerance levels |
| 1% | 4. ensures no prescription dose penetrates into PTV-OAR overlap regions |
| 84% | 5. can flag clinically significant excess dose to critical structures |
1. eliminates plans that will fail IMRT QA at the treatment machine
 2. highlights dose calculation errors due to inhomogeneities
 3. guarantees that patients will not receive dose to critical structures that exceeds tolerance levels
 4. ensures no prescription dose penetrates into PTV-OAR overlap regions
 5. can flag clinically significant excess dose to critical structures

Correct Answer: 5

Can flag clinically significant excess dose to critical structures

[Experience-Based Quality Control of Clinical IMRT Planning](#)

Moore, Kevin L.; Brame, R. Scott; Low, Daniel A.; Mutic, S.;
INTERNATIONAL JOURNAL OF RADIATION ONCOLOGY * BIOLOGY *
PHYSICS Volume: 81 Issue: 2 Pages: 545-551

[Radiotherapy Dose-Volume Effects on Salivary Gland Function](#) Deasy,

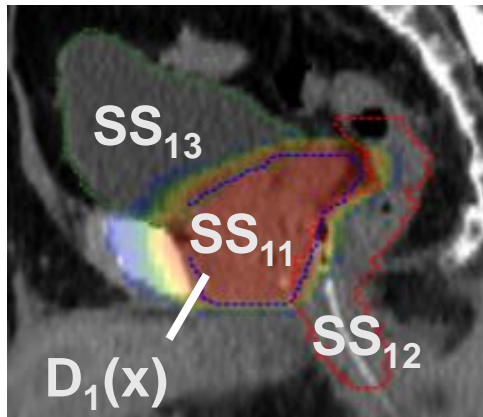
Joseph O.; Moiseenko, Vitali; Marks, Lawrence; Chao, K.S. Clifford;
Nam, Jiho; Eisbruch, Avraham; INTERNATIONAL JOURNAL OF
RADIATION ONCOLOGY * BIOLOGY * PHYSICS Volume: 76 Issue: 3
Pages: S58-S63

0D knowledge-based (single-variable) dose prediction

Step 1

- Identify a set of site similar training patients

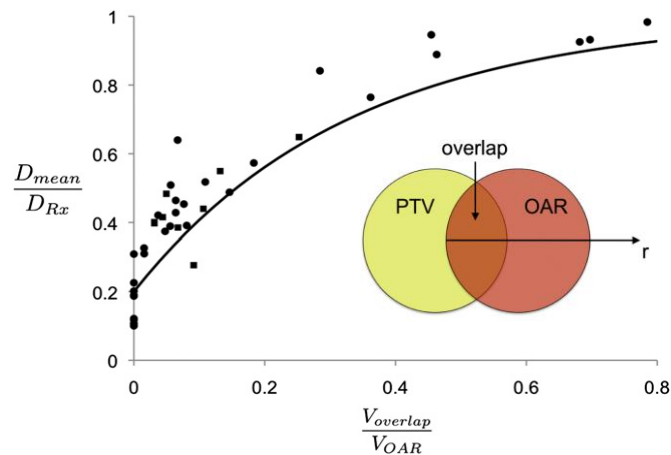
Patient 1



Patient N

Step 2

- Correlate mean dose with input geometry



Step 3

- Utilize prediction to obtain mean dose estimation for new patients

LT PAROTID Overlap

Volume = 13.5454 cc

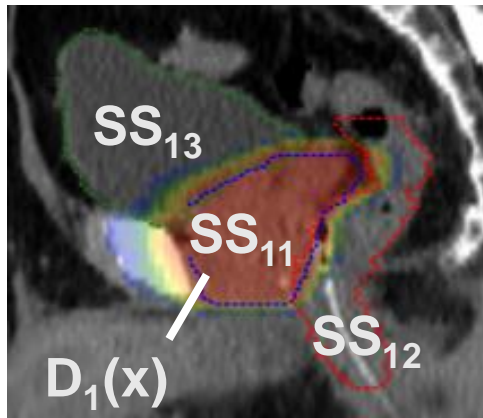
- PTV 5400 overlap volume = 0.967021 cc
Suggested Mean Dose = 1912.86 cGy
Current Mean Dose = 2038.5 cGy

0D \rightarrow 1D (DVH) knowledge-based dose prediction

Step 1

- Identify a set of site similar training patients

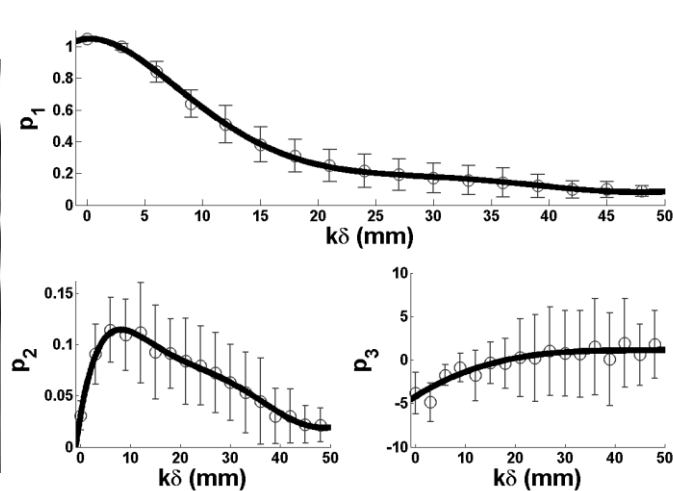
Patient 1



Patient N

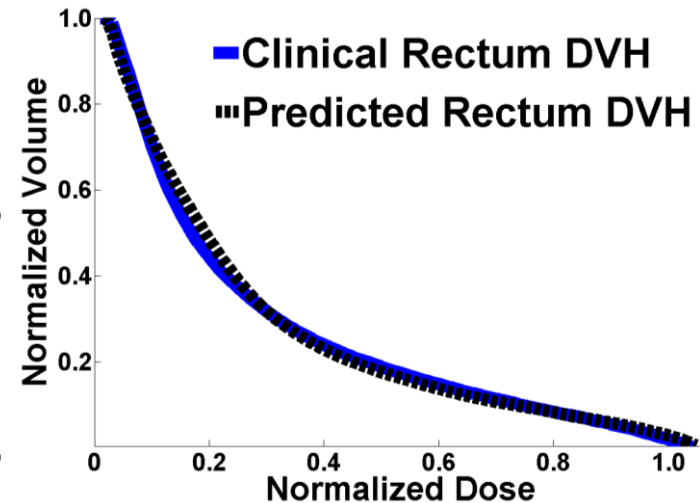
Step 2

- Generate pDVH model from training cohort



Step 3

- Utilize pDVH model to obtain DVH prediction for new patient

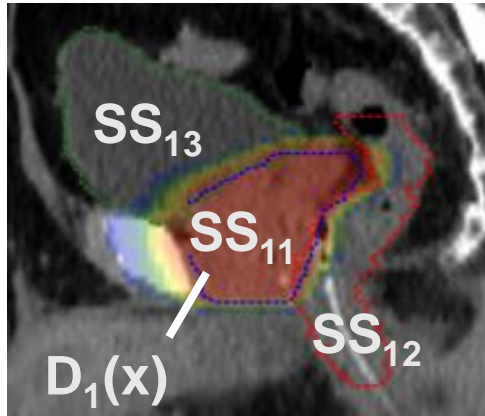


0D \rightarrow 1D \rightarrow 3D knowledge-based dose prediction

Step 1

- Identify a set of site similar training patients

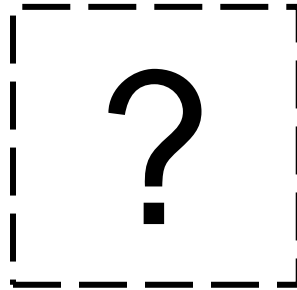
Patient 1



Patient N

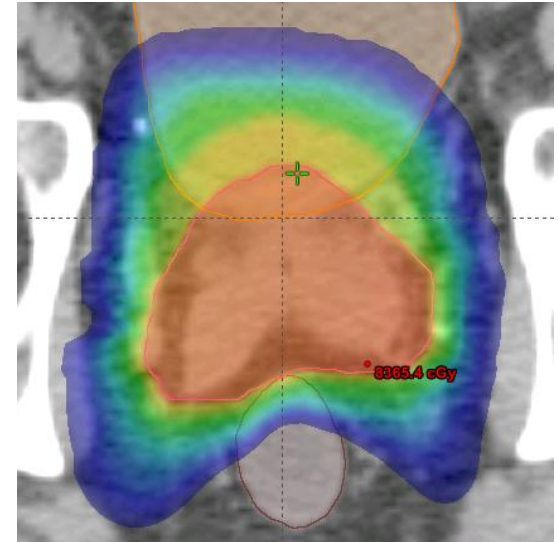
Step 2

- Generate 3D prediction model



Step 3

- Utilize model to obtain 3D dose prediction



IMRT QC = knowledge-based plan assessment

Key features of a “knowledge base”:

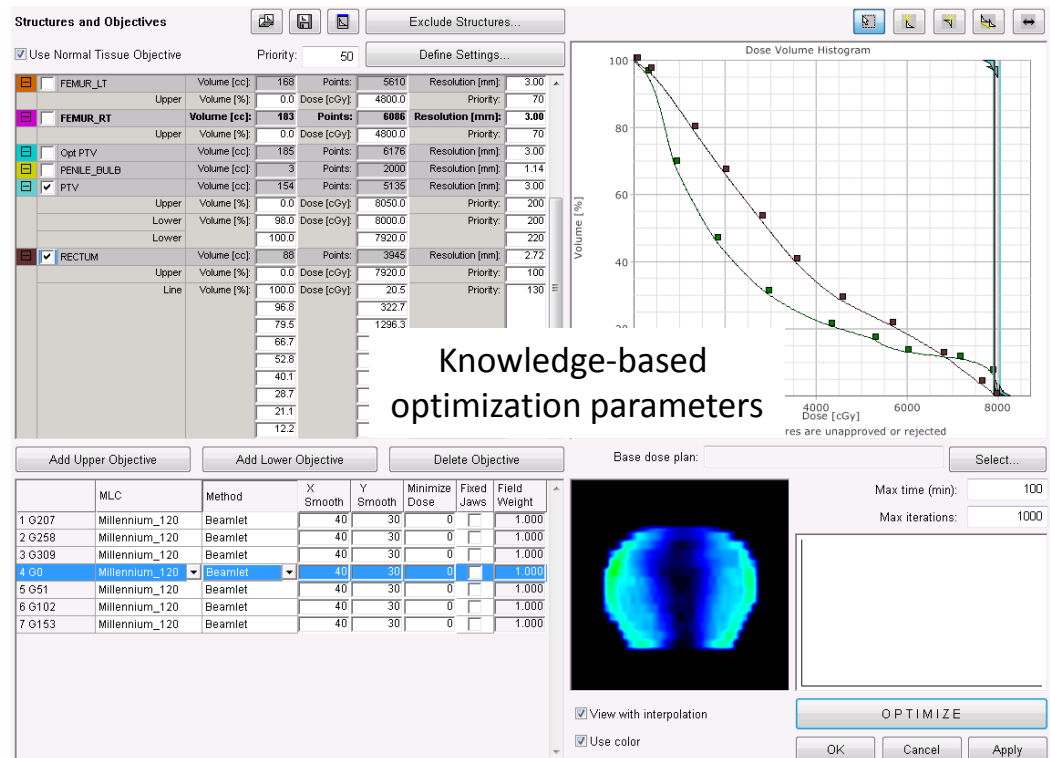
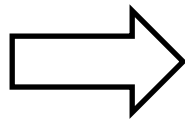
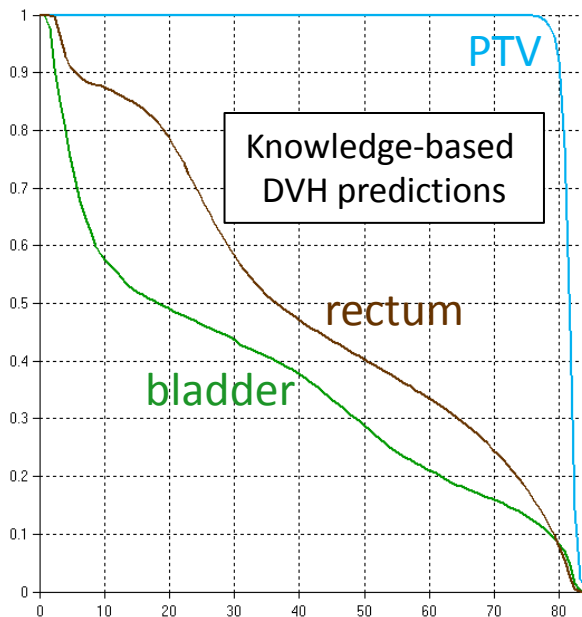
1. Must be quantitative
1. Must have discernable *correlations*
 - e.g. larger bladder = lower bladder DVH
3. Must provide a sufficient range of previous experience

With these ingredients, one has everything needed to make patient-specific predictions

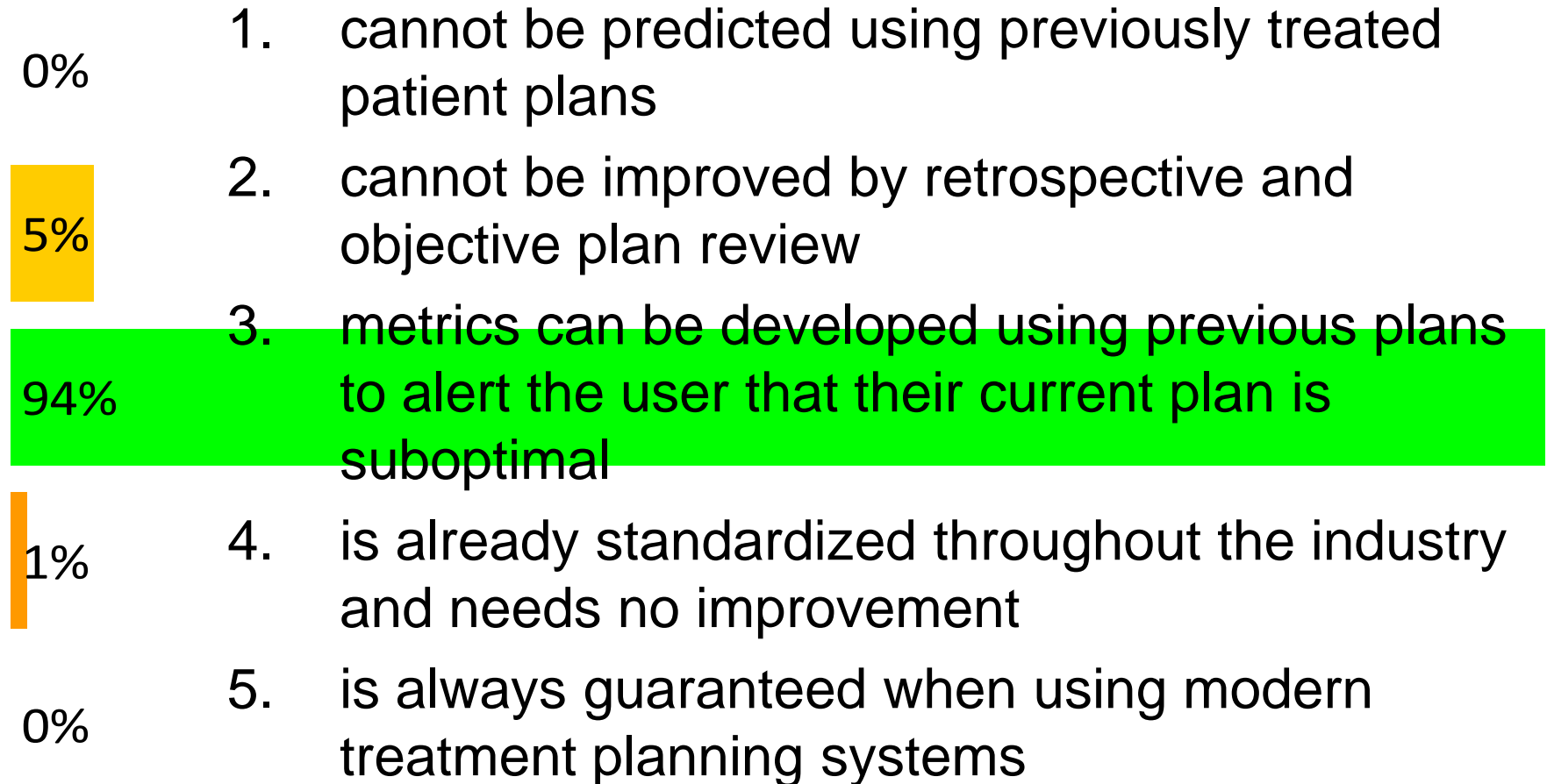


Knowledge-based planning “by hand”

- Knowledge-based planning involves nothing more than incorporating the dose-volume predictions directly into the optimization loop



Treatment plan quality:



Correct Answer: 3

Metrics can be developed using previous plans to alert the user that their current plan is suboptimal

[Quantitative Metrics for Assessing Plan Quality](#)

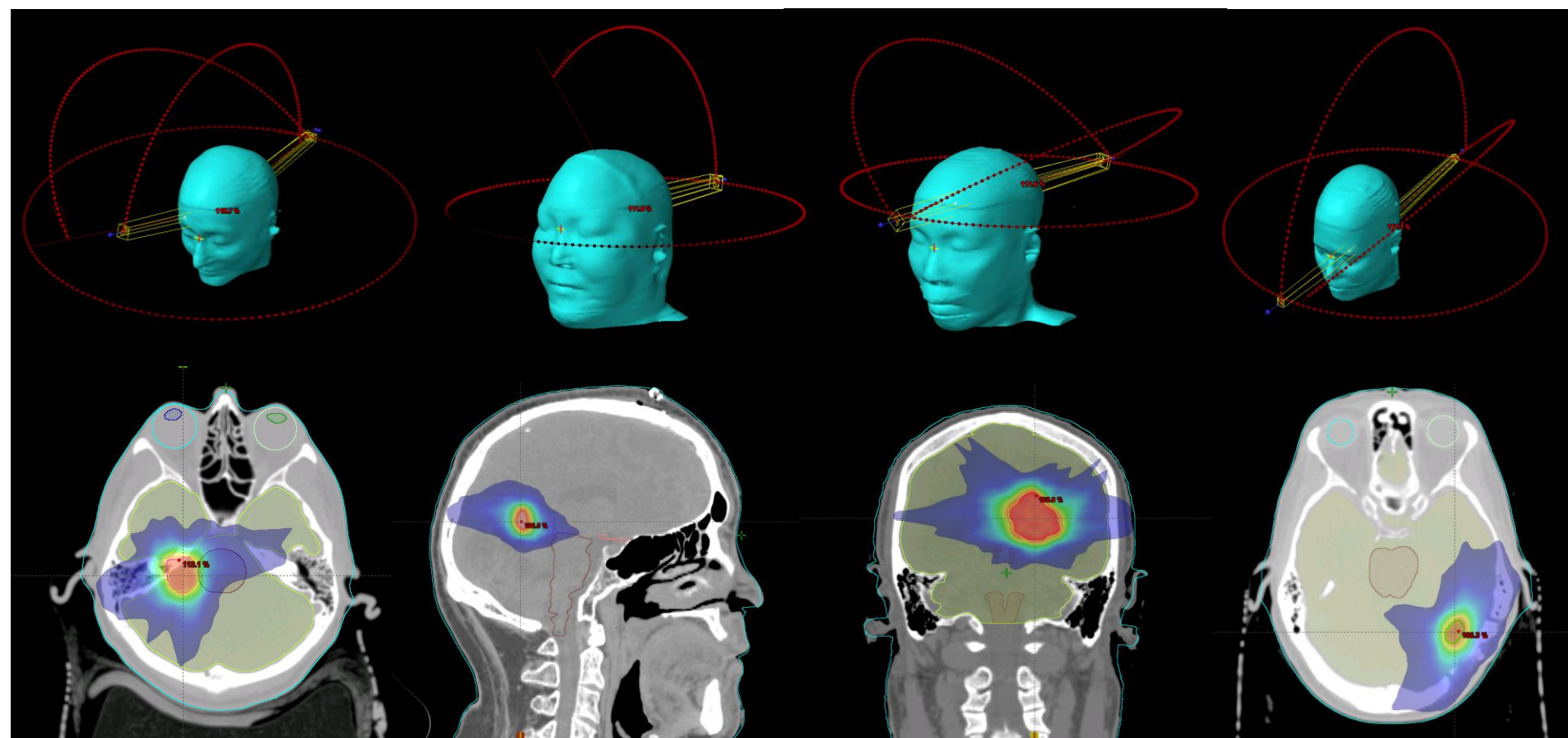
Moore, Kevin L.; Brame, R. Scott; Low, Daniel A.; et al. SEMINARS IN RADIATION ONCOLOGY Volume: 22 Issue: 1 Pages: 62-69

[Predicting dose-volume histograms for organs-at-risk in IMRT planning](#),

Appenzoller, Lindsey M.; Michalski, Jeff M.; Thorstad, Wade L.; et al. MEDICAL PHYSICS Volume: 39 Issue: 12 Pages: 7446-7461

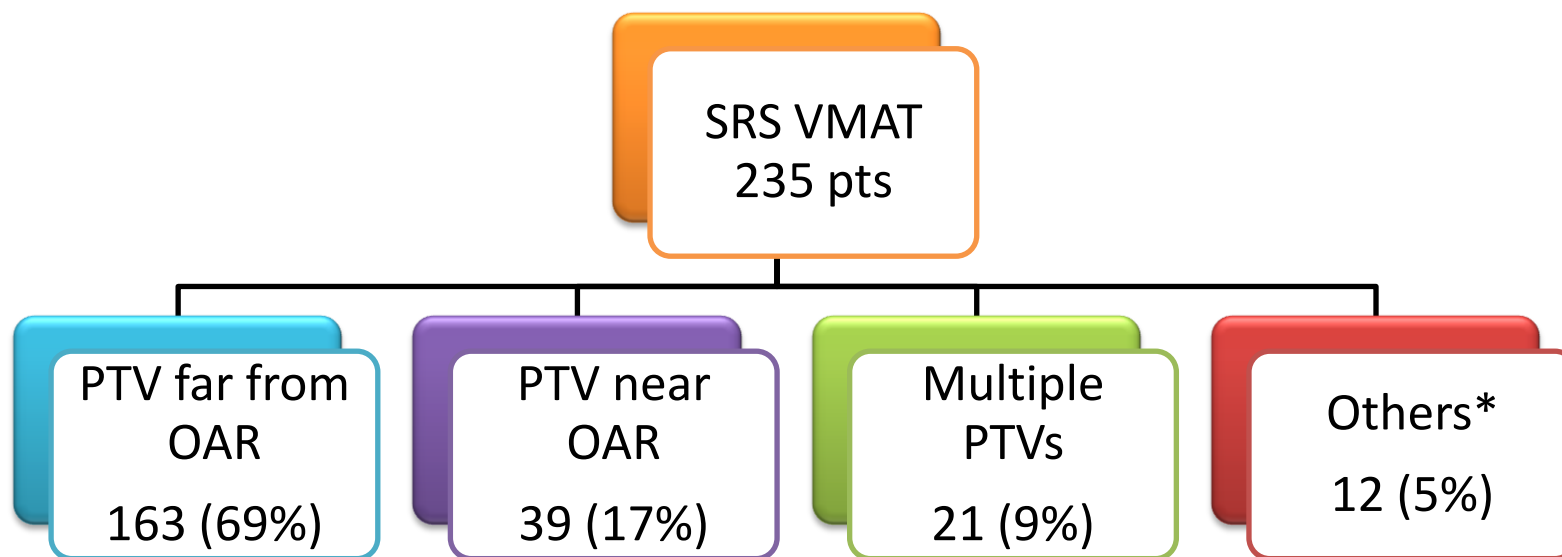
KBP in SRS: The UCSD experience

- For several years, standard treatment for SRS/SRT at UCSD has been multi-arc non-coplanar RapidArc



UCSD SRS experience

- SRS: Target size, shape, and location show enormous variation
 - PTV volume (0.1 cc - 60 cc)
 - Malignant vs. benign disease
 - Fractionation schedule and clinical priorities
 - Proximity to OARs (brainstem, optic nerves, cochlea) highly variable (0-10cm)
 - Multiple PTVs

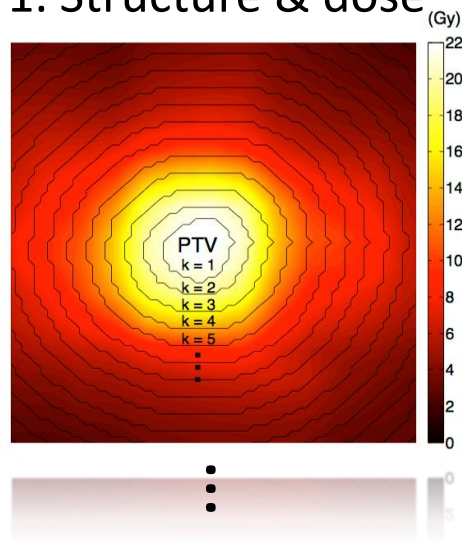


* Overlapping retreatment, staged approach for AVM



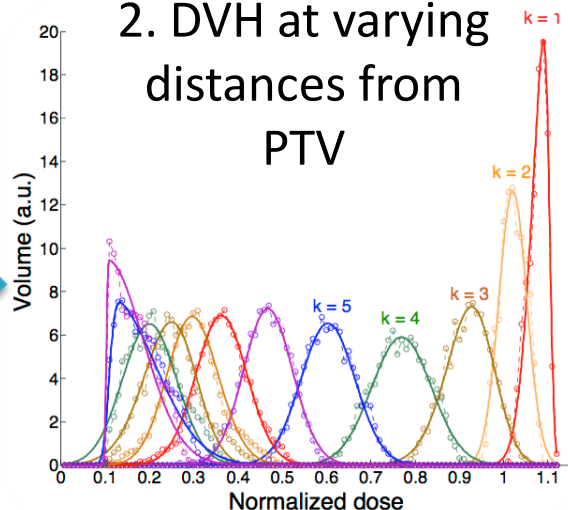
KBP in SRS

1. Structure & dose



N training plans

2. DVH at varying distances from PTV

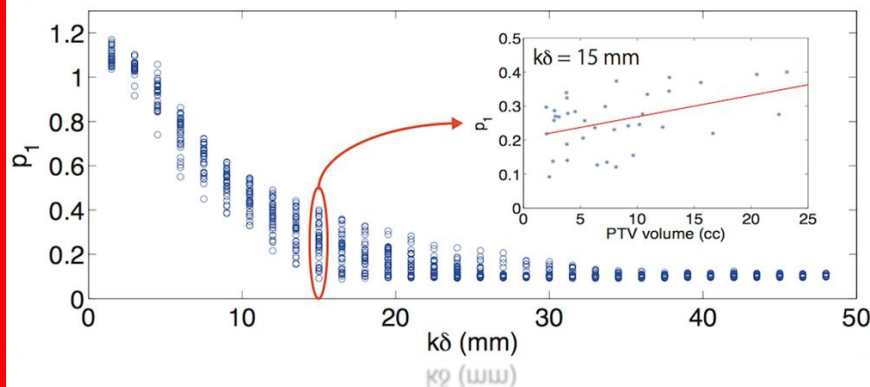


3. Fit with skew-normal PDF

$$f(p_1, p_2, p_3; D) = \frac{1}{\pi p_2} \exp\left(-\frac{(D-p_1)^2}{2p_2^2}\right) \times \int_{-\infty}^{p_3(D-p_1)/p_2} \exp(-t^2/2) dt,$$

Three fit parameters:
location, scale, shape

4. Parameterize fit parameters

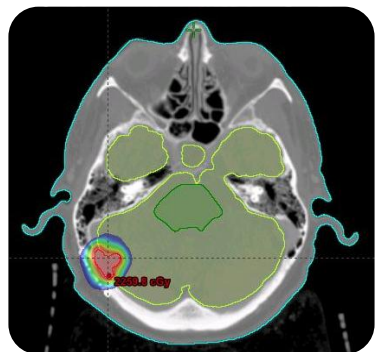


Fits include:
Distance from PTV
PTV volume



KBP in SRS

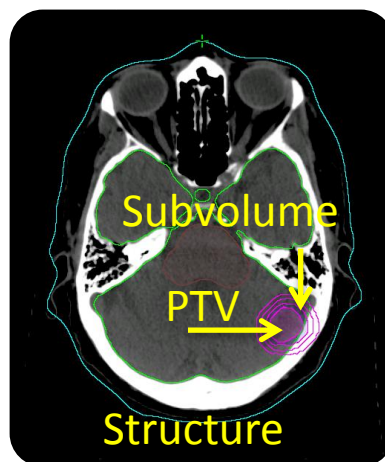
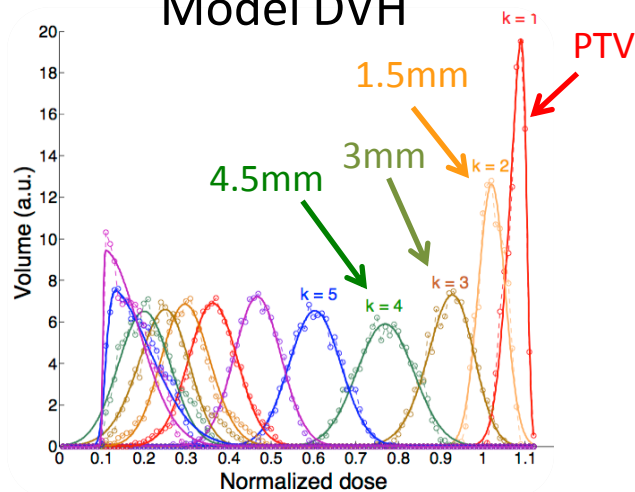
N training plans



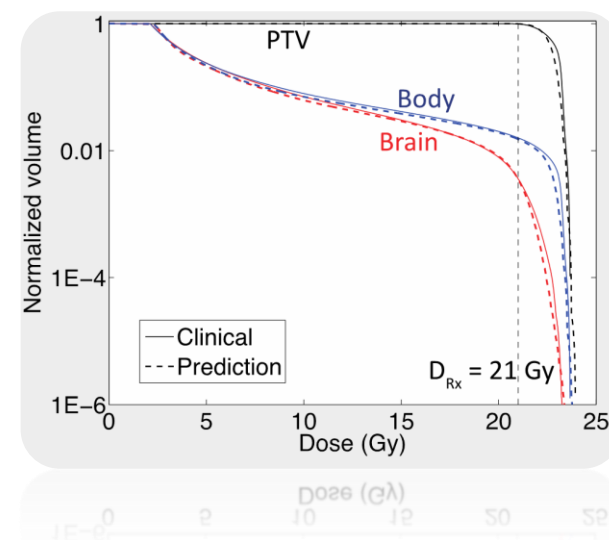
Structure & dose



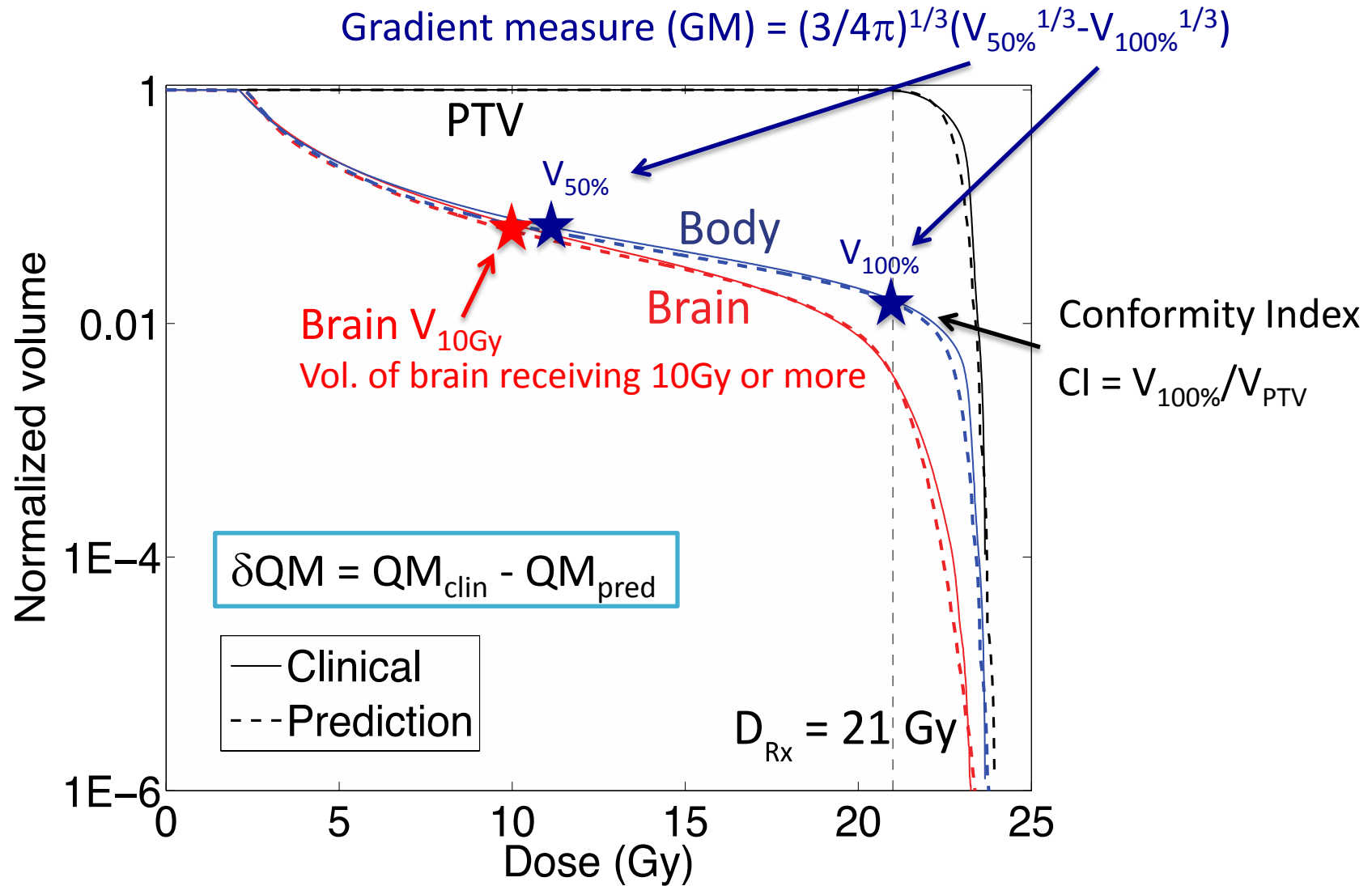
Model DVH



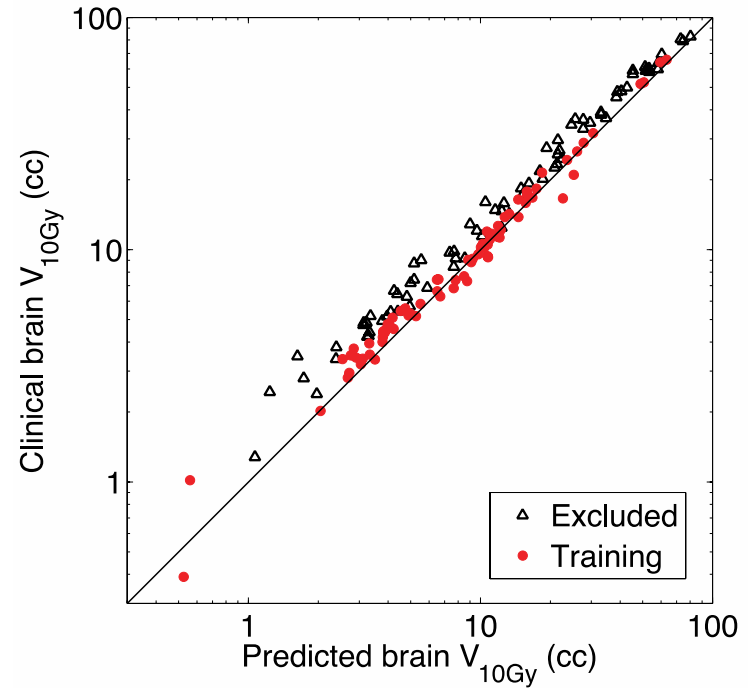
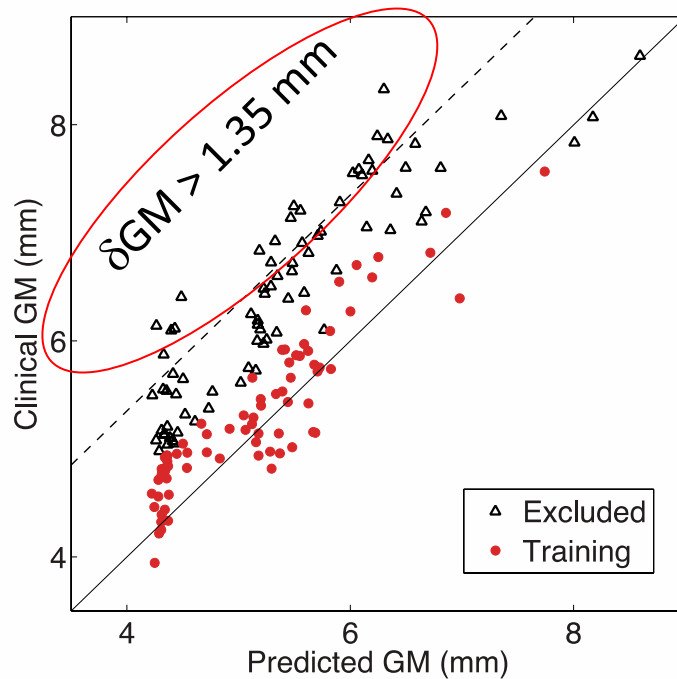
Predict DVH



SRS plan quality metrics are DVH-based



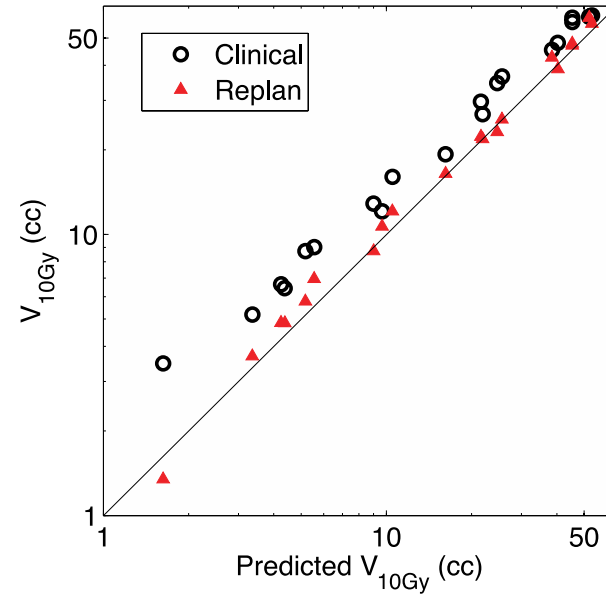
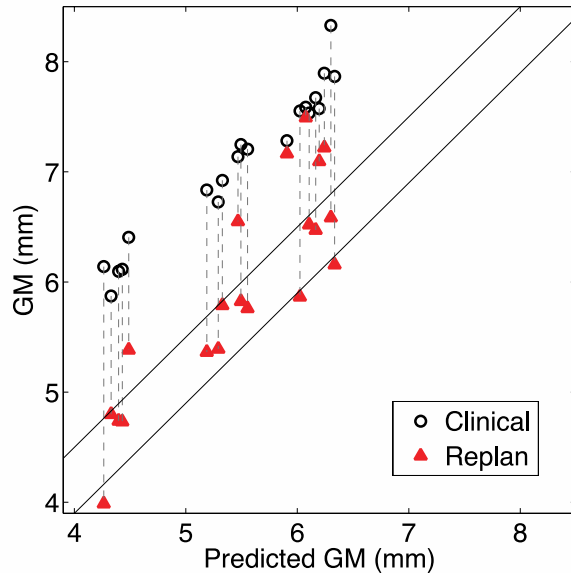
Accurate QM predictions and outlier identification



δQM	Training	Excluded	p-value
δGM (mm)	0.2 ± 0.3	1.1 ± 0.5	<0.001
$\delta V_{10\text{Gy}}/V_{10\text{Gy}}$	0.04 ± 0.12	0.20 ± 0.11	<0.001
δCI	-0.02 ± 0.12	-0.03 ± 0.10	0.19



KBP replanning confirms predicted clinical gains



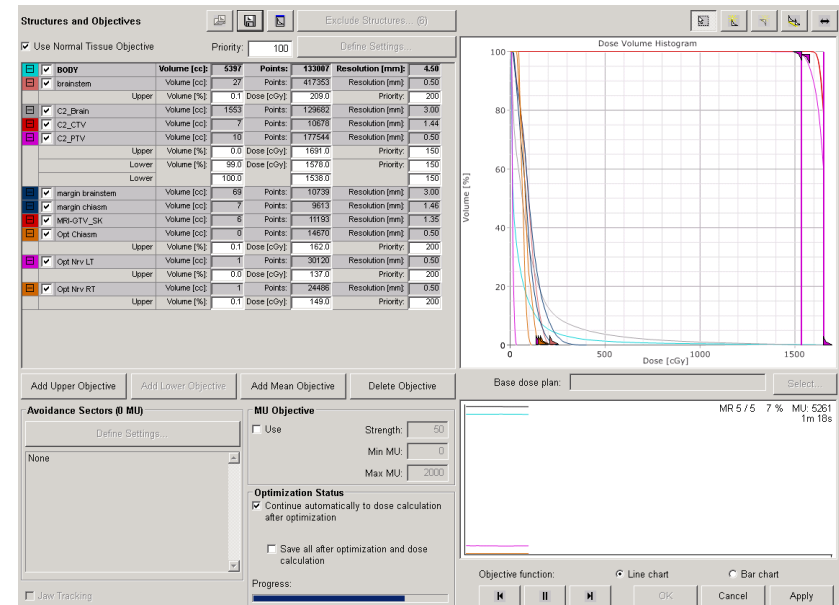
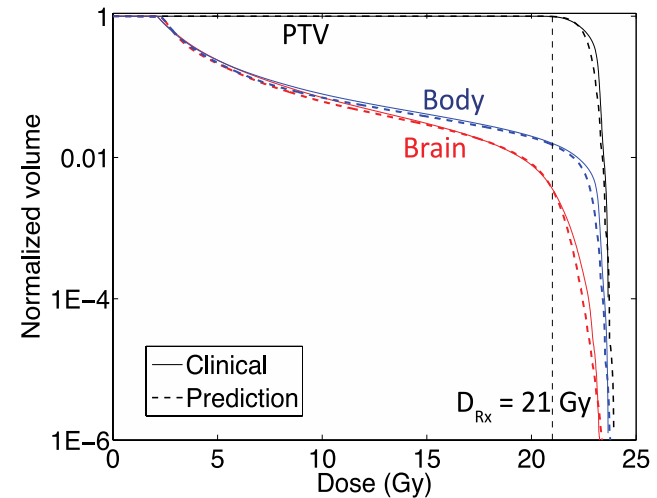
δQM	Clinical	Replan
δGM (mm)	1.6 ± 0.2	0.5 ± 0.5
$\delta V_{10Gy}/V_{10Gy}$	0.27 ± 0.11	0.04 ± 0.06
δCI	1.12 ± 0.09	1.08 ± 0.11
Max dose	1.10 ± 0.03	1.18 ± 0.04

Improved QMs,
Higher max dose



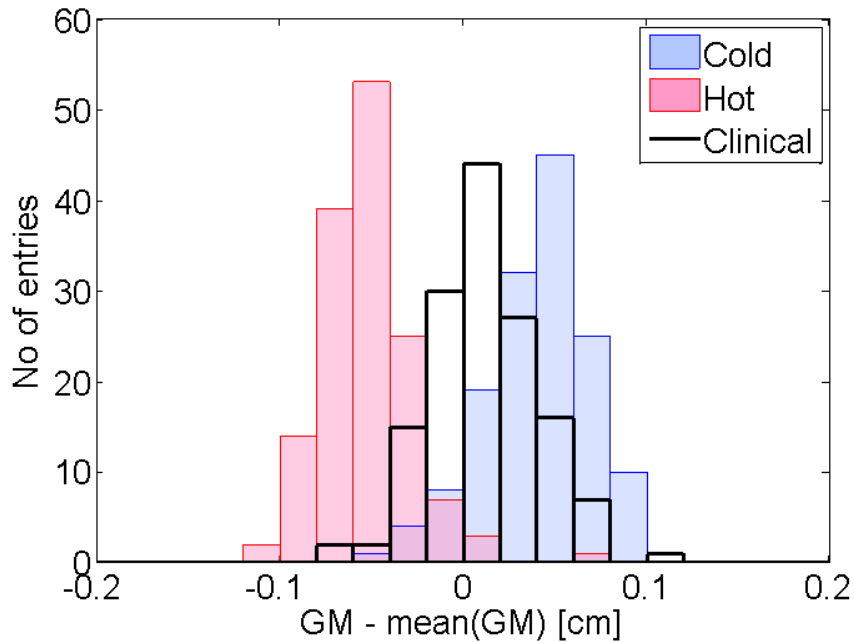
KBP SRS in Eclipse

- Use original plan's arc arrangement
- DVH predictions feed two different optimization routines, coded as patient-specific templates
 - HOT: for brain metastases, reduces penalty for hot spots and prioritizes GM
 - COLD: for use in benign disease and retreatments where hot spot is clinically important
- All plans are normalized to the same PTV coverage ($V100\%=98\%$ typically)

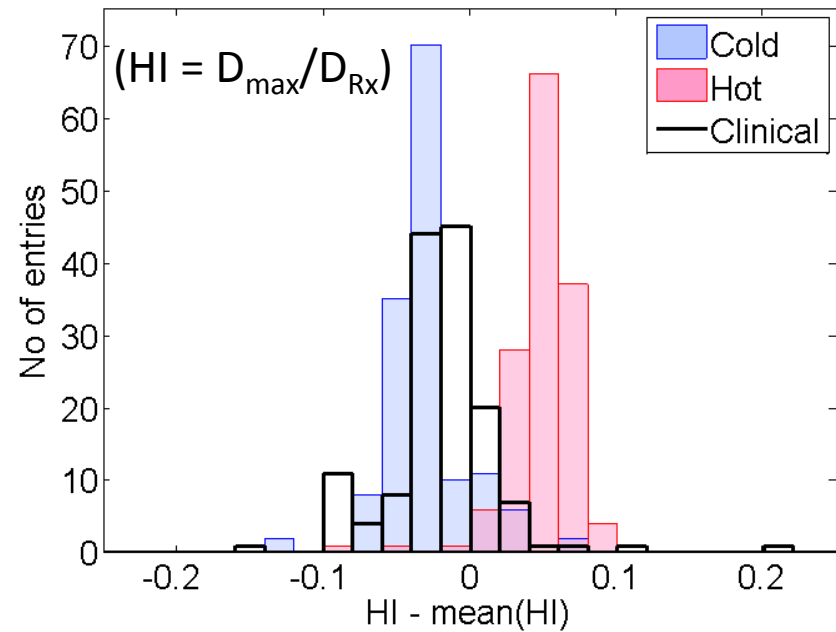


Tuning up autoplanning routines

GM: Hot < Clinical <= Cold



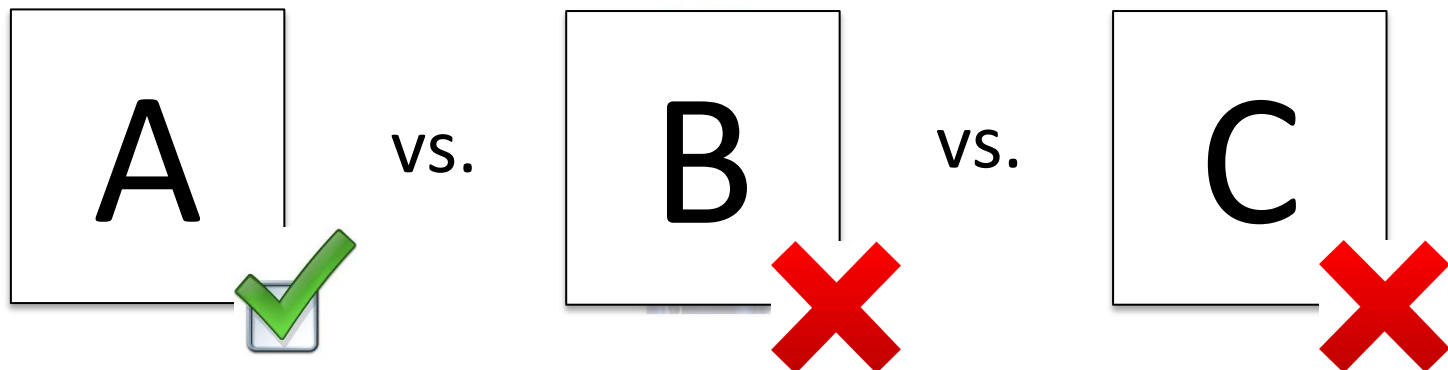
HI: Cold <= Clinical < Hot



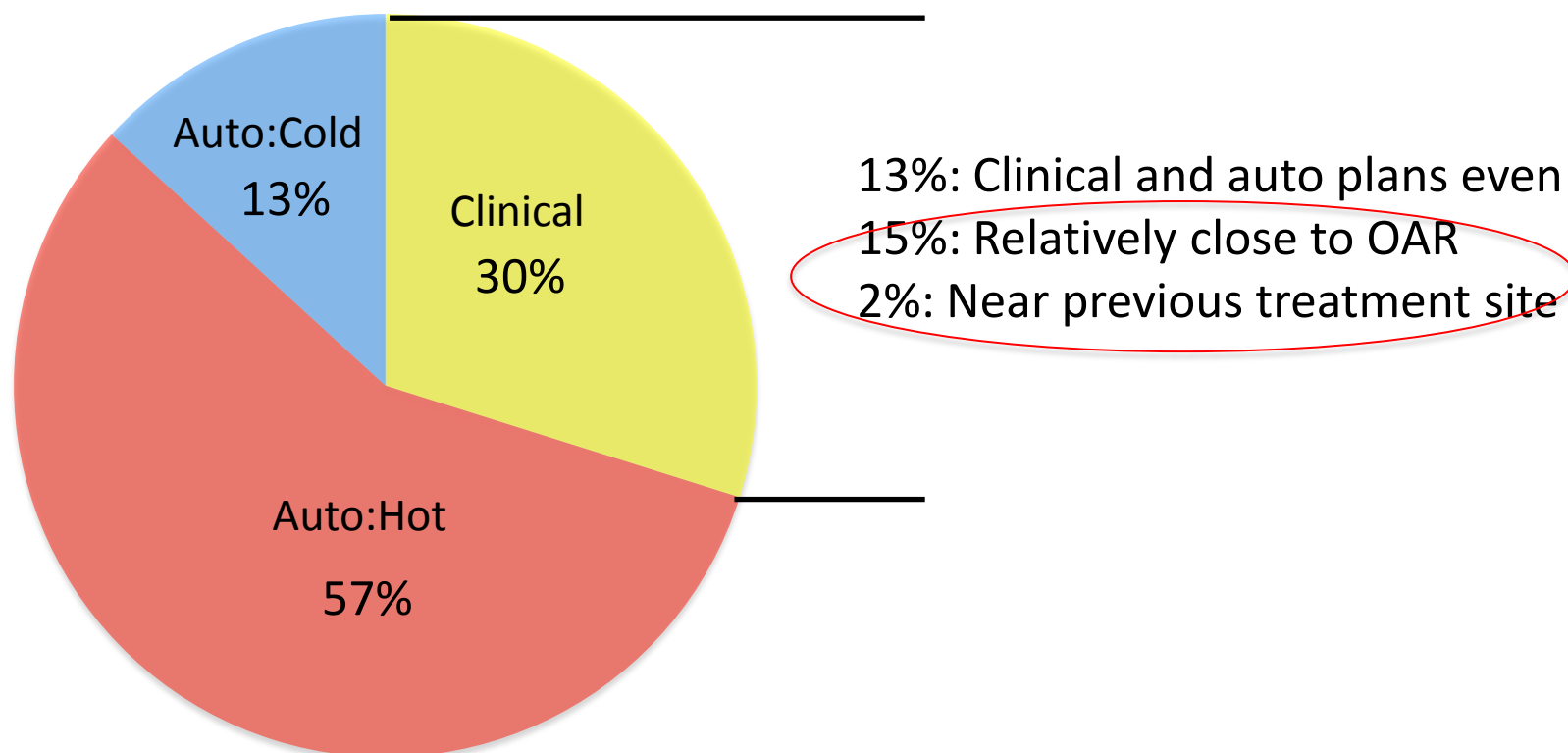
Single-blind study of autoplans vs. manual plans

Study schema:

1. Automatically replan 200+ SRS cases with *HOT* and *COLD* routines
2. Clinically approved plan and autoplans are de-identified (A, B, C randomly)
3. SRS physicians review plans with relevant clinical information and selects the preferred plan



Preliminary results



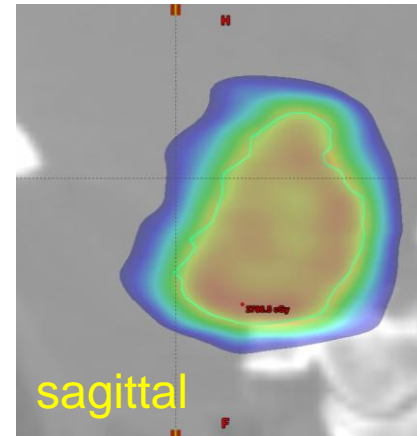
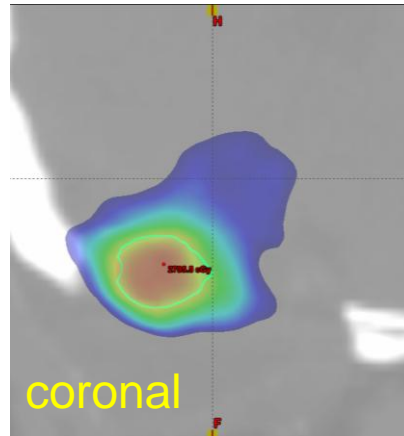
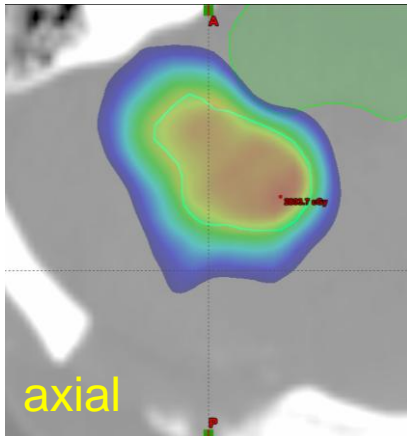
	Clinical	Auto: Hot	Auto: Cold	Total
Dr. Sanghvi	18 (26%)	43 (61%)	9 (13%)	70
Dr. Hattangadi	25 (34%)	39 (53%)	10 (14%)	74
TOTAL	43 (30%)	82 (57%)	19 (13%)	144



Preliminary results

- Autoplan sequences took ~15 minutes on average
- In the (17%) 24/144 cases where the manually-planned treatments were preferred
 - 21 plans were selected because of more aggressive OAR sparing (brainstem, cochlea, or optic nerve) at max dose level
 - 3 plans were selected because the manual plans better spared a nearby volume that received prior radiotherapy

clinical plan



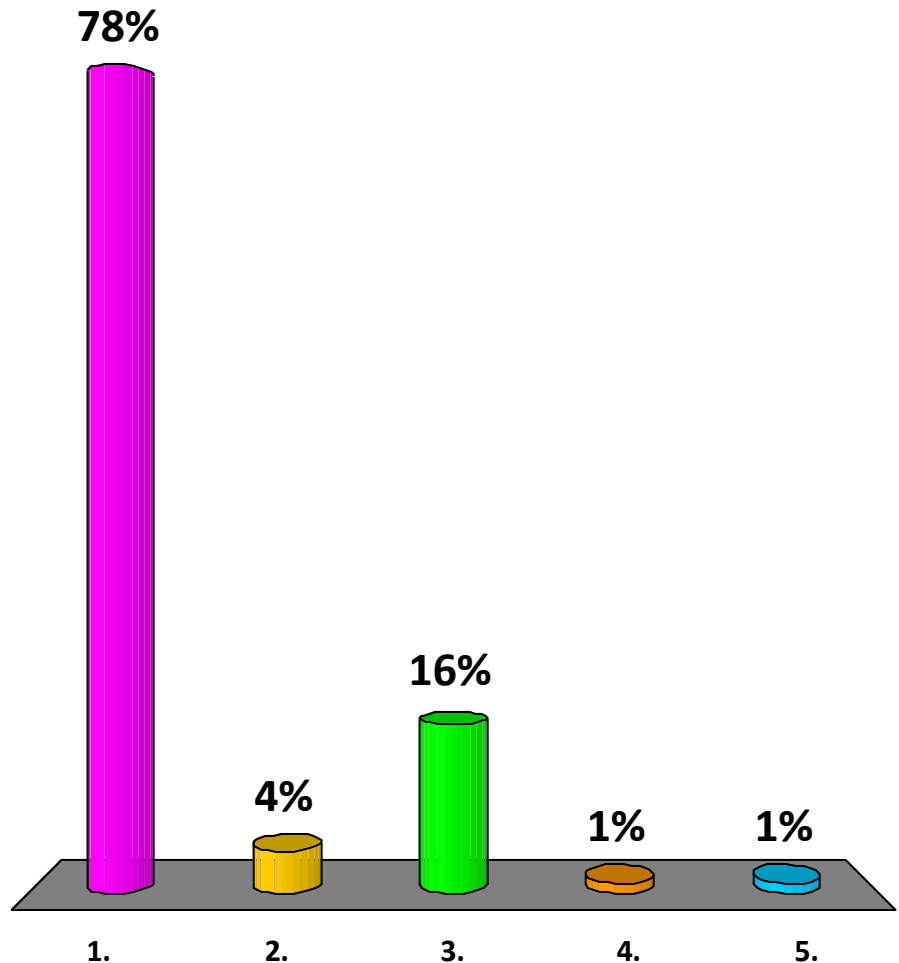
Summary of current KBP system in SRS

- Already have solution that yields superior or equivalent results for 83% of SRS cases
- Focusing on that remaining 17%...
 - Robust multi-met solution (forthcoming)
 - Robust neighboring OAR solution (underway)
 - Prior tx solution (underway)
 - Clinical “go live” after completion of blind study
- When possible, such a benchmarking study should be used before clinical implementation of automated planning



Knowledge-based planning in SRS:

1. can predict plan quality metrics and automate the planning process based on accurate dose-volume predictions
2. automatically loads standard planning templates for patients
3. guides the planning process by continually adjusting dose objectives during optimization
4. can only be used for inverse optimized planning
5. saves time but likely at the expense of plan quality



Correct Answer: 1

1. Can predict plan quality metrics and automate the planning process based on accurate dose-volume predictions

Knowledge-based prediction of plan quality metrics in intracranial stereotactic radiosurgery
S Shiraishi, J Tan, LA Olsen, KL Moore
Medical physics 42 (2), 908-917

Conclusion

- SRS and SBRT* are extremely well suited to knowledge-based techniques
- Knowledge-based quality metric prediction is useful for both quality control and planning automation
- Clinical KBP is still in its infancy, but in some form these techniques will be part of the treatment planning process
- KBP can also help inform clinical decision making (when to fractionate, benefits of different treatment techniques, e.g. 4π vs. static field vs. coplanar VMAT vs. protons)

** Abstracts at AAPM 2015:*

- *Foy et al SU-ET-97*
- *Snyder et al MO-F-CAMPUS-T-04*

