Comparison of T2 and FLAIR MR imaging for target delineation in Glioblastoma: impact on target coverage, normal tissue exposure and pattern of recurrence.

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 Glioblastoma Multiforme (GBM) accounts for 70% of new adult cases of malignant brain tumors

• Standard primary therapy for GBM includes maximal safe resection followed by adjuvant radiation and chemotherapy

• High recurrence rates (>90%) with a median overall survival (OS) of 15-18 months

- Features of GBM:
 - Rapid progression
 - Enhancing tumor
 - Surrounding oedema (Contains tumor)





CTV - EORTC: residual tumor and/or resection cavity plus 2 cm without the intentional inclusion of peritumoral edema

CTV - RTOG: post-operative peritumoral edema plus 2 cm

Background



Current RTOG protocols advise using CT and either FLAIR or T2 images to identify CTV . . . the differences between these sequences have not been clearly defined

T2-CTV	T2-CTV
FLAIR-CTV	FLAIR-CTV
• T ternational guidelines rec ent eclesion on T1 - weighted h	he use of cost-operation ast- entry the gr is tumor veluin
• I titutional preference generally define tumor volumes and associated	ctos whether 12 or FLAIP is u d to enc
 There are limited data comparing the e imaging sequences for RT plann 	desimetric and clinical impact of sing

FLAIR MRI





The aim of this study is to evaluate the impact on target coverage and normal tissue exposure of FLAIR and T2 weighted MRIs to delineate the surrounding edema for radiation treatment planning in GBM.

Furthermore we analyzed the patterns of recurrence in relation to its peritumoral edema to gain a better understanding of whether peritumoral edema should be intentionally included within the CTV

Methods and Materials



• We used treatment planning images of adult patients with GBM treated between 2008 and 2012 at the Radiation Therapy Unit-ASMN in whom a complete pretreatment MRI with contrast-enhanced T1, T2 and FLAIR sequences was currently available for review

 \cdot All patients were treated with post-operative IMRT and a total dose of 60 Gy in 30 fx and concomitant chemotherapy (TMZ)

 $\boldsymbol{\cdot}$ For each patients the MRI scans were fused with the planning CT and used for target definition

• The GTV encompassed the resection cavity and any residual tumor as seen on a contrastenhancing T1 postoperative MRI

• Delineation of CTV was carried out by adding 2 cm margins to the GTV

• The CTV was expanded by 0.3 cm to create the PTV to compensate for variability in treatment set-up and patient motion

• All patients were followed after RT with MRI scan and recurrence was delineated by neuro-radiologist

Methods and Materials



• The CTVs studies were contoured on the T2 (CTV-T2) and FLAIR (CTV-FLAIR) sequences without comparison to the alternative sequence by a single observer



- The PTVs (PTV-T2 and PTV-FLAIR) were created with a standard 0.3 cm volumetric expansion
 - Using the "calculate volume" function, the CTVs and PTVs in cc were recorded
 - \cdot The differences between the CTVs/PTVs $\,$ were tabulated and a mean percent difference was calculated $\,$
 - The *Dice coefficient* (defined as the volume of overlap between two sets of contour, divided by their mean volume) was calculated to evaluate overlap between the CTVs/PTVs

Methods and Materials



• For each patients two IMRT plans were calculated for PTV-T2 and PTV-FLAIR to evaluate the potential consequences in respect to target coverage and normal tissue exposure using the PTVs generated with different MRI sequences



 Pattern of recurrence in terms of their relationship to the tumor volumes delineated using the different MRI sequences was evaluate for each patients







... to date were evaluated 34 cases

• The CTV_{T2} volumes and CTV_{FLAIR} were significantly different (p=0.03)



- The CTV_{FLAIR} volumes were significantly larger than CTV_{T2} (p=0.04):
 - mean volumes = 84.181 cc vs 77.535 cc
 - \cdot mean percent deviation between CTV $_{\rm FLAIR}$ and CTV $_{\rm T2}$ = 25%
 - average overlap volume between CTV_{FLAIR} and CTV_{T2} = 54.6 cc
 - \cdot average **union** volume between CTV_{FLAIR} and CTV_{T2} = 80.57 cc
 - Mean Dice Coefficient = 0.74



• The PTV_{T2} volumes and PTV_{FLAIR} were significantly different (p=0.03), with a mean volumes of 173 and 191.2 cc, respectively

 \cdot For each pt two **IMRT plans** were created by optimization on PTV_{T2} and $\text{PTV}_{\text{FLAIR}}$:

Parameters	PTV _{FLAIR}	PTV _{T2}	p
T2 IMRT plan			
V95 сс	99.8	99.9	n.s.
D98 Gy	47.79	59.8	0.02
D2 Gy	63	63.2	n.s.
CI	0.71	0.76	n.s.
FLAIR IMRT plan			
V95 сс	99.9	99.9	n.s.
D98 Gy	59.7	59.8	n.s.
D2 Gy	63.1	63.27	n.s.
CI	0.83	0.62	n.s.













 The median volume of brain irradiated to 60 Gy according to plan type was significantly higher for original plan compared with the FLAIR and T2 plans (p=0.001)

• However for pts with peritumoral edema > 75 cc the median percent volume of brain irradiated to 60 Gy was similar in the three groups





Recurrence was defined as:

- "in field" if more than 95% of the recurrence volume was in 95% isodose line of 60 Gy;
- "**marginal**" when less than 95% of the recurrence volume was outside the high dose volume;
- "distant" when recurrence was outside the RT field (<20% isodose line).



30 pts (88%) failed in central or marginal localization



Distribution of the Recurrence Volume (RV) with different plans



Conclusions



• Based on our comparison of T2 and FLAIR imaging for radiation treatment planning, both techniques are important and not interchangeable

• Each technique can help distinguish normal parenchyma from edema and abnormal tissue

• Until we have sufficient imaging to determine where a recurrence is most likely to occur, the use of a statistical margin that accounts for the pattern of failure of GBM within 2 cm of the primary tumor is supported



Thank you !