

A data-driven method to predict achievability of clinical objectives in IMRT

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A data-driven method to predict achievability of clinical objectives in IMRT

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Abstract:

When specifying a clinical objective for a target volume and normal organs/tissues in IMRT planning, the user may not be sure if the defined clinical objective could be achieved by the optimizer. To this end, we propose a novel method to predict the achievability of clinical objectives upfront before invoking the optimization. A new metric called “Geometric Complexity (GC)” is used to estimate the achievability of clinical objectives. Essentially GC is the measure of the number of “unmodulated” beamlets or rays that intersect the Region-of-interest (ROI) and the target volume. We first compute the geometric complexity ratio (GCratio) between the GC of a ROI in a reference plan and the GC of the same ROI in a given plan. The GCratio of a ROI indicates the relative geometric complexity of the ROI as compared to the same ROI in the reference plan. Hence GCratio can be used to predict if a defined clinical objective associated with the ROI can be met by the optimizer for a given case. We have evaluated the proposed method on six Head and Neck cases using Pinnacle3 (version 9.10.0) Treatment Planning System (TPS). Out of total of 42 clinical objectives from six cases accounted in the study, 37 were in agreement with the prediction, which implies an agreement of about 88% between predicted and obtained results. The results indicate the feasibility of using the proposed method in head and neck cases for predicting the achievability of clinical objectives.

A data-driven method to predict achievability of clinical objectives in 25 IMRT

INTRODUCTION

Intensity modulated radiation therapy (IMRT) has grown as an effective way of producing a conformal dose to tumor, while effectively sparing the surrounding normal tissues and organs. The main goal of optimization in IMRT is to find parameters that will yield the best possible treatment plans under given clinical and technical conditions. In the current practice of IMRT planning, one of the common approaches is that a user would first define a set of initial clinical objectives for target volumes (E.g. Minimum PTV Dose of 6300 cGy) and perform the optimization. After achieving the target volume objectives, user would start including Organs-at-risk (OAR) objectives (E.g. spinal cord Maximum Dose of 4500 cGy) one by one and perform several re-optimizations. From here on, the process gets more complicated and the user is required to carefully tweak the objective parameters in order to strike a balance between target coverage, target homogeneity and OAR sparing [1]. This generally involves several optimizations to arrive at an optimal objective setting. Often, for difficult treatment plans, the user may not be sure if the defined clinical objective could be achieved by the optimizer. In many situations, the defined clinical objective goes unachieved by the optimizer. But this realization happens only after performing one or many optimizations. This leads to several backtracking steps and hence the process becomes ineffective and time consuming. Moreover, due to these difficulties, the resulting plan quality becomes highly dependent on the ability of the treatment planners to meet the specified objectives [2].

Researchers have investigated algorithmic methods to make IMRT planning more efficient and less dependent on the expertise of the treatment planners [3-9]. Some
 50 researchers have explored data-driven approaches as well with the same research intent [10-13]. Such data-driven methods have been proven successful for predicting achievable dose levels for clinical objectives before invoking the actual optimization.

We propose a novel data-driven method to improve the efficiency of IMRT optimization
 55 process. Our method allows predicting the achievability of clinical objectives upfront before invoking the optimization, thereby eliminating the need for several trials and errors in fine tuning the objective parameters. This study evaluates the feasibility of the proposed method on Head and Neck cases.

60 MATERIALS AND METHODS:

Geometric Complexity (GC):

Geometric complexity (GC) is the measure of the number of “unmodulated” beamlets or rays that intersect the Region-of-interest (ROI) and the composite target volume (i.e. the volume containing all target volumes). The GC computed for a ROI is given by

65

$$GC = \frac{n + 1}{N_T/V} = \frac{n + 1}{N} \quad (1)$$

Where, n is the number of beamlets that pass through the ROI and target volume for a
 70 given plan, V is the volume of the target volume (e.g. the volume of the tumor to be

irradiated), N_T is the total number of beamlets passing through the target volume, and $N = N_T/V$ is the number of available beamlets per unit volume of the target.

Figures 1 diagrammatically illustrates two examples of a target volume and an OAR being irradiated by a single radiation beam. In this example, the number N_T of available beamlets is four. (This is diagrammatic: in practical IMRT cases N_T is typically on the order of hundreds or thousands). In the example of **Figure 1a**, all four available beamlets that intersect the target volume also intersect the OAR, so that $n = 4$. This results in a high value for the GC metric of Equation (1), indicating high geometrical complexity and a low likelihood of achieving the objectives associated with the OAR after optimization.

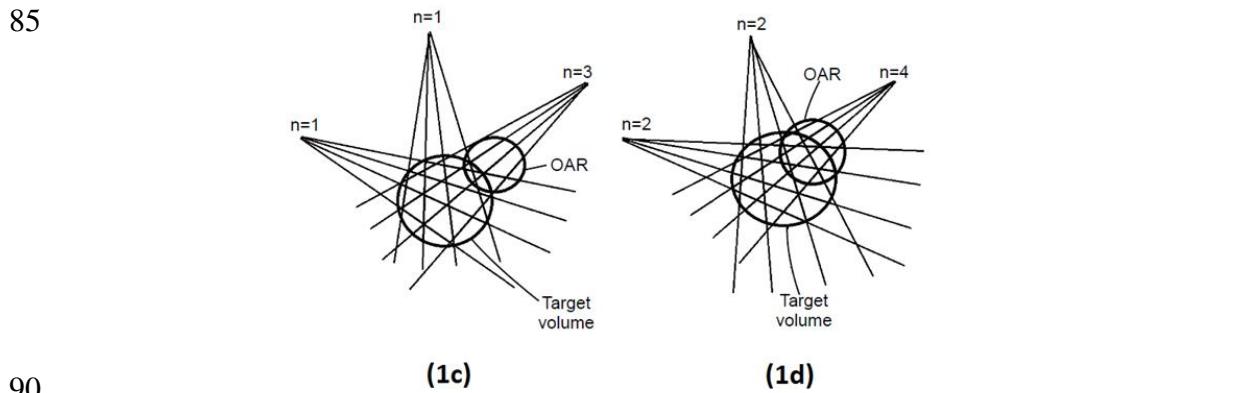
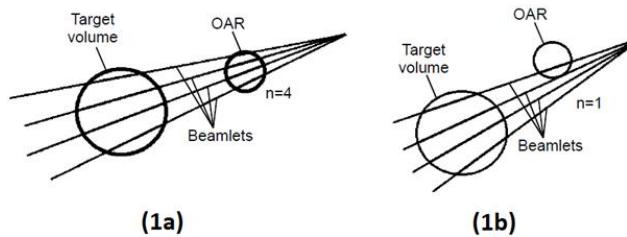


Figure 1: Diagrammatically illustrated examples (1a-1d) of geometric complexity estimation based on the position of OAR with respect to target volume (1a-1b) and the level of target-OAR overlap (1c-1d)

95 By comparison, **Figure 1b** shows a different IMRT geometry in which only one of the beamlets that intersect the target volume also intersects the OAR, leading to $n = 1$. This results in a low value for the GC metric of Equation (1), indicating low geometrical complexity and a high likelihood of achieving the objectives associated with the OAR after optimization. (For example, that single beamlet intersecting the OAR could be
100 shuttered off by the optimized MLC configuration).

In another example illustrated in **Figures 1c and 1d**, each radiation beam is modulated into four beamlets, so that for the three radiation beams $N_T = 4 \times 3 = 12$. In the example of **Figure 1c**, one beamlet of the leftmost radiation beam also intersects the OAR, one beamlet of the middle radiation beam also intersects the OAR, and three beamlets of the rightmost radiation beam also intersect the OAR. Thus,
105 $n = 1 + 1 + 3 = 5$, and from Equation (1) the GC metric is $\left(\frac{6}{12}\right)V$. In the example of **Figure 1d**, two beamlets of the leftmost radiation beam also intersects the OAR, two beamlets of the middle radiation beam also intersects the OAR, and all four beamlets of
110 the rightmost radiation beam also intersect the OAR. Thus, $n = 2 + 2 + 4 = 8$, and from Equation (1) the GC metric is $\left(\frac{9}{12}\right)V$. It follows that the IMRT geometry of **Figure 1c** is more likely to yield an achievable IMRT plan optimization (because of its lower GC) as compared with the IMRT geometry of **Figure 1d** (because of its higher GC).

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Geometric complexity ratio

We first create a reference plan in which all the clinical objectives have been met through the optimization process. The GC associated with each segmented clinical structure or
 120 Region of Interest (ROI) is computed for the reference plan. The GC obtained per ROI for the reference plan is considered as the base or reference value i.e. GC_{ref} .

$$125 \quad GC_{ref} = \frac{n_{ref}+1}{N_{T,ref}/V_{ref}} = \frac{n_{ref}+1}{N_{ref}} \quad (2)$$

Where, n_{ref} is the number of beamlets that pass through the ROI and target volume for a reference plan, V_{ref} is the volume of the target volume (e.g. the volume of the tumor to be irradiated), $N_{T,ref}$ is the total number of beamlets passing through the target volume in
 130 the reference plan and $N_{ref} = N_{T,ref}/V_{ref}$ is the number of available beamlets per unit volume of the target volume in the reference plan.

Dividing equation 1 by equation 2, we get geometric complexity ratio (GC_{ratio}) of a ROI as shown below.

$$135 \quad GC_{ratio} = \frac{\frac{n+1}{N}}{\frac{n_{ref}+1}{N_{ref}}} \quad (3)$$

140 It is to be note that if the ROI is a target volume, then ‘n’ refers to the number of beamlets that intersect target volume and all other ROIs getting included in the optimization.

145 Predicting the achievability of clinical objectives

Essentially our approach involves the comparison of geometric complexity of a given plan to that of a reference plan to estimate the achievability of clinical objectives. The GC_{ratio} of a ROI indicates the relative geometric complexity of the ROI as compared to the same ROI in the reference plan. Achievability of a clinical objective associated with the ROI can be perceived as the inverse of GC_{ratio} of the ROI. Hence GC_{ratio} can be used to predict if a defined clinical objective associated with the ROI can be met by the optimizer for a given case. Basically a higher GC_{ratio} indicates a lesser likelihood for the optimizer to achieve the clinical objective defined for a given ROI. Similarly, a lower GC_{ratio} indicates a higher likelihood for the optimizer to achieve the clinical objective defined for the given ROI.

In the examples illustrated in **Figures 1a to 1d**, assume that 1a and 1c are reference plans and 1b and 1d are given or current plans. Also assume that the OAR objectives in the reference plans 1a and 1c have been met by the optimizer. The estimated GC_{ratio} for OAR in plan 1b with respect to plan 1a is 0.25, which implies that if a planner defines same objective setting for the OAR in current plan, it is highly likely that the objective will be met by the optimizer. Similarly, the estimated GC_{ratio} for OAR in plan 1d with respect to plan 1c is 1.5, which implies that if a planners defines same objective setting for the OAR in current plan, it is highly likely that the objective will not be met by the optimizer.

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However, these numerical values may not be intuitive for the user. Hence, we have interpreted the achievability of the objectives (i.e. likelihood of achieving an objective)

based on the GC_{ratio} value using different schemes as given in **Table I**. The basic assumption when predicting the achievability of a given objective is that the objective parameters (i.e. dose, volume and weight) associated with that objective is same as that used in the reference plan. It is to be noted that the clinical objectives in the reference plan(s) must have been achieved in order to make accurate predictions. **Figure 2** shows the flowchart of the proposed method.

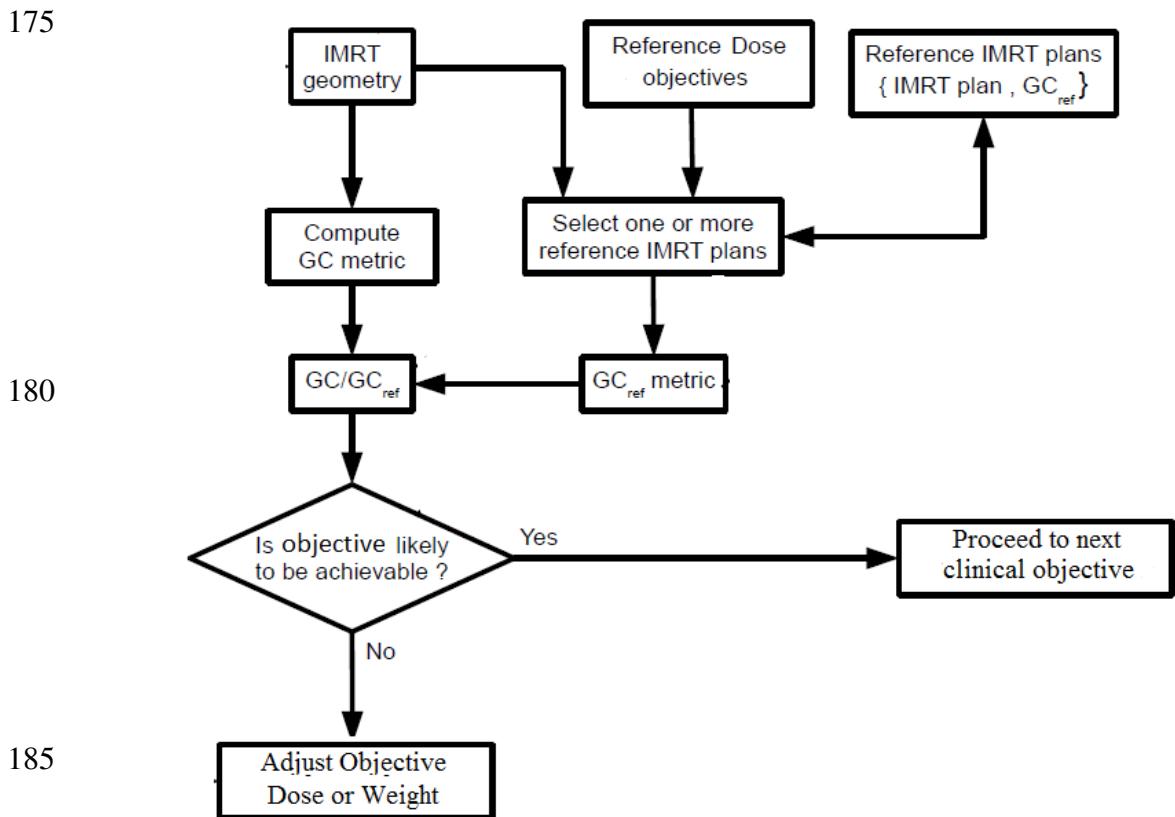


Figure 2: Flowchart of the proposed method for predicting the achievability of clinical objectives

Study methodology

Pinnacle³ (version 9.10.0) Treatment Planning System (TPS) was used for the whole study for contouring, dose computation and optimization. The optimization was done using DMPO module in which a minimum segment MU of 4 and minimum segment area
195 of 4 cm² was used throughout the study. Varian Tx HDMLC machine was used for simulation purposes. The dose grid resolution was set to 0.3 cm in X, Y and Z planes for dose computation.

First, a reference head and neck case with 7 beams was selected. For the reference plan, a
200 manual planning in a conventional manner was performed by adding clinical objectives one-by-one (in the order of target objectives, cord and brainstem objectives and parotid objectives) and tweaking the objective parameters over many optimizations (as described in the introduction) and reached an acceptable plan, wherein all defined objectives were met. The GC_{ref} values for each ROI included in the reference plan was computed. Six
205 head and neck cases of varying complexity was selected for the study. The beams angles were manually selected for all the cases. The GC values were computed for target volumes, spinal cord, parotids and brainstem in these plans. Our objective is to predict the achievability of clinical objectives associated with these ROIs by making use of the reference GC_{ref} values obtained in the reference plan. **Equation 3** was used to compute
210 GC_{ratio} for all ROIs (including target volumes). The schemes listed in **Table I** were used to interpret the GC_{ratio} values. Different schemes were used for interpretation because at the time of study we were not sure which one will be suitable with respect to the selected reference plan. Our intention is to find an interpretation scheme that gives best agreement

Table I: Interpretation of the achievability of the objectives (i.e. likelihood of achieving an objective) based on the GC_{ratio} value using different schemes. Here A refers to Achievable, PA refers to possibly achievable, PNA refers to possibly not achievable and NA refers to not achievable.

	Scheme #	Scheme
220		$GC_{ratio} \leq 1 = A$
	1	$1 < GC_{ratio} \leq 1.1 = PA$
		$1.1 < GC_{ratio} \leq 1.2 = PNA$
		$GC_{ratio} > 1.2 = NA$
225		$GC_{ratio} \leq 1.15 = A$
	2	$1.15 < GC_{ratio} \leq 1.2 = PA$
		$1.2 < GC_{ratio} \leq 1.25 = PNA$
		$GC_{ratio} > 1.25 = NA$
230		$GC_{ratio} \leq 1.2 = A$
	3	$1.2 < GC_{ratio} \leq 1.3 = PA$
		$1.3 < GC_{ratio} \leq 1.35 = PNA$
		$GC_{ratio} > 1.35 = NA$
235		$GC_{ratio} \leq 1.2 = A$
	4	$1.2 < GC_{ratio} \leq 1.35 = PA$
		$1.35 < GC_{ratio} \leq 1.45 = PNA$
		$GC_{ratio} > 1.45 = NA$

between predicted and obtained results with respect to the selected reference plan.

Basically the achievability is interpreted as “Achievable (A)”, “Possibly Achievable

(PA)", "Possibly Not Achievable (PNA)" and "Not Achievable (NA)" based on GC_{ratio} values. After making the predictions with four different schemes, we included the clinical objectives one-by-one in the same order they were added in reference plan and performed optimization (without changing objective parameters) to get the actual results for each clinical objective. The same was repeated for all six cases.

245 **RESULTS:**

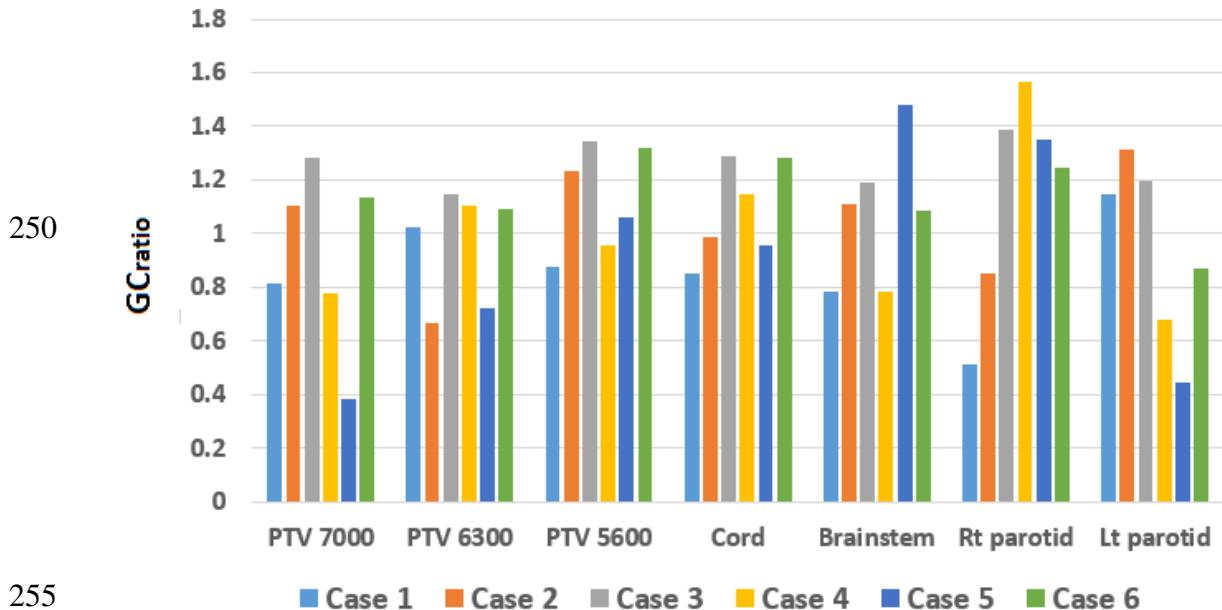


Figure 3: The variations in the geometric complexity for different ROIs in the six cases included in the study.

260 **Table II:** Comparison of GC_{ratio} values, estimated achievability using different schemes and obtained (actual) results after optimization for six head and neck cases. Here 'Ach' refers to Achieved and 'Not Ach' refers to Not Achieved.

Table IIa: Head & Neck – Reference case

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	1	--	--	--	--	Ach
PTV 6300	Target Dose	6300	95	1	--	--	--	--	Ach
PTV 5600	Target Dose	5600	95	1	--	--	--	--	Ach
Cord	Max Dose	4500	0	1	--	--	--	--	Ach
B.Stem	Max Dose	5400	0	1	--	--	--	--	Ach
Parotid Rt	Mean Dose	2600	--	1	--	--	--	--	Ach
Parotid Lt	Mean Dose	2600	--	1	--	--	--	--	Ach

Table IIb: Head & Neck –Case 1

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	1.104	PA	A	A	A	Ach
PTV 6300	Target Dose	6300	95	0.667	A	A	A	A	Ach
PTV 5600	Target Dose	5600	95	1.231	NA	PNA	A	A	Ach
Cord	Max Dose	4500	0	1.112	PA	A	A	A	Ach
B.Stem	Max Dose	5400	0	0.989	A	A	A	A	Ach
Parotid Rt	Mean Dose	2600	--	0.854	A	A	A	A	Ach
Parotid Lt	Mean Dose	2600	--	1.313	NA	NA	PNA	PNA	Not Ach

265 **Table IIc:** Head & Neck – Case 2

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	0.811	A	A	A	A	Ach
PTV 6300	Target Dose	6300	95	1.024	PA	A	A	A	Ach
PTV 5600	Target Dose	5600	95	0.877	A	A	A	A	Ach
Cord	Max Dose	4500	0	0.852	A	A	A	A	Ach
B.Stem	Max Dose	5400	0	0.781	A	A	A	A	Ach
Parotid Rt	Mean Dose	2600	--	0.514	A	A	A	A	Ach
Parotid Lt	Mean Dose	2600	--	1.144	PA	A	A	A	Ach

Table IIId: Head & Neck – Case 3

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	1.285	NA	NA	PA	PA	Not Ach
PTV 6300	Target Dose	6300	95	1.145	PA	A	A	A	Not Ach
PTV 5600	Target Dose	5600	95	1.341	NA	NA	PNA	PNA	Not Ach
Cord	Max Dose	4500	0	1.291	NA	NA	PNA	PNA	Not Ach
B.Stem	Max Dose	5400	0	1.188	NA	PA	A	A	Not Ach
Parotid Rt	Mean Dose	2600	--	1.385	NA	NA	NA	PNA	Not Ach
Parotid Lt	Mean Dose	2600	--	1.193	PNA	PA	A	A	Ach

Table IIe: Head & Neck – Case 4

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	0.775	A	A	A	A	Ach
PTV 6300	Target Dose	6300	95	1.105	PA	A	A	A	Ach
PTV 5600	Target Dose	5600	95	0.953	A	A	A	A	Ach
Cord	Max Dose	4500	0	1.145	PA	A	A	A	Ach
B.Stem	Max Dose	5400	0	0.782	A	A	A	A	Ach
Parotid Rt	Mean Dose	2600	--	1.564	NA	NA	NA	NA	Ach
Parotid Lt	Mean Dose	2600	--	0.678	A	A	A	A	Ach

270 **Table IIIf:** Head & Neck – Case 5

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	0.385	A	A	A	A	Ach
PTV 6300	Target Dose	6300	95	0.724	A	A	A	A	Ach
PTV 5600	Target Dose	5600	95	1.061	PA	A	A	NA	Ach
Cord	Max Dose	4500	0	0.958	A	A	A	A	Ach
B.Stem	Max Dose	5400	0	1.481	NA	NA	NA	NA	Not Ach
Parotid Rt	Mean Dose	2600	--	1.349	NA	NA	PNA	PNA	Not Ach
Parotid Lt	Mean Dose	2600	--	0.446	A	A	A	A	Ach

Table IIg: Head & Neck – Case 6

Organ	Objective	Dose	Volume	GC _{ratio}	Scheme	Scheme	Scheme	Scheme	Obtained
	Type	(cGy)	(%)		1	2	3	4	Result
PTV 7000	Target Dose	7000	95	1.135	PA	A	A	A	Ach
PTV 6300	Target Dose	6300	95	1.094	PA	A	A	A	Ach
PTV 5600	Target Dose	5600	95	1.317	NA	NA	PNA	PNA	Not Ach
Cord	Max Dose	4500	0	1.282	NA	NA	PA	PA	Not Ach
B.Stem	Max Dose	5400	0	1.088	PA	A	A	A	Ach
Parotid Rt	Mean Dose	2600	--	1.244	NA	PNA	PA	PA	Ach
Parotid Lt	Mean Dose	2600	--	0.872	A	A	A	A	Ach

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Table II gives the comparison between predicted and obtained results for the four interpretation schemes. **Figure 3** shows the geometric complexity ratio for different ROIs in the six cases included in the study. The cumulative GC_{ratio} values (i.e. sum of all GC_{ratio} values greater than 1) for Case 1 to Case 6 are 4.8, 2.2, 8.8, 4.5, 3.9, and 7.2 respectively.

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We analyzed the results using Pearson correlation method. In order to perform correlation tests, a value of 1 was assigned to ‘A’, 0.5 to ‘PA’, 0.5 to ‘PNA’ and 0 to ‘NA’. Also a value of 1 was assigned to the obtained result if the objective was met and a value of 0 if the objective was not met. Though the prediction using all four schemes have a positive correlation with the obtained results, scheme # 2 has the maximum positive correlation

285 (correlation was 0.81 with r^2 value of 0.66 and $p < 0.005$) with the obtained results. It is
evident from **Table II** that, out of the total of 42 clinical objectives from all six cases, 37
were in agreement with the prediction for scheme # 2 which implies an agreement of
about 88% between predicted and obtained results. **Table III** lists the percentage
correlation and strength of correlation between predicted and obtained results for all
290 interpretation schemes. Please note that we have previously presented these results in
AAPM 2016 annual conference [14] and also published as Patent [15].

DISCUSSION

The results indicate that it is feasible to use the proposed method to predict the
295 achievability of clinical objectives before invoking optimization. Except for Case 3, the
prediction was accurate for other cases. For Case 3, the cumulative GC_{ratio} score is
significantly higher, which indicates that the geometry of Case 3 is considerably different
from that of reference plan. Basically a poor selection of reference plan can lead to
inaccurate predictions. Hence, it is recommended to choose an appropriate reference plan
300 with respect to a given case to make accurate predictions. In practice, one can run a
search on a database of previously optimized plans with estimated GC_{ratio} values for
different ROIs and preferably select a plan for which the cumulative score is less than or
equal to the number of ROIs (including target volumes) included in the optimization. The
plan selected hereby can be used as a reference plan for making the predictions on
305 clinical objectives. This process is illustrated in **Figure 4**.

Table III: The percentage agreement and strength of correlation between predicted and obtained results for different interpretation schemes.

Scheme #	Scheme	% agreement	Correlation	r^2
<hr/>				
1	$GC_{ratio} \leq 1 = A$			
	$1 < GC_{ratio} \leq 1.1 = PA$	86%	0.63	0.40
	$1.1 < GC_{ratio} \leq 1.2 = PNA$			
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2	$GC_{ratio} \leq 1.15 = A$			
	$1.15 < GC_{ratio} \leq 1.2 = PA$	88%	0.81	0.66
	$1.2 < GC_{ratio} \leq 1.25 = PNA$			
<hr/>				
3	$GC_{ratio} \leq 1.2 = A$			
	$1.2 < GC_{ratio} \leq 1.3 = PA$			
	$1.3 < GC_{ratio} \leq 1.35 = PNA$	86%	0.62	0.39
<hr/>				
4	$GC_{ratio} \leq 1.2 = A$			
	$1.2 < GC_{ratio} \leq 1.35 = PA$			
	$1.35 < GC_{ratio} \leq 1.45 = PNA$	86%	0.74	0.55
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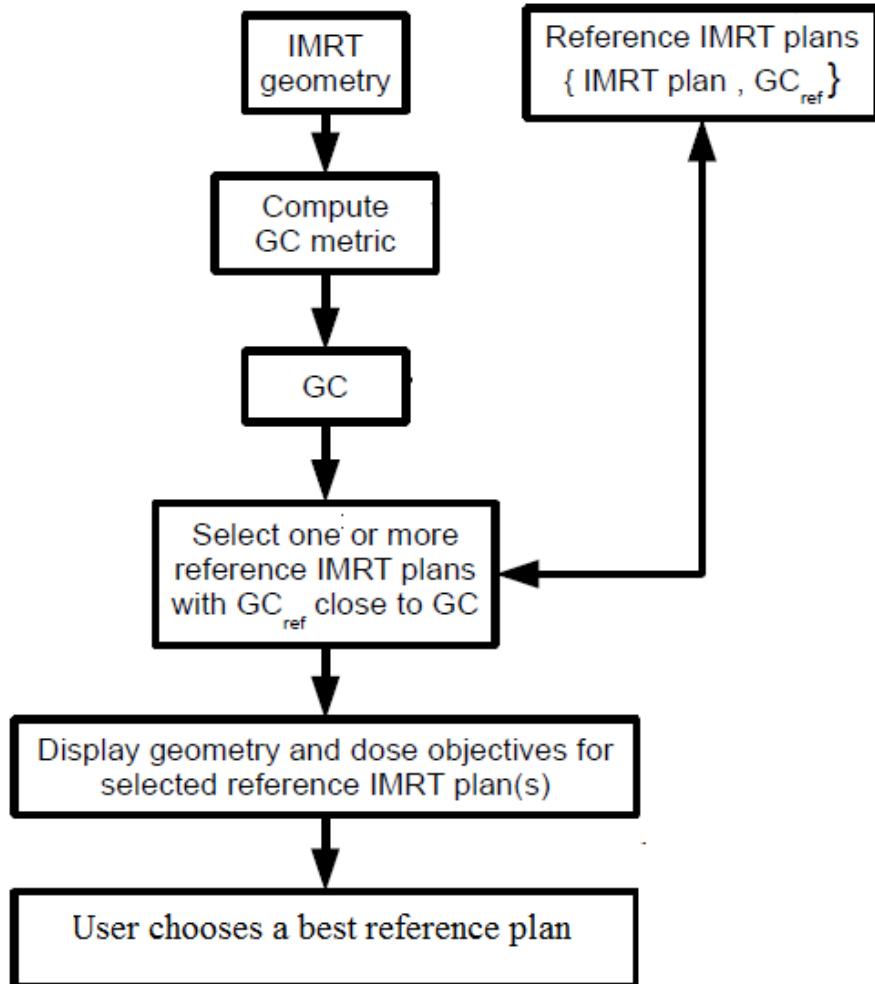


Figure 4: An illustration of the process of selecting a good reference plan from a database containing several plans

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Also it is to be noted that the scheme # 2, which was found to be suitable for interpretation, might not be suitable if a different reference plan is used for prediction.
For instance, if Case 3 is used as a reference plan, scheme # 2 results in an agreement of
340 about 73% with the obtained results, whereas scheme # 1 results in an agreement of about
83%. Hence it is important to select an appropriate scheme for interpretation based on the
reference plan to make accurate predictions. To select an appropriate scheme, the average
GC_{ratio} value of a given reference plan can be compared against that of a master reference
plan, whose interpretation scheme has been established. Based on the outcome of this
345 comparison, an appropriate interpretation scheme can be picked from a library of
schemes for the given reference plan. A unique advantage of the proposed method is that
the beam angle configuration between a reference plan and a given plan need not be the
same to make predictions. This is because, the computation of geometric complexity
directly accounts for the beam configurations.

350 Typically the computation of GC_{ratio} takes a few seconds even for the cases involving
numerous ROIs. Since, it takes several optimization loops before reaching an acceptable
plan in complicated cases, knowing the probable results before invoking optimization can
be of very useful. In several different ways a user can respond after knowing the
355 achievability of objectives. For instance, the clinical objectives with lower achievability
can be initialized with higher importance weights to ensure they are most likely achieved
after optimization, thereby avoiding backtracking steps. This will save a lot of time spent
in tweaking the parameters and also reduce the inter-user variability in plan quality.
Recently, some researchers have proposed a method for a fully automated solutions for

360 IMRT optimization [14, 15], which has been implemented in commercial TPS [16-19].

Our approach of predicting the achievability of clinical objectives can also be used in conjunction with the existing automatic approach [16-19] for initializing the objective parameter settings before starting the optimization loops, which can reduce the overall time spent during optimization.

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CONCLUSION

We have proposed a method to predict the achievability of clinical objectives in IMRT planning. Basically our method allows benchmarking a given plan against a high quality reference plan through the use of geometric complexity metric. The study demonstrates 370 the feasibility of using the proposed method for predicting the achievability of clinical objectives with reasonable accuracy. Though we have demonstrated the feasibility for head and neck anatomy, this method should be applicable for other anatomic sites such as brain, thorax, abdomen and pelvis. Also, the proposed method can be directly applied to VMAT scenarios.

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CONFLICT OF INTEREST

NO CONFLICT OF INTEREST

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