



The dosimetric comparison using anisotropic analytical algorithm (AAA) and Acuros XB algorithms in the target overlapping with air cavity of pituitary carcinoma

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Background: This study was to investigate the dose variation between anisotropic analytical algorithm (AAA) and Acuros XB (AXB) calculation algorithms in heterogeneous inside planning target volume (PTV) of pituitary carcinoma.

Methods: AAA and AXB algorithms were used for dose calculation by Eclipse treatment planning system in Varian TrueBeam linear accelerator machine with 120 MLC. Twelve cases of pituitary carcinoma were planned by intensity-modulated radiotherapy (IMRT) and volumetric modulated arc therapy (VMAT) techniques. The target overlapped with the air cavity. IMRT plans were done by 9 static fields and VMAT plans were done by 2–3 arc rotation fields. The PTV was prescribed the dose of 50.4 Gy. The dosimetry was analyzed by dose to 95% and 98% of volume of PTV, maximum dose, dose coverage, and conformity index (CI) of PTV. The normal organ was analyzed by mean dose (Dmean) and maximum dose (Dmax).

Results: The dose coverage of PTV using AXB calculation was less than using AAA calculation in the air cavity region for both IMRT and VMAT. The dose coverage of PTV was any statistical significantly difference between AAA and AXB algorithms. For the normal organs, the mean and maximum dose agreed well for AAA and AXB algorithms.

Conclusions: AXB can calculate the dose more accurate than AAA for dose calculation in inhomogeneous region. AXB algorithms should be selected for dose calculation in clinical.

Keywords: Anisotropic analytical algorithm (AAA); Acuros XB (AXB); treatment planning, inhomogeneity, air cavity

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Introduction

In radiotherapy, the prescription dose delivered to the patient must be accurate with $\pm 5\%$ of overall (1). The treatment planning is an important system to reduce the variation of the dose to patient in the dose calculation process which depends on the dose calculation algorithms.

Eclipse treatment planning (Varian Medical Systems, Palo

Alto, CA, USA) provides two mainly algorithms to calculate dose which the one is anisotropic analytical algorithm (AAA) and the other one is Acuros XB (AXB) algorithm. The AAA which has been implemented in Eclipse TPS is one of a 3D pencil beam convolution-superposition algorithm. AAA model configuration is based on basic physical parameters which are determined by Monte-Carlo. This algorithm

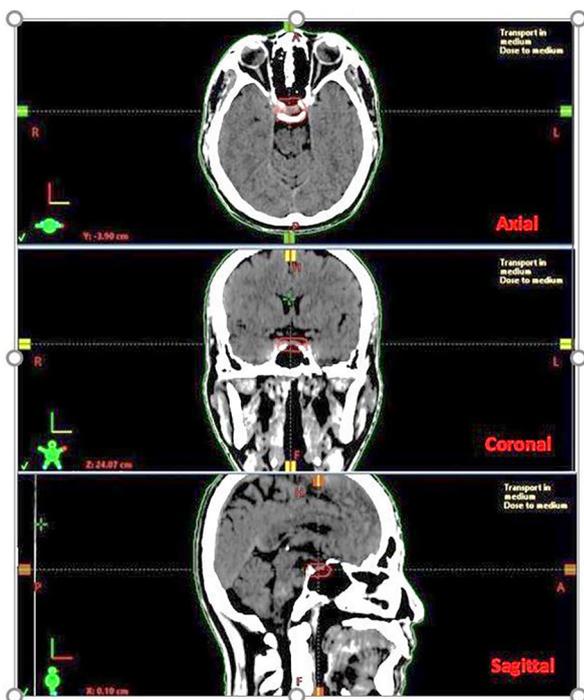


Figure 1 The PTV volume inside the air cavity shown in the red line for axial, coronal and sagittal planes. PTV, planning target volume.

model calculates tissue inhomogeneities anisotropically by using 13 lateral photon scatter kernels. Superposition of the doses determined from photon and electron convolutions gives the final dose distribution. In many cases, the overestimate dose between air and water interface was reported for AAA (2-6). On the other hand, the AXB applies the Linear Boltzmann Transport Equation (LBTE) Solvers in dose calculation modelling method which describes the macroscopic behavior of radiation particles as they travel through and interact with the material (7). Thus, AXB for dose estimation inside and around the inhomogeneity region agrees well with the Monte Carlo (MC) calculation method. AAA is an overestimate dose in the inhomogeneity region compared with MC (4). Moreover, some studies have been published the dose comparison between AAA and AXB, the results shown the dose calculation of AXB was more accurate dose prediction than AAA in inhomogeneity phantom (4,8-11). The aim of AXB is to approve for accurate modeling of the absorb dose in medium.

The dose comparison between the two algorithms has been studied and concerned in past few years. Many publications have evaluated the dose difference of AXB in

many regions of carcinoma with AAA calculation. They found the highest dose variation in heterogeneity medium interface between air and tissue. They considered the dose difference impacted in the homogenous target volume only (9,10). Although Zhang *et al.* (12) published the dose different of AAA and AXB in the lung tumor which it contains low-density tissue, the whole target volume is still homogeneous. The dose coverage of target volume is no statistically significant difference between AAA and AXB. In addition, the effect of dose for heterogeneity as lung and air is difference. From Kim *et al.* (13) studied the dose comparison in heterogeneity phantom slab between AAA and AXB of about 2.8% and 20% in lung tissue and in air, respectively. AAA calculation is more overestimate dose in air than in lung too much that the results agreed with Fogliata *et al.* (14). So, in the clinical, the cases which contain the air cavity in the treatment field should be concerned the dose in air inhomogeneity They found the dose different between AAA and AXB enhancing in light lung and air material than in normal lung tissue for inhomogeneity phantom.

This study investigated the dose difference impacted in the heterogenous target volume by IMRT and VMAT treatment techniques in a pituitary carcinoma region. When the gross target volume (GTV) is expanded to the planning target volume (PTV), it is commonly overlapping with the nasal air cavity. The PTV has filled in some part by air cavity inside the volume. From the previous studies (4,8-11), the dose difference was more impact in near interface of air and tissue. The dose should be concerned in heterogeneous tissue. Especially, the tumor has filled with the air cavity as shown in *Figure 1*. It affects with the dose coverage of the target when the plan is evaluated.

Therefore, the objective of this study is to investigate the dose variation between AAA and AXB calculation algorithms in heterogenous inside the PTV of pituitary carcinoma cases. PTV has the air cavity filled inside which effects with the dose distribution and dose coverage in plan. The dose differences between two algorithms were also determined and discussed before using in clinic.

We present the following article in accordance with the MDAR reporting checklist (available at <http://dx.doi.org/10.21037/tro-20-57>).

Methods

Treatment planning algorithm

The Eclipse treatment planning systems was used to all plans

Table 1 The volumes of PTV and air cavity filled within PTV

Case No.	PTV volume (cm ³)	Air cavity volume (cm ³)	% air cavity inside PTV volume
1	25.20	3.24	12.86
2	4.21	1.02	24.23
3	14.04	1.99	14.17
4	34.55	3.10	8.97
5	35.49	4.01	11.30
6	20.69	3.75	18.12
7	18.00	1.20	6.67
8	20.45	2.01	9.83
9	54.23	5.57	10.27
10	6.36	0.32	5.03
11	16.81	0.64	3.81
12	22.22	2.64	11.88

PTV, planning target volume.

Table 2 The dose constraints criteria for PTV and normal organs optimization

Structures	Dose constraints (cGy)
PTV	D _{95%} >5,040
Brainstem	D _{max} <5,400
Optic chiasm	D _{max} <5,400
Optic nerves	D _{max} <5,400
Cochlear	D _{mean} <4,500

PTV, planning target volume.

version 15 for AAA and AXB. AAA and AXB were configured in the treatment planning system with the same beam data set. All plans employed the 6 MV photon beam of Varian Clinac TrueBeam machine which was attached with the 120 multileaf collimator (Varian Medical Systems, Palo Alto, CA, USA). The MLC width is 0.5 cm at the central 20 cm of field and 1 cm for the outer 20 cm. Both algorithms were used to calculate dose in 12 pituitary carcinoma cases for intensity-modulated radiotherapy (IMRT) and volumetric modulated arc therapy (VMAT) techniques.

Simulation

All patients took individually a thermo-plastic head

mask (Civco medical solution, Coralville, IA, USA) for immobilization which was attached to a base plate indexed bar on the treatment table. The reference points were marked on the mask. The patients were scanned by computed tomography (CT) simulator (Siemens Somatom definition AS 64-slice, Erlangen, Germany). The 1-mm slice thickness was scanned along the skull vertex to the clavicles. The images were imported to Eclipse treatment planning system. The magnetic resonance imaging (MRI) was registered with CT imaging for target volume delineation.

Target volume and organs at risk (OARs) delineation

The target volume and OARs of 12 cases were delineated by radiation oncologist. The GTV was expanded to the PTV overlapping with the nasal air cavity. The PTV had filled in some part by air cavity inside the volume. The air cavity has the Hounsfield unit (HU) of -1,000 HU. Target volumes and air cavity volume in each case are shown in *Table 1*. Brainstem, optic chiasm, optic nerves, and cochleares was contoured as the OARs. The dose constraints and dose prescription are shown in *Table 2* according to a radiation oncologist's guide for organ at risk in brain and their dose constraints (15). Conventional fractionation was applied for total dose of 50.4 Gy in 28 fractions, once a day, 5 days per week (16).

Treatment planning

All IMRT plans were provided using 6 MV photon beam of nine fields around patients. The gantry angle started from 200° to 160° with 40° intervals. The collimator rotation was 0° for all fields. The dose rate was set at 400 monitor unit (MU)/min. The smart MLC was selected as MLC sequence mode for leaves motion calculation after the optimization finished.

For VMAT technique, all plans were created with double Arcs for full rotation (360°) with 6 MV photon beam, which started from the counter clockwise arc in the first arc plan. The maximum dose rate was set to 600 MU/min and then it was controlled automatic by the optimization process. The same dose constraints for optimization with IMRT plans were set.

All plans were calculated by using AAA and AXB. The dose optimization used the photon optimizer (PO) algorithm version 15 in Eclipse treatment planning. Dose correction for tissue heterogeneity was applied. AXB

Table 3 The average dosimetric comparison of AAA and AXB for both IMRT and VMAT in PTV

PTV	IMRT (n=12)			VMAT (n=12)		
	AAA	AXB	P value	AAA	AXB	P value
D ₉₅ (Gy)	51.1±0.6	48.9±3.1	0.028	50.2±1.8	47.6±4.8	0.020
D ₉₈ (Gy)	50.6±0.7	46.3±3.7	0.002	49.4±1.9	46.6±5.3	0.001
D _{max} (Gy)	53.7±0.5	54.9±0.9	<0.001	53.9±1.0	54.7±1.0	<0.001
CI	0.98±0.03	0.92±0.06	0.004	0.95±0.05	0.88±0.09	0.002

AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy; PTV, planning target volume; CI, conformity index.

calculation utilized the same MU and MLC pattern the same as the AAA calculation. The dose calculation grid was set to 2.5 mm for all cases. For AXB, the dose to medium mode was used for dose calculation. Then, no normalization mode was automatically selected for all plans of IMRT and VMAT calculated by both algorithms. The dose was verified by measurement using ArcCheck diode array detector before all VMAT plans were sent to treat patient. The dose comparison was analyzed by Gamma index (3%, 3 mm) under 95% passing rate.

For analysis, although the plans used 2 treatment techniques as IMRT and VMAT, the dose comparison between AAA and AXB algorithm was determined in each same treatment technique.

Statistical analysis

For the PTV, the dose parameters generated by the dose-volume histogram (DVH) using Eclipse treatment planning are conformity index (CI), dose coverage, maximum dose, D₉₈ and D₉₅. The maximum dose (D_{max}) of plan must not be more than 110% of prescription dose. Therefore, D_{max} was determined for PTV. The CI was used to evaluate the treatment plans calculated by Equation [1] (17).

$$CI = \frac{PTV_p}{TV} \quad [1]$$

where PTV_p is the volume of target receiving the prescription dose and TV is the target volume (17-19). CI is a parameter to explain the good dose conformed in PTV. The perfect conformity value is a unity.

For the OARs, the analytic parameters applied the mean dose and the maximum dose (1 cc for brainstem and 1% for the other organs). The results of dose difference between AAA and AXB were compared using *t*-test for statistical analysis. The threshold value was set at P<0.05 for

significant difference. The percentage of dose difference (% dose diff) was computed using Equation [2].

$$\% \text{ Dose diff} = \left(\frac{\text{Dose of AAA} - \text{Dose of AXB}}{\text{Dose of AXB}} \right) \times 100 \quad [2]$$

Ethical statement

The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by institutional review board (IRB) of Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand of No. 268/62 and informed consent was taken from all the patients.

Results

Target volume

The doses were analyzed with DVH. The average dosimetry results of PTV calculated by AAA and AXB for IMRT and VMAT techniques are shown in *Table 3*. The example DVHs of PTV comparisons between AAA and AXB calculations for IMRT and VMAT techniques are shown in *Figure 2*. The dose of PTV was normalize at 95% of isodose line, for dose coverage of target compared between AAA and AXB calculation. The dosimetric comparisons of PTV between AAA and AXB at D₉₅, D₉₈, and maximum dose for each case are shown in *Figure 3*. The highest differences of dose at D₉₅, D₉₈, and maximum dose were 21.38%, 25.79% and 5.46% for IMRT technique, and 27.92%, 33.25% and 2.33% for VMAT technique, respectively. The dose of AAA was higher than AXB in all cases at D₉₅ and D₉₈ supported by P value. On the other hand, the maximum dose of AXB was significantly higher than AAA in all cases. The highest difference in dose was found in case no. 2, which the dose coverage was dropped when it was calculated by the AXB

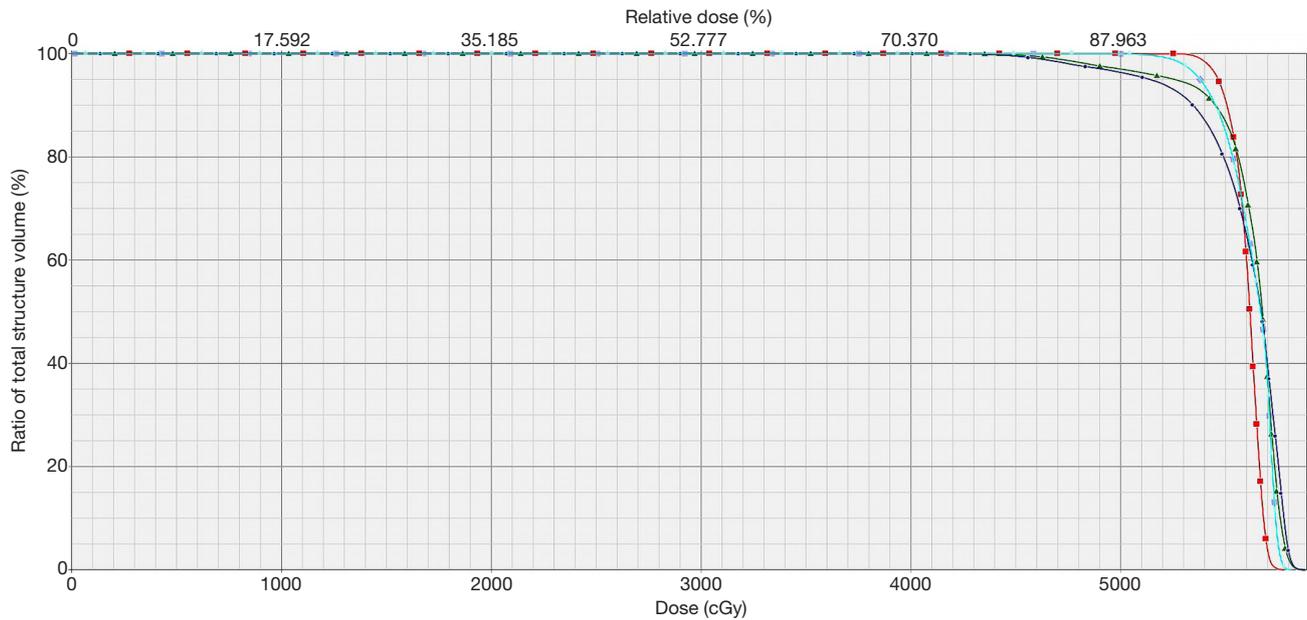


Figure 2 The example DVH of PTV comparisons between AAA and AXB calculations for IMRT and VMAT techniques (red line: AAA for IMRT, Green line: AXB for IMRT, cyan line: AAA for VMAT and blue line: AXB for VMAT). DVH, dose-volume histogram; PTV, planning target volume; AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy.

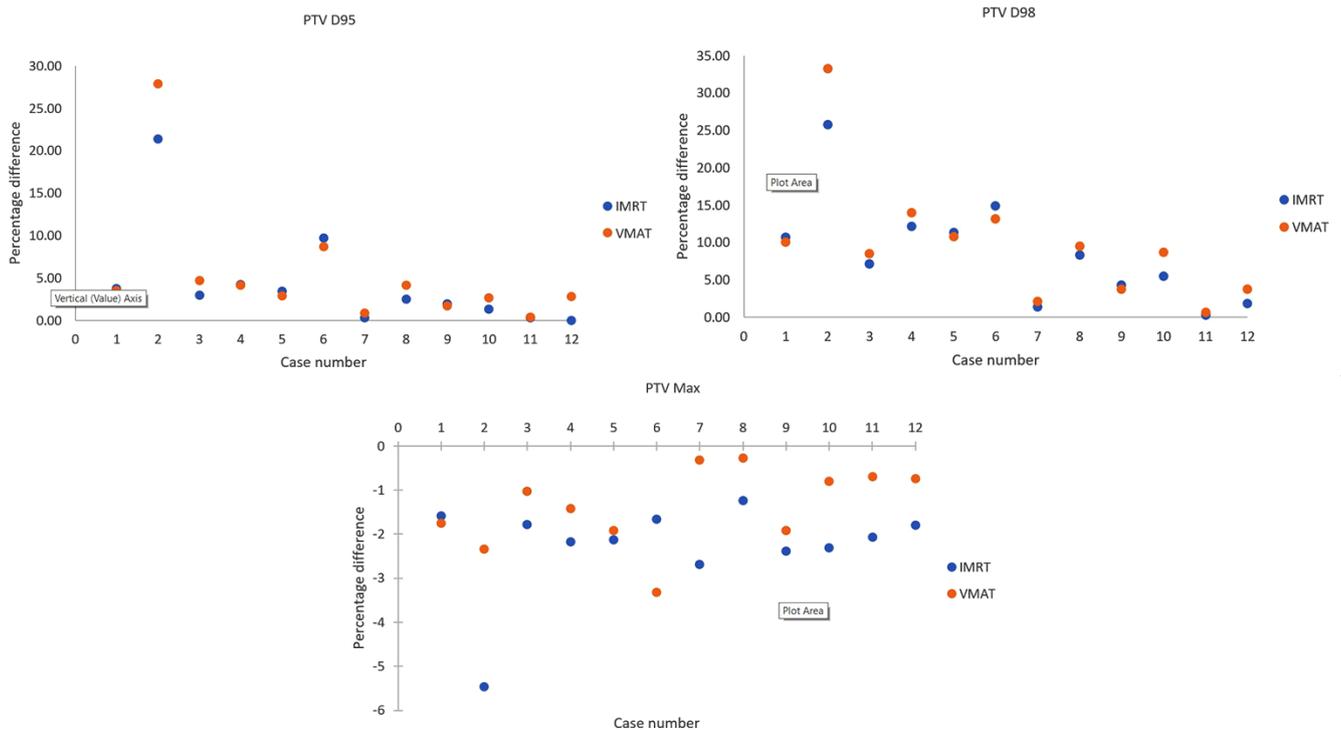


Figure 3 The dose differences between AAA and AXB calculation for both IMRT and VMAT plans for each case of patient. AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy; PTV, planning target volume.

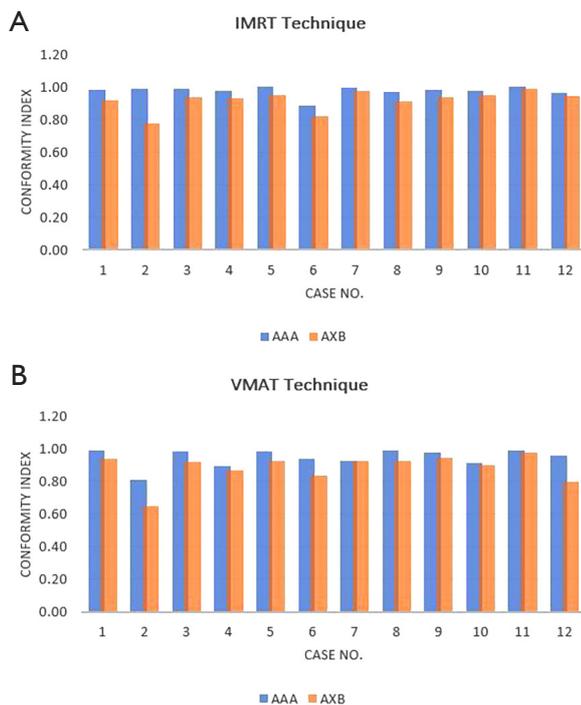


Figure 4 The conformity index value of AAA and AXB. (A) Using IMRT technique. (B) Using VMAT technique. AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy.

algorithm. This case is a small target volume and a high volume of air cavity filled.

For plan quality evaluation, the CIs for both dose algorithms are presented in *Table 3*. The CI comparison between two algorithms for both IMRT and VMAT plans are displayed in *Figure 4*. The CI value of 1.00 represents the perfect coverage of prescribed dose to PTV. The CI decreased when the dose calculation algorithm was changed from AAA to AXB in both IMRT and VMAT treatment techniques. The difference of CI value between AAA and AXB was significant with P value <0.05 in both treatment techniques. So, the dose conformity and dose coverage of AAA was better than AXB in all cases.

The dose distributions of axial plane displayed at the central axis of IMRT and VMAT plans for both calculation algorithms are shown in *Figure 5*. It was found that isodose line of dose prescription did not cover a whole of PTV, especially in the overlapping area with the air cavity. Therefore, when the AXB was used to calculate in the same plans, the prescribed isodose line was not cover the PTV

compared with the AAA for both IMRT and VMAT plans.

Normal organs

The doses of normal organs were calculated by AAA and AXB for IMRT and VMAT techniques shown in *Table 4*. The maximum doses as well as D_{1cc} for brain stem in IMRT plans between AAA and AXB was 40.6 ± 5.9 and 40.7 ± 5.9 Gy, respectively. On the other hand, the D_{1cc} in VMAT plans was 38.8 ± 6.6 and 38.7 ± 6.5 Gy for AAA and AXB calculations, respectively. For optic chiasm, the maximum doses (D_{max}) between AAA and AXB was 50.8 ± 2.9 and 51.1 ± 3.1 Gy for IMRT techniques, while VMAT techniques received the dose of 50.8 ± 3.1 and 51.0 ± 3.2 Gy.

In the case of both optic nerves, for IMRT plans, the D_{1cc} of right optic nerve calculated by AAA and AXB was 34.3 ± 9.5 and 34.6 ± 9.7 Gy, respectively. For VMAT plans, the right optic nerve received dose of 32.4 ± 8.6 and 32.5 ± 8.7 Gy for AAA and AXB algorithms, respectively. For left optic nerve, AAA and AXB algorithms provided the D_{1cc} of 37.1 ± 10.3 and 37.3 ± 10.5 Gy, respectively, for IMRT plans. For VMAT plans, the left optic nerve received dose of 35.4 ± 10.2 and 35.7 ± 10.3 Gy for AAA and AXB algorithms, respectively. The maximum dose of both cochlear was very less compared with tolerance dose.

For the normal organs, the doses calculated by AAA was slightly less than by AXB in all organs for both IMRT and VMAT techniques. Moreover, mostly normal organs dose by using IMRT technique was higher than by using VMAT technique in the same calculation algorithms. However, the dose of normal organs were not any significant statistic differences in both algorithms and treatment techniques by P value >0.05 .

Discussion

In this study, the determination of dose calculation algorithm AAA against AXB was compared in clinical situation for pituitary cancer cases. The dose comparison between both algorithms was determined in the same treatment technique. This is an interesting case because some area of PTV was filled with the air cavity. The validation was performed by IMRT and VMAT treatment techniques. The dose result calculated by AXB covered the target lesser than AAA in air cavity region for both treatment techniques. It can be analyzed by D_{95} , D_{98} and CI values from DVH. So, the doses at D_{95} and D_{98} values

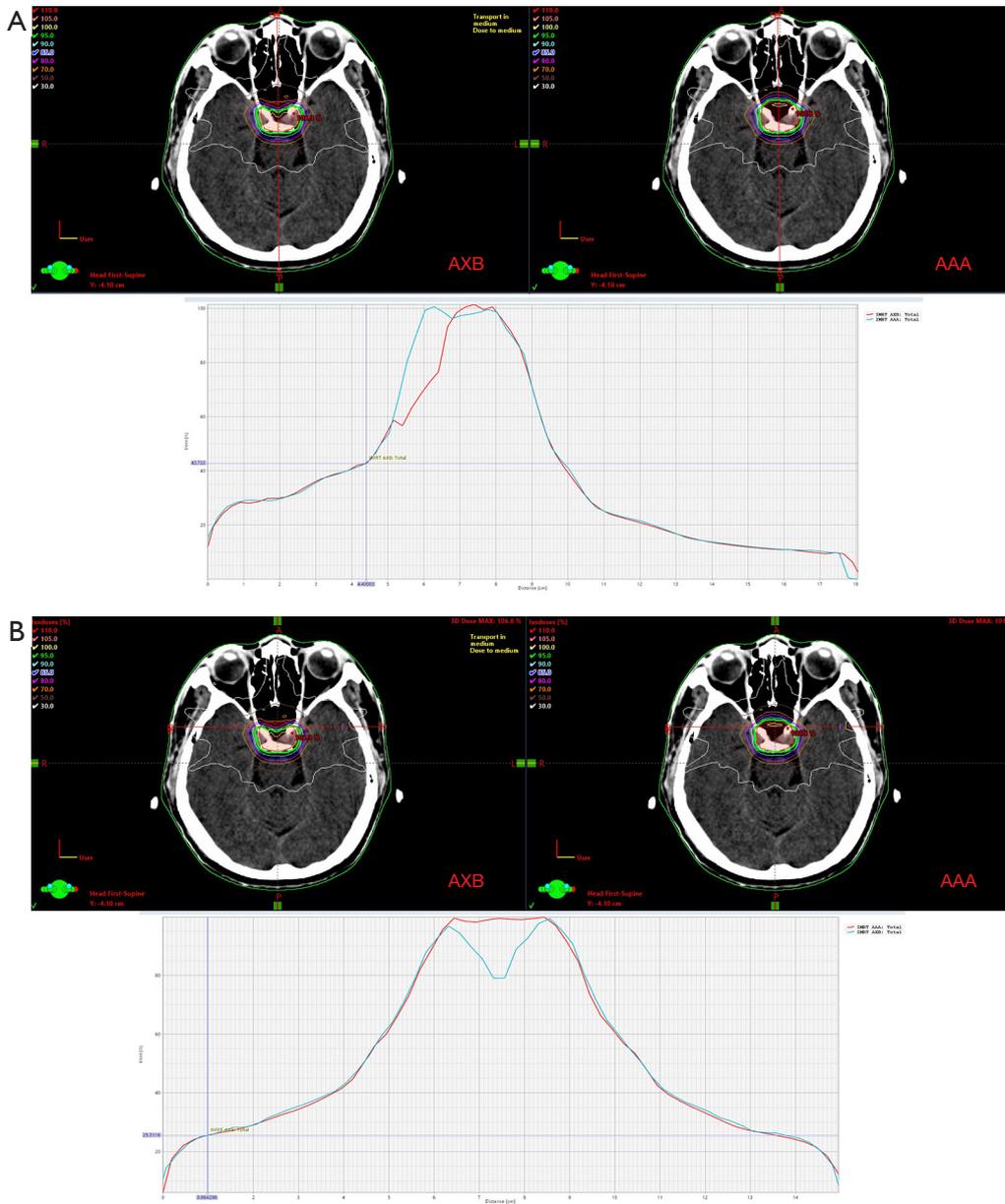




Figure 5 The dose distribution and dose profile of AAA and AXB for IMRT and VMAT plans. For dose profile, blue line is AXB calculation and red line is AAA calculation. (A) The dose distribution of axial plane and the dose profile of anteroposterior axis comparison between AXB and AAA for IMRT plans. (B) The dose distribution of axial plane and the dose profile of right-left lateral axis comparison between AXB and AAA for IMRT plans. (C) The dose distribution of axial plane and the dose profile of anteroposterior axis comparison between AXB and AAA for VMAT plans. (D) The dose distribution of axial plane and the dose profile of right-left lateral axis comparison between AXB and AAA for VMAT plans. AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy.

from AXB were significantly lower than dose from AAA. Moreover, CI values of AXB was lower than of AAA for both treatment techniques, the different values were significant. When the dose was determined by visual isodose

distribution, the prescribed dose was dropped in the cavity region. Especially, in the case of a small target volume and a high volume of air cavity filled, the dose difference was too high such as the case number 2. This effect caused

Table 4 The average dosimetric comparison of AAA and AXB for both IMRT and VMAT techniques in normal organs

Organ at risk	IMRT (n=12)		VMAT (n=12)	
	AAA	AXB	AAA	AXB
Brain stem				
D _{mean} (Gy)	18.2±5.8	18.3±5.8	17.1±6.2	17.2±6.2
D _{1cc} (Gy)	40.6±5.9	40.7±5.9	38.7±6.1	38.8±6.1
Optic chiasm				
D _{mean} (Gy)	44.3±7.8	44.6±7.9	44.1±7.2	44.3±7.5
D _{max} (Gy)	50.8±2.9	51.1±3.1	50.8±3.1	51.0±3.2
Right optic nerve				
D _{mean} (Gy)	17.3±5.5	17.5±5.6	17.1±5.3	17.2±5.2
D _{max} (Gy)	34.3±9.5	34.6±9.7	32.3±9.5	32.4±9.7
Left optic nerve				
D _{mean} (Gy)	19.2±6.5	19.3±6.5	19.9±6.6	20.0±6.7
D _{max} (Gy)	35.7±8.6	35.8±8.7	34.1±8.6	34.4±8.6
Right cochlear				
D _{mean} (Gy)	10.7±5.4	10.9±5.7	10.0±5.6	10.2±5.8
D _{max} (Gy)	15.6±6.4	16.0±6.6	14.5±6.4	14.8±6.7
Left cochlear				
D _{mean} (Gy)	10.8±5.1	10.9± 5.2	10.1±5.2	10.3±5.4
D _{max} (Gy)	17.2±8.4	17.6±8.6	16.7±9.7	17.0±9.7

AAA, anisotropic analytical algorithm; AXB, Acuros XB; IMRT, intensity-modulated radiotherapy; VMAT, volumetric modulated arc therapy.

the AAA overestimated dose in air or low density due to lack of scattering when compared with AXB (11,14,20). Accordingly, AXB algorithm was more accurate than AAA from calculated dose in heterogeneity medium (3,8,9,11,14). Many studies reported the dose comparison of AAA and AXB with MC simulation. They found that the dose of AXB agreed with MC better than the dose of AAA for inhomogeneous medium (14,21).

The dose differences between AAA and AXB were larger in the very low-density region for small field size and large air gap. Fogliata *et al.* (14) studied the dose difference of AAA and AXB comparison with MC in low density (normal lung of -790 HU) and very low density (light lung of -942 HU). They found the dose difference of AAA and MC in very low density higher than low density. On the other hand, the dose difference of AXB and MC was good agreement in both tissue densities. Inside the most internal light lung material the differences between AXB

and MC were small. So, The HU affected with AAA more than AXB algorithms in low density. The HU of air cavity filled in this study was -1,000 HU. Rana *et al.* (6) showed the dose differences between calculation algorithms and measurement. The differences were larger in small field sizes for both AAA and AXB. These effects were enhanced for AAA than AXB. For dose comparison between different air gap distance, the dose difference was higher when the air gap was thicker. There are the reasons for the dose differences between AAA and AXB enhanced in the case that had the large air cavity filled and a small volume of PTV. Thus, it affects to dose coverage of the target in this study according to CI value. From the results, the CI values calculated by AXB were less than calculated by AAA for all cases. The comparison of dose coverage between AAA and AXB showed a statistical difference for both IMRT and VMAT plans by P value. Although AAA seems better than AXB for dose coverage of PTV, AXB is more accurate

than AAA for dose calculation in inhomogeneous medium. Thus, the dose which shows the good coverage of AAA is incorrect in the air cavity region. The dose predicted by AAA and AXB depends on the combined effects of field size, the thickness of the air cavity, and photon energy (6,14). However, this study did not investigate the different energy.

Normal organs dose was similar for both AAA and AXB, and for all treatment techniques. The dose difference in normal organs did not show any statistical difference because the dose calculated by AAA and AXB was good agreement in homogeneous tissue (9). The doses calculated by AAA was less than by AXB in all organs.

Conclusions

The dose was compared in air cavity region of target volume for pituitary carcinoma by AAA and AXB algorithms. The dose coverage of target using AXB was less than using AAA for calculation in the air cavity region which it depended on the size of target and volume of air cavity. In general, the AXB can calculate dose accurately than AAA for inhomogeneous region. The good dose coverage to target displayed by AAA calculation should be determined carefully in clinical when evaluation dose in the target is filled by air cavity. To be able to accept the best plan in normal practice. The evaluation parameters such as D_{95} , D_{98} , D_{max} and CI for target volume, and dose tolerance for normal tissues must be used to determine. Together with other tools (DVH, isodose distribution etc.), it should be determined within the accepted criteria. An attempt should be made to avoid large air gaps when smaller field sizes are used for the radiation treatment.

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Footnote

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Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at <http://dx.doi.org/10.21037/tro-20-57>).

[org/10.21037/tro-20-57](http://dx.doi.org/10.21037/tro-20-57)). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by institutional review board (IRB) of Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand of No. 268/62 and informed consent was taken from all the patients.

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