

# CT Perfusion: The Basics

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*Chief of Neuroradiology and Head & Neck Radiology  
University of Michigan Health System*



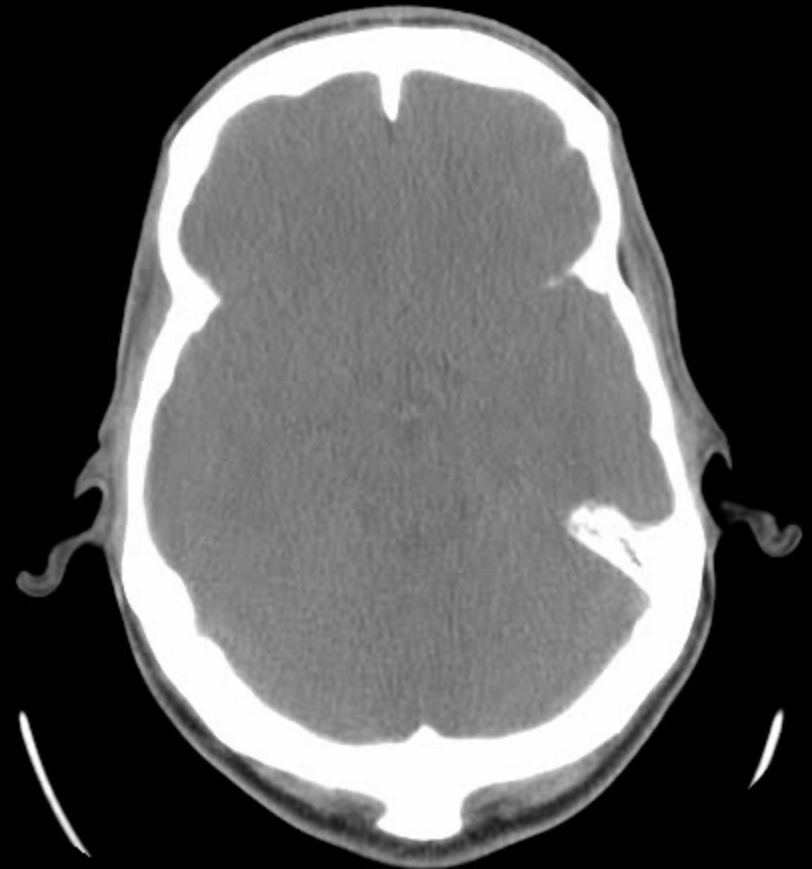
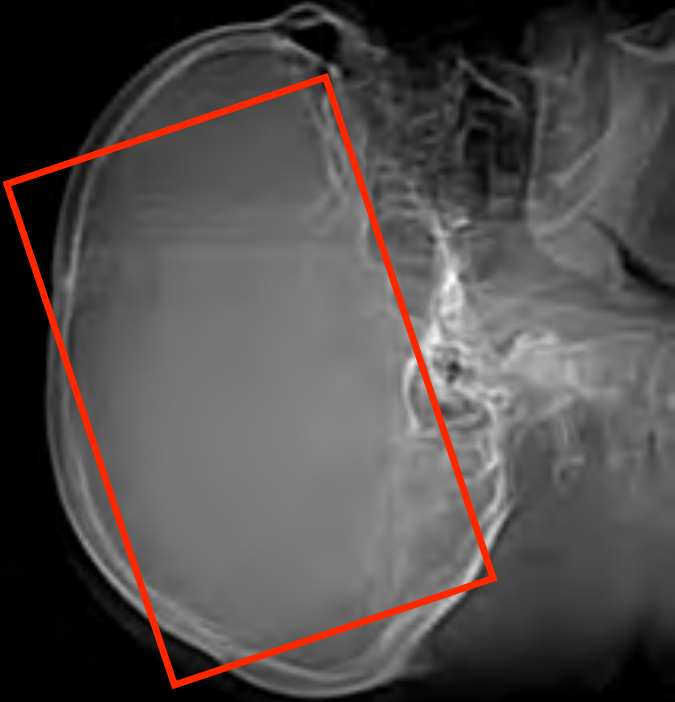
# Outline

- **Technique**
- **Clinical Applications**
  - **Stroke**
  - **Brain Tumors**
  - **Head & Neck**

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# CT Perfusion Protocol

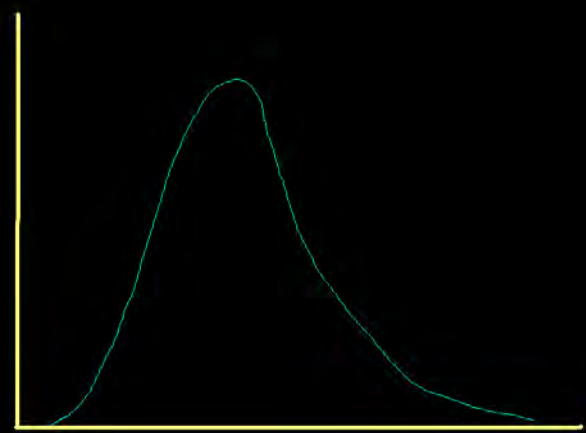
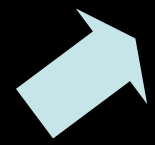
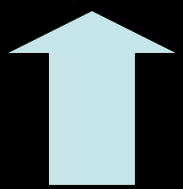
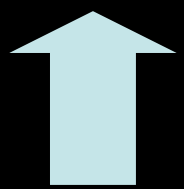


# CT Perfusion

FOV 25.0 cm

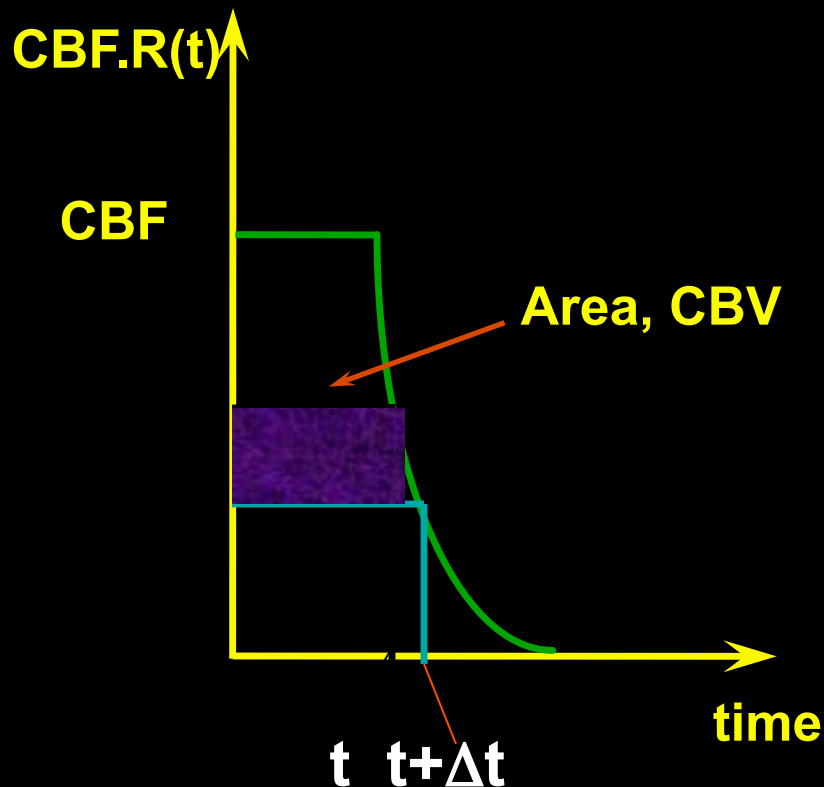


- Define ROIs for:
  - Vein
  - Artery
- Software has automated vessel selection capability



# CT Perfusion – What is behind it?

$$Q(t) = CBF \cdot C_a(t) * R(t)$$



## Deconvolution

- Technique described by Leon Axel, M.D., Ph.D. in 1983
- Measure Time-concentration in an input artery.
- Use mathematical process called “deconvolution” to separate effect of input from observed contrast time in tissue.

# Computation

- Calculations are made based on the “central volume principle” which relates blood flow, blood volume and mean transit time.

$$BF = BV / MTT$$

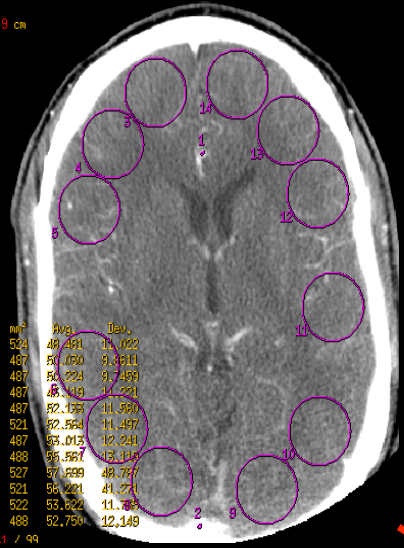


S47.7  
IFOV 20.9 cm

R  
9  
7

ROI	mm <sup>2</sup>	Avg.	Dev.
13	524	48.491	11.022
14	487	50.050	9.4611
3	487	50.224	9.7499
4	487	48.349	11.221
5	487	52.137	11.360
6	521	52.564	11.467
7	487	53.013	12.041
8	488	55.567	13.115
9	527	57.699	14.767
10	521	56.221	11.231
11	522	53.822	11.333
12	488	52.756	12.149

rank = 41 / 99

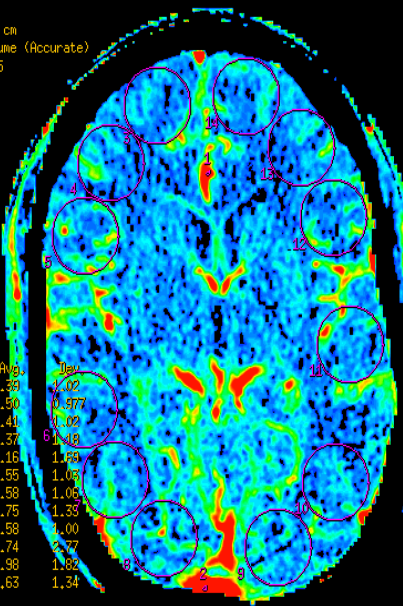


## CBV

S47.7  
IFOV 20.9 cm  
Blood Volume (Accurate)  
cursor 5x5

10  
R  
9  
7

ROI	Avg.	Dev.
13	1.39	1.02
14	1.50	1.577
3	1.41	1.02
4	1.37	1.48
5	2.16	1.63
6	1.55	1.05
7	1.58	1.05
8	1.75	1.35
9	1.58	1.00
10	1.74	1.77
11	1.98	1.82
12	1.63	1.34

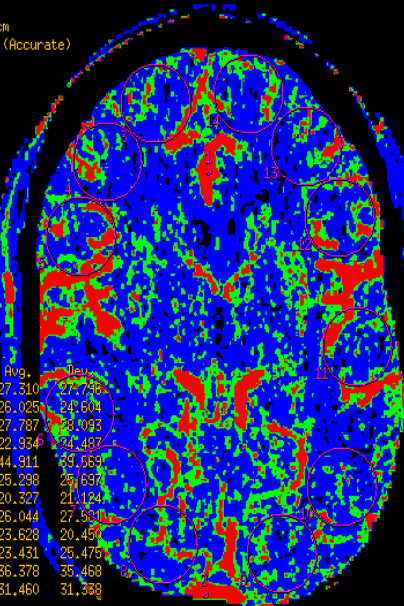


## CBF

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Blood Flow (Accurate)  
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100  
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ROI	Avg.	Dev.
13	27.310	27.736
14	26.025	24.604
3	27.787	28.093
4	22.936	21.487
5	44.911	38.689
6	25.298	29.637
7	20.327	21.124
8	26.044	27.631
9	23.628	20.450
10	23.431	25.475
11	36.378	35.468
12	31.460	31.388

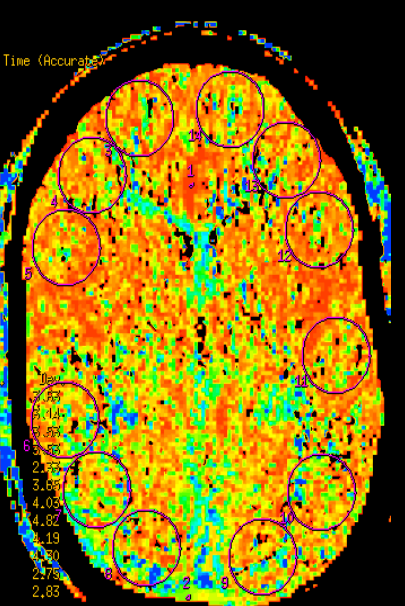


## MTT

S47.7  
IFOV 20.9 cm  
Mean Transit Time (Accurate)  
cursor 5x5

15.010  
R  
9  
7

ROI	Avg.	Dev.
13	4.35	5.55
14	4.67	5.14
3	4.51	5.53
4	4.60	5.53
5	3.73	2.77
6	5.39	3.85
7	6.07	4.03
8	5.99	4.82
9	5.45	4.19
10	5.24	4.30
11	4.04	2.73
12	3.91	2.83



# Technique

- **Non-enhanced Brain**
- **CT Perfusion acquisition acquired at level of the basal ganglia**
  - **8cm total coverage**
  - **50cc of 370 contrast**
  - **4cc/sec for 12.5 sec**

# Outline

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- **Radiation Dose Update**

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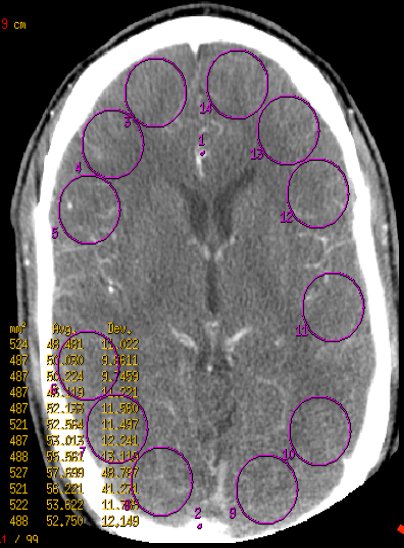
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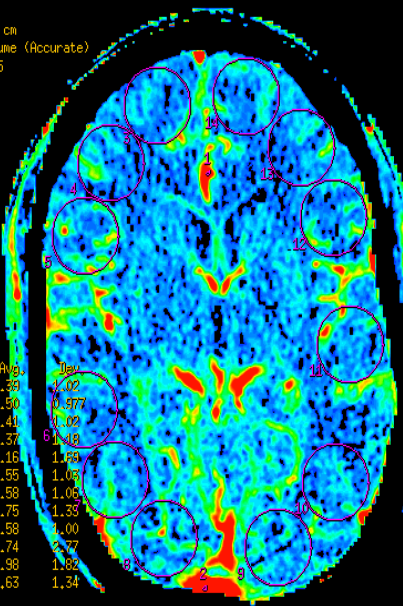


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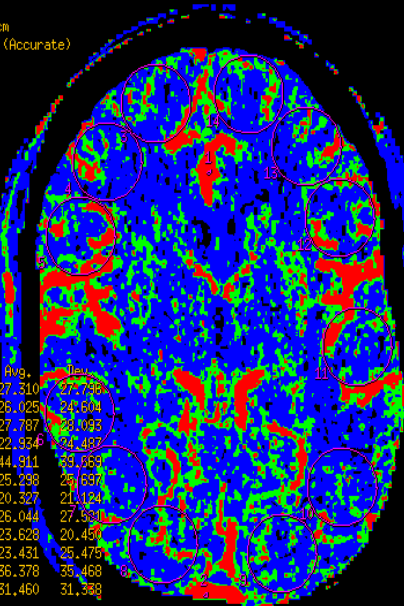


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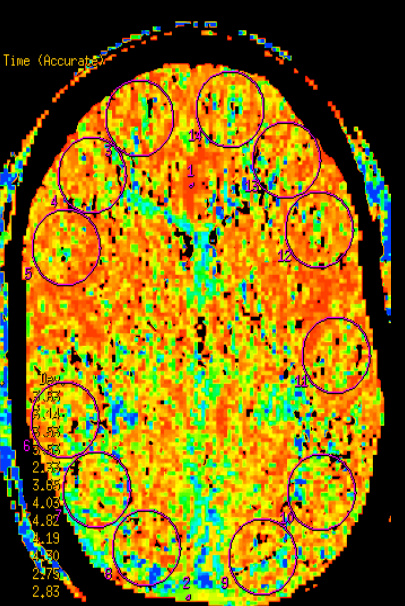


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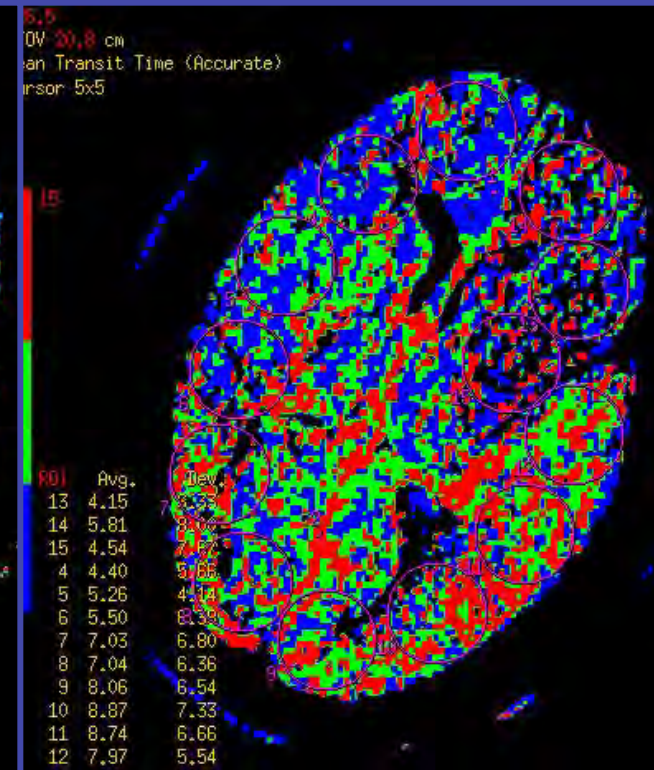
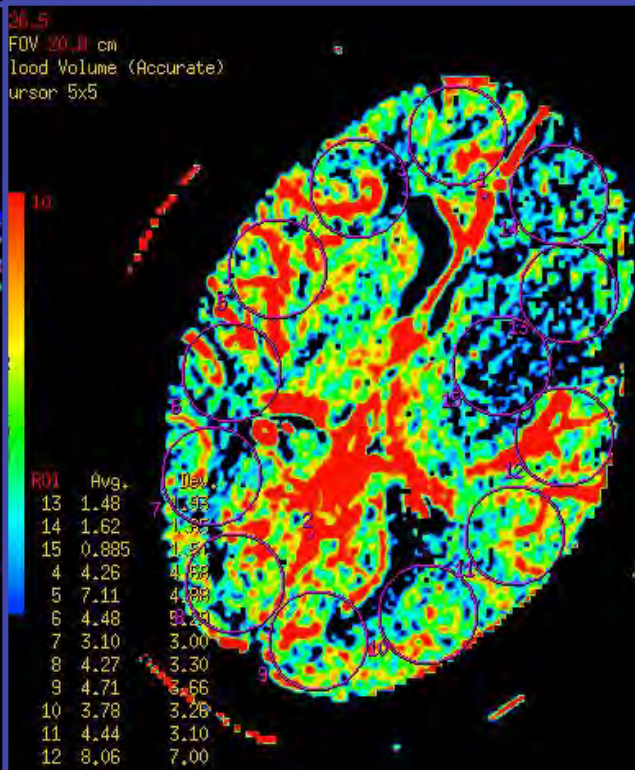
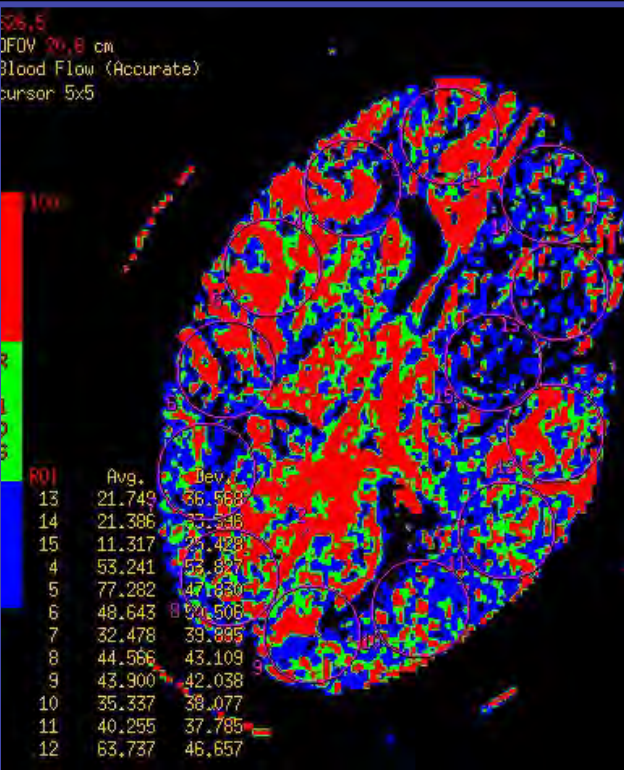
# CT Perfusion

## Qualitative Assessment

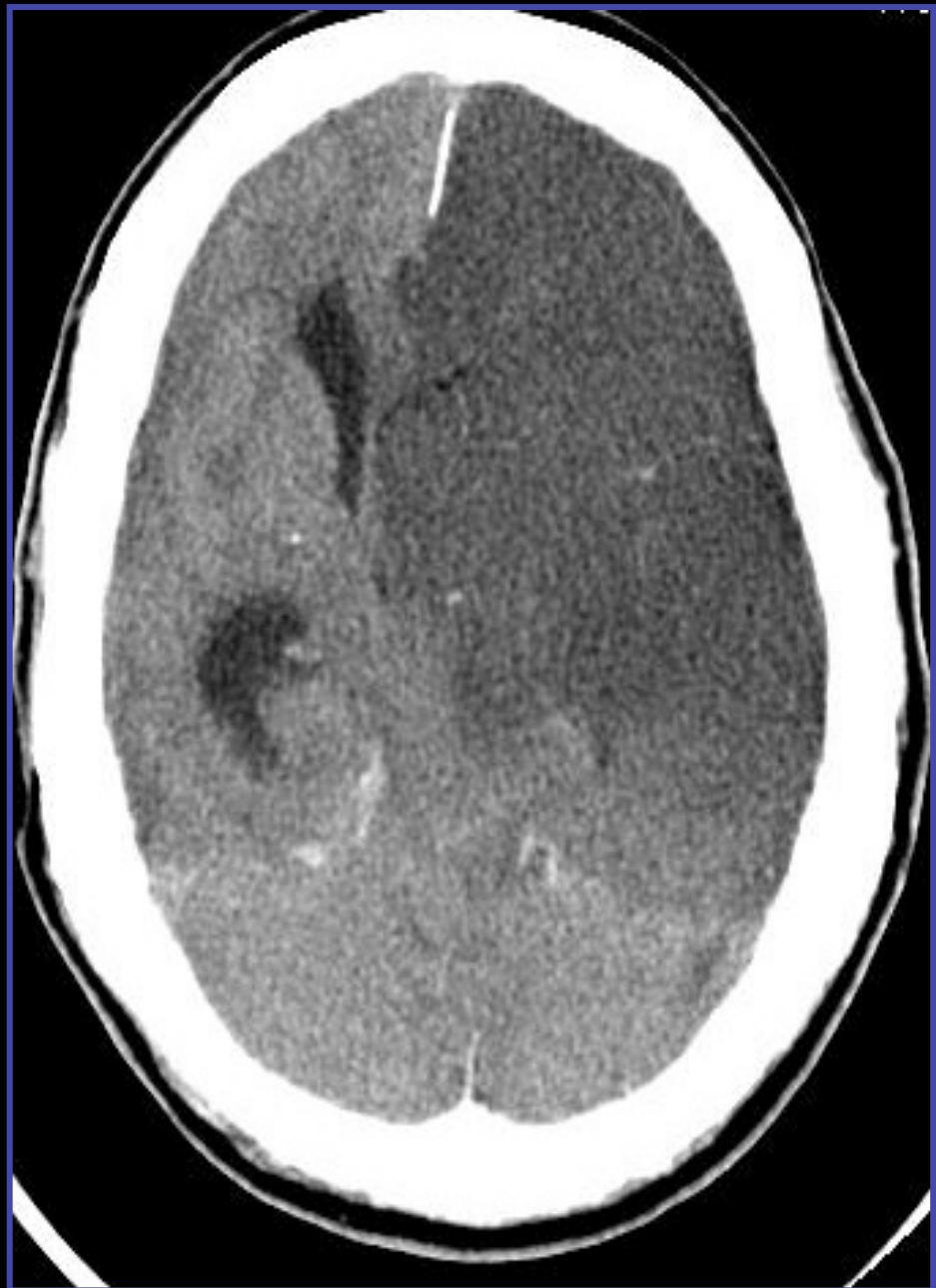
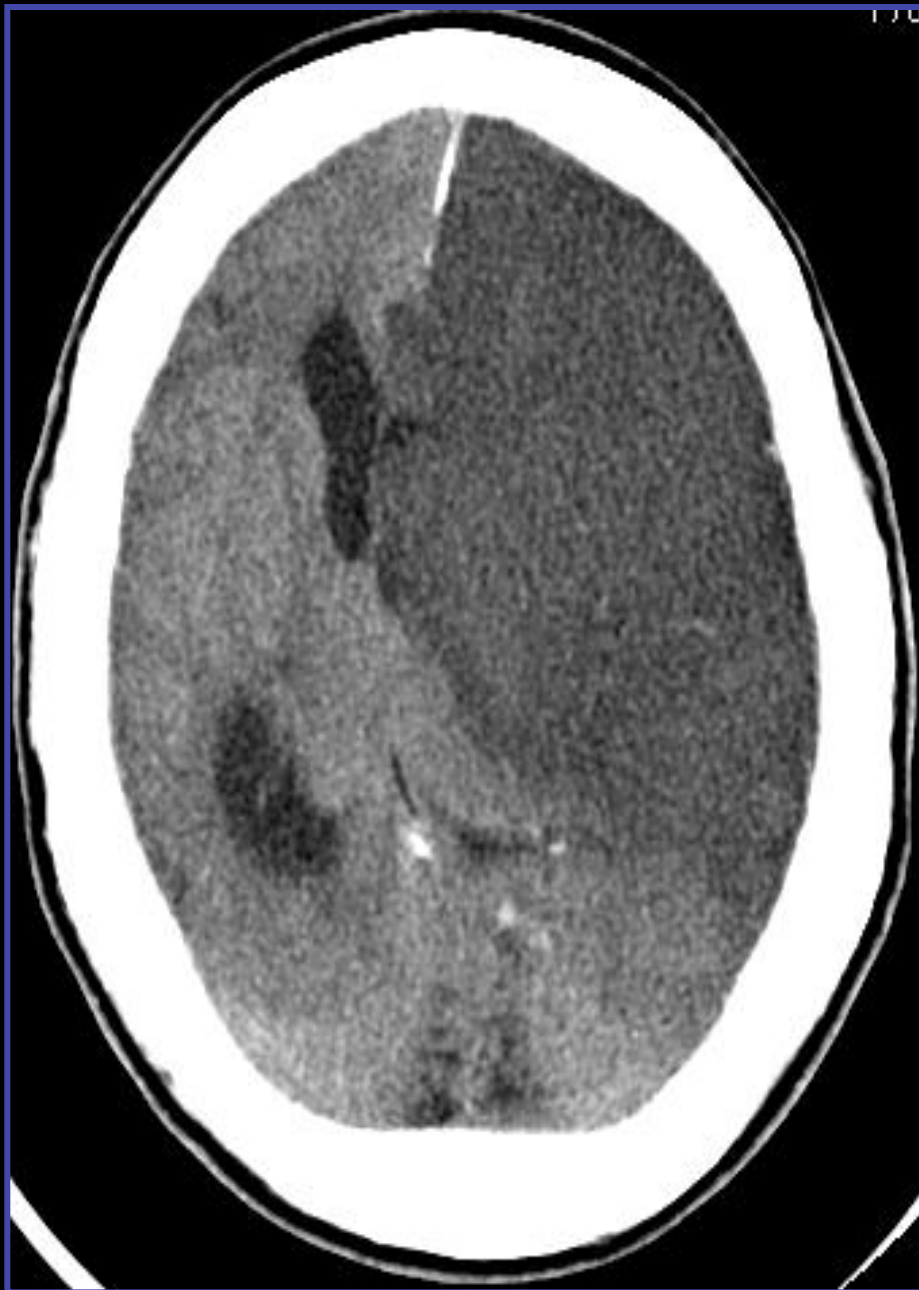
*Salvageable tissue: ↓CBF, ↔↑CBV ↑MTT*

*Infarct: ↓CBF, ↓CBV and ↑MTT*

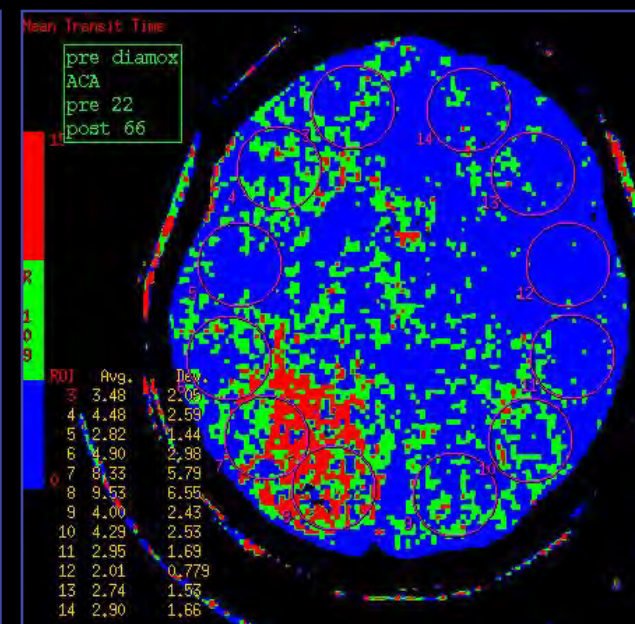
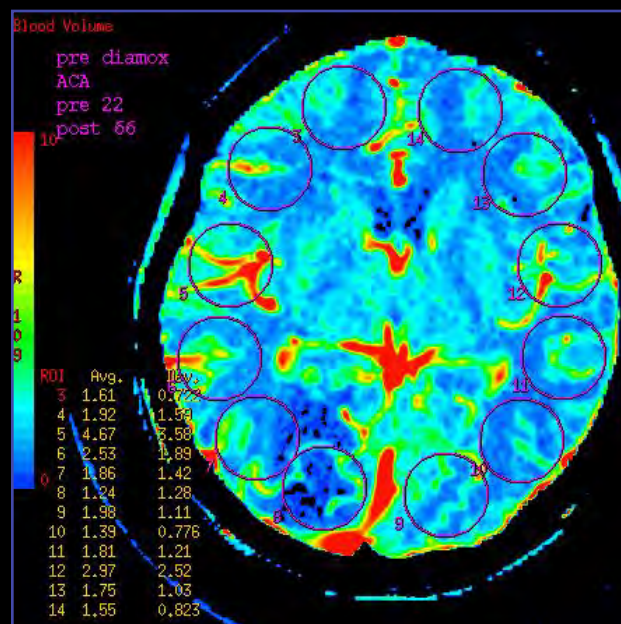
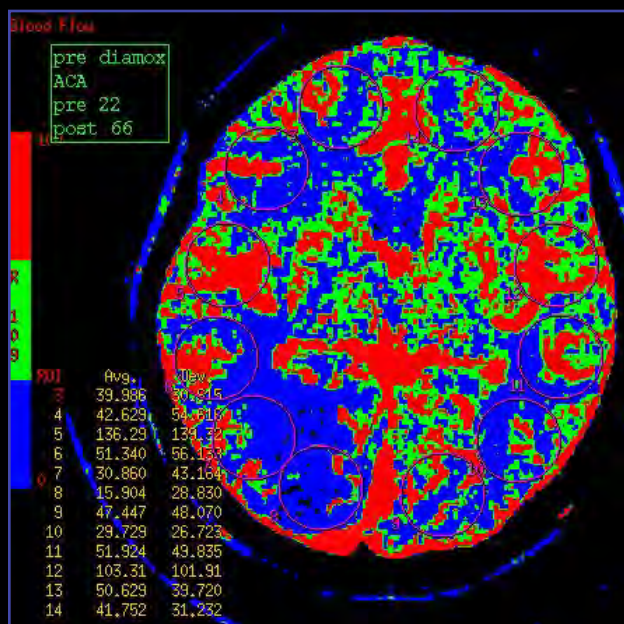
*Hunter et al Radiology 2003;227:725-730*



↓ CBF, ↓ CBV and ↑ MTT in left ACA and most of left MCA territories compatible with infarction







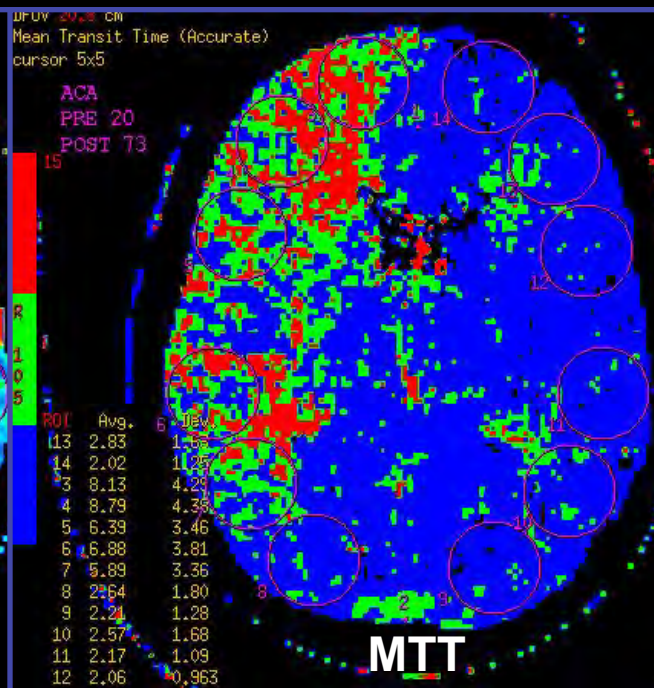
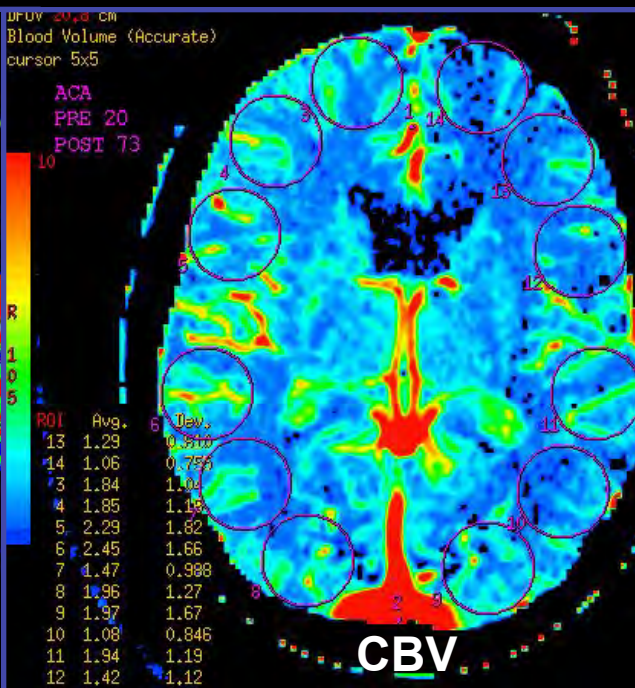
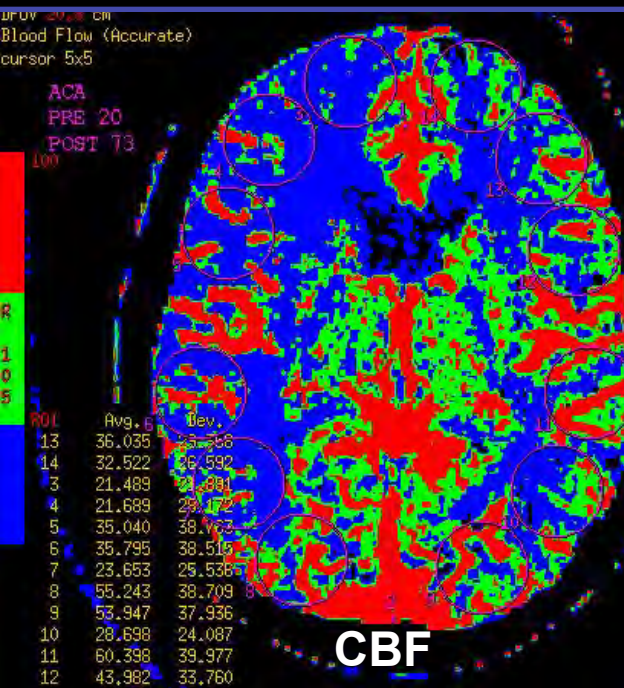
**Findings compatible with right PCA infarct**

↓ CBF

↓ CBV

↑ MTT





Perfusion CT changes compatible with ischemia

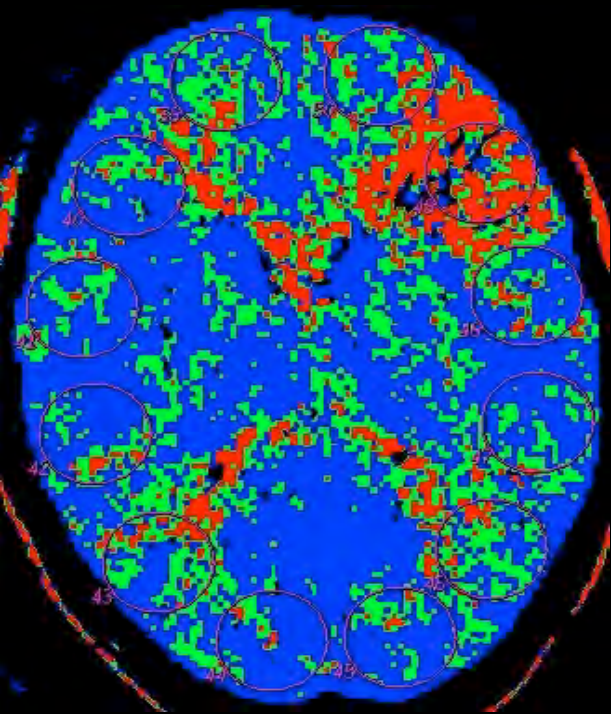
↓ CBF

→ CBV

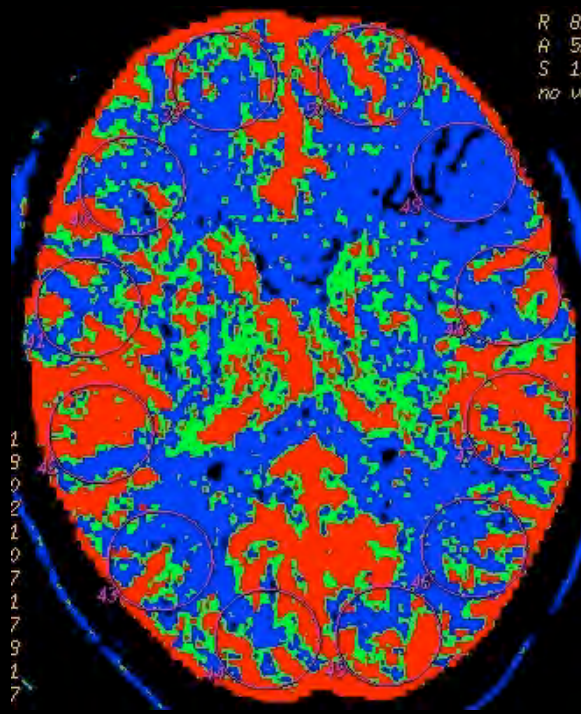
↑ MTT

No cortical infarct on 8 mo. follow up CT

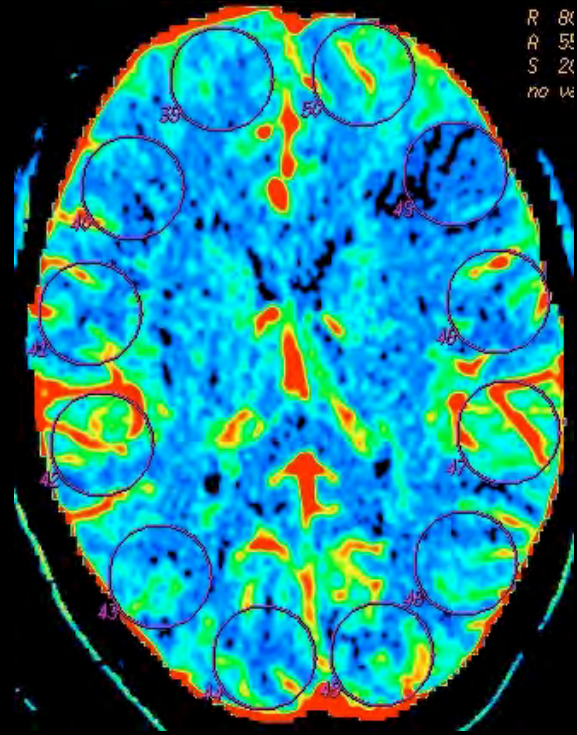




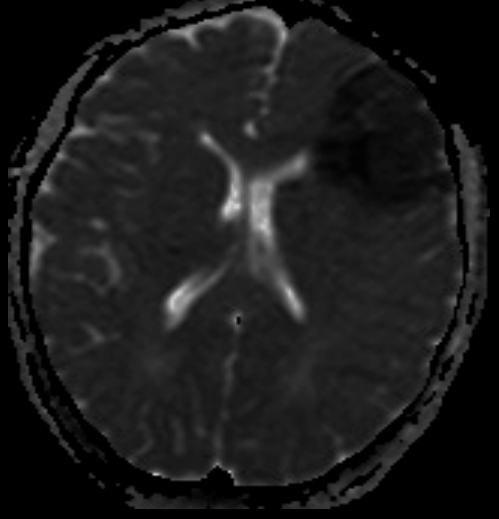
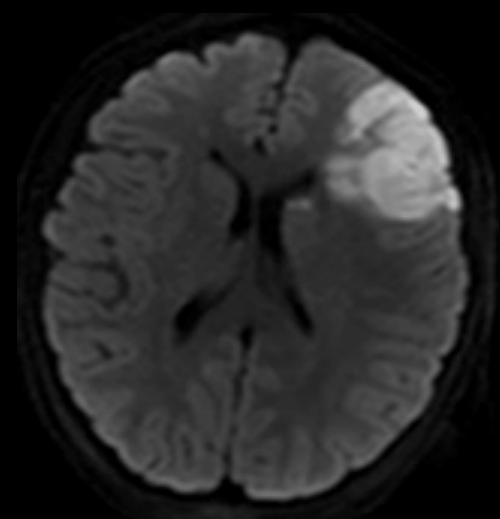
MTT

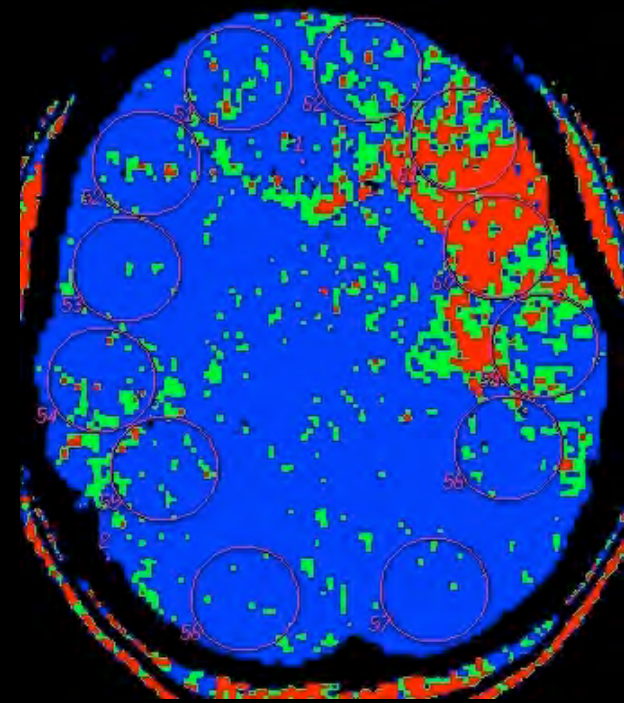


CBF

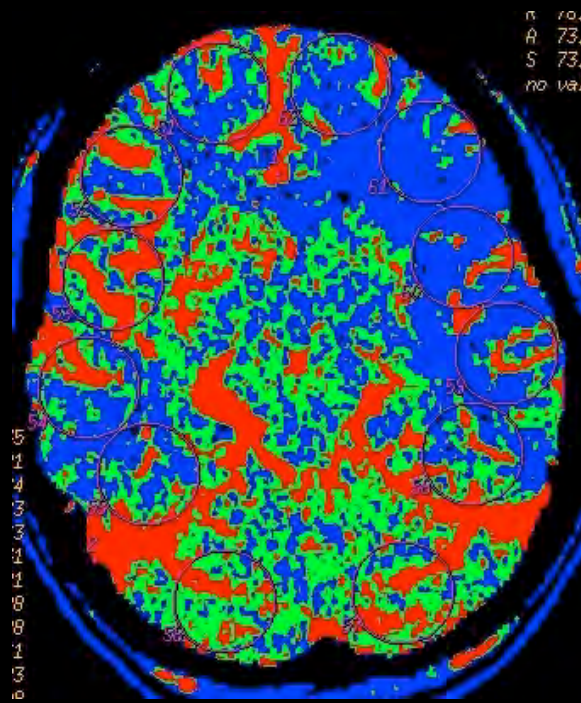


CBV

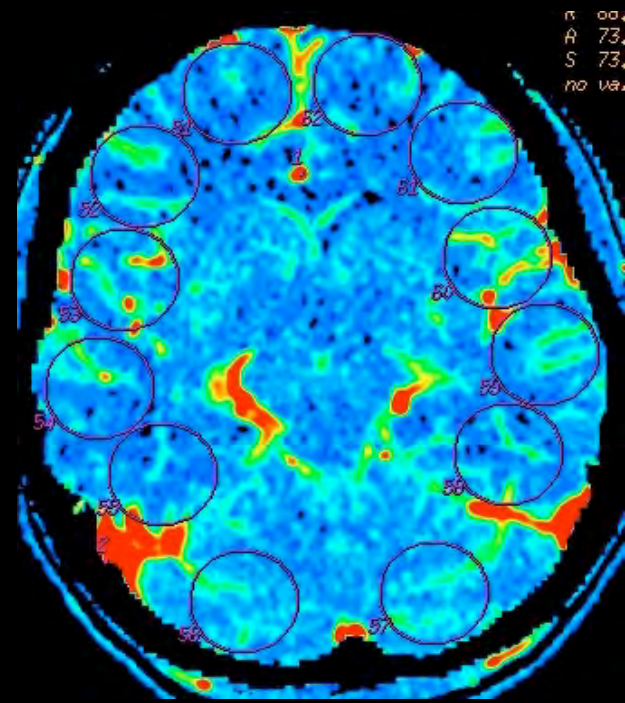




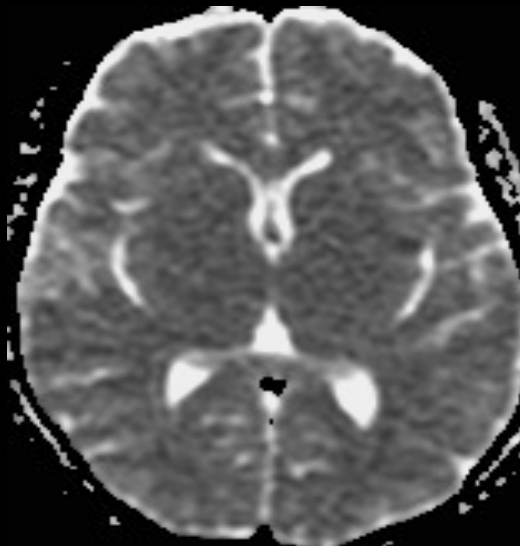
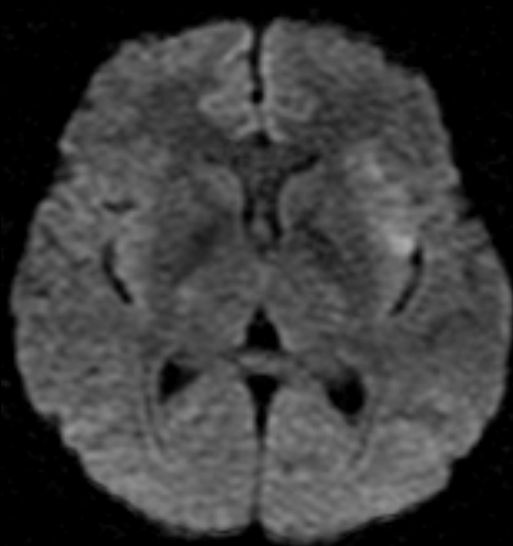
MTT



CBF



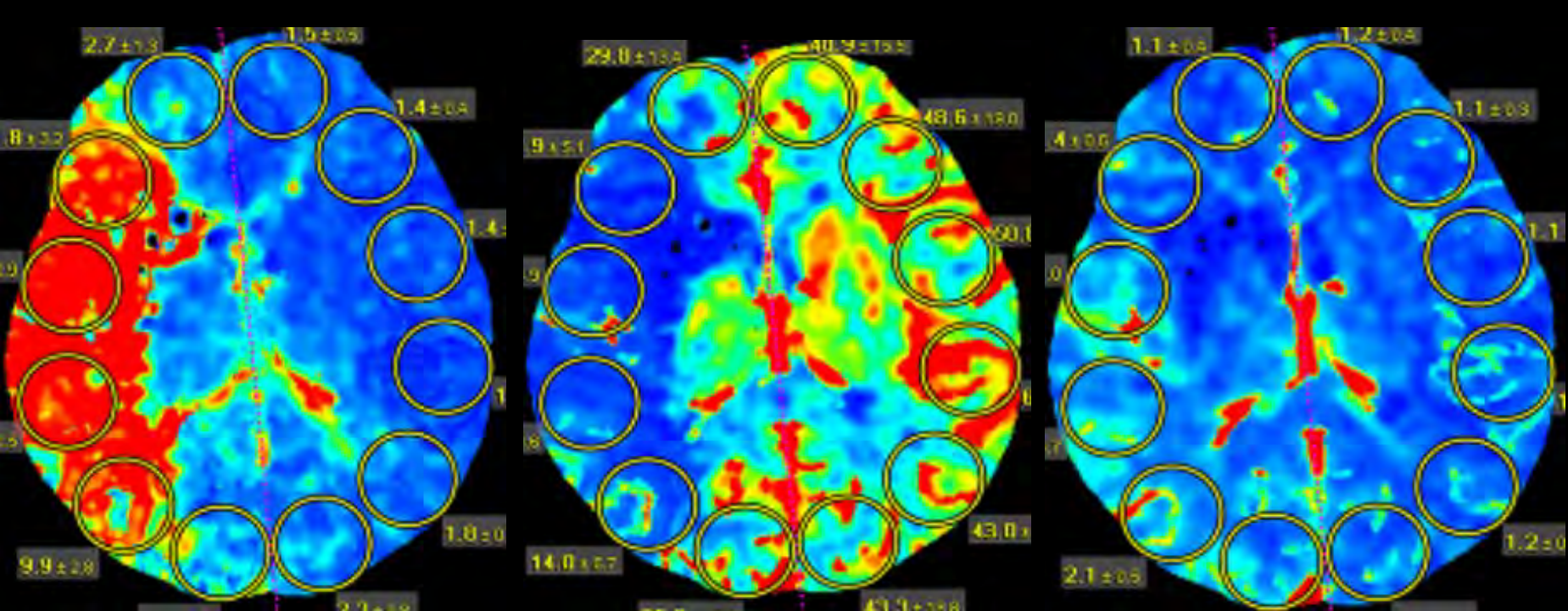
CBV



n 70.  
A 73.  
S 73.  
no val

n 00.  
A 73.  
S 73.  
no va.

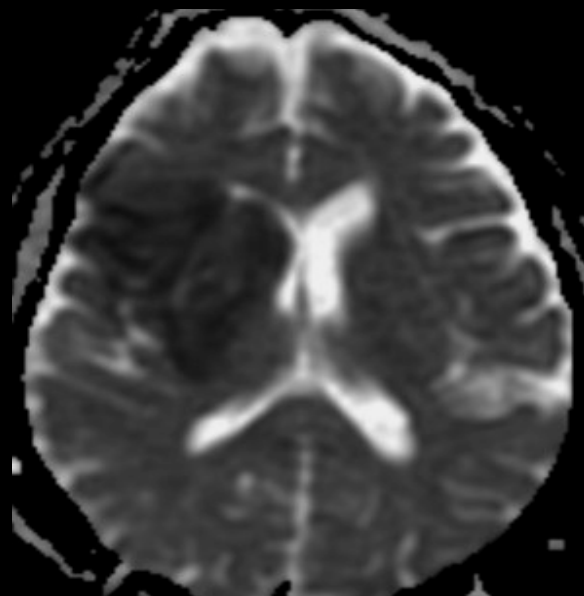
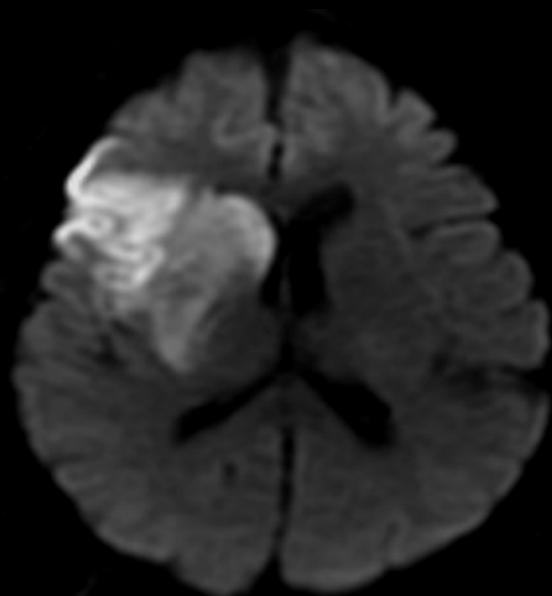
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MTT

CBF

CBV



# Outline

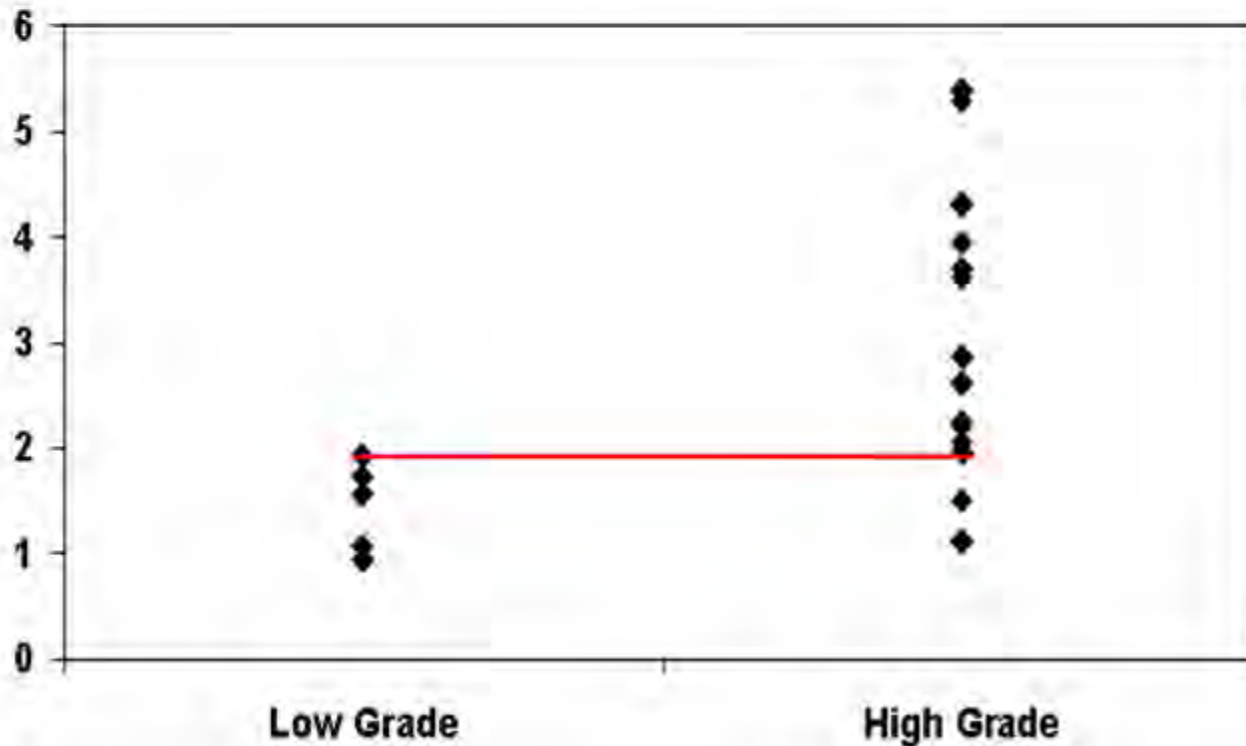
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# Perfusion CT Imaging: Glioma Grading

# patients	nCBV Mean (SD)	nCBF Mean (SD)	nMTT Mean (SD)
Low Grade (5)	<b>1.44 (0.42)</b>	<b>1.16 (0.36)</b>	<b>1.69 (1.12)</b>
High Grade (14)	<b>3.06 (1.35)</b>	<b>3.03 (2.16)</b>	<b>1.29 (0.55)</b>
<b>p-value</b>	<b>0.005</b>	<b>0.045</b>	<b>0.559</b>

# Perfusion CT Imaging: Glioma Grading

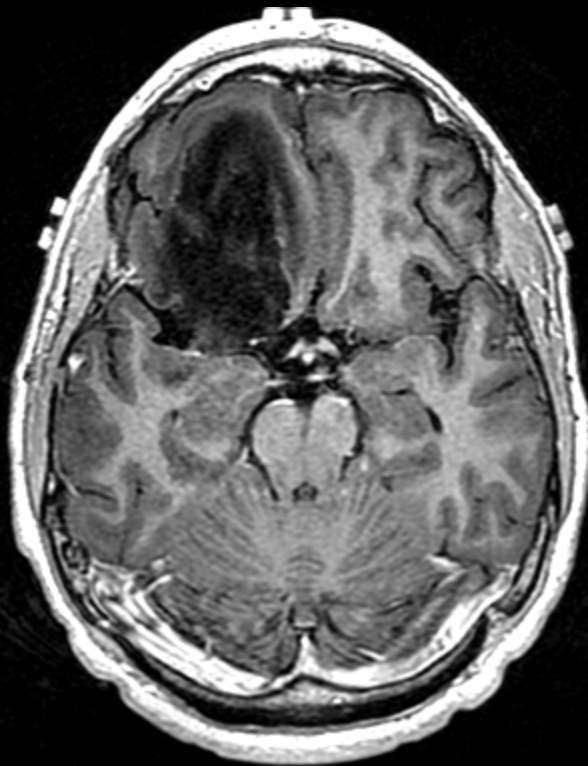
## Normalized CBV values



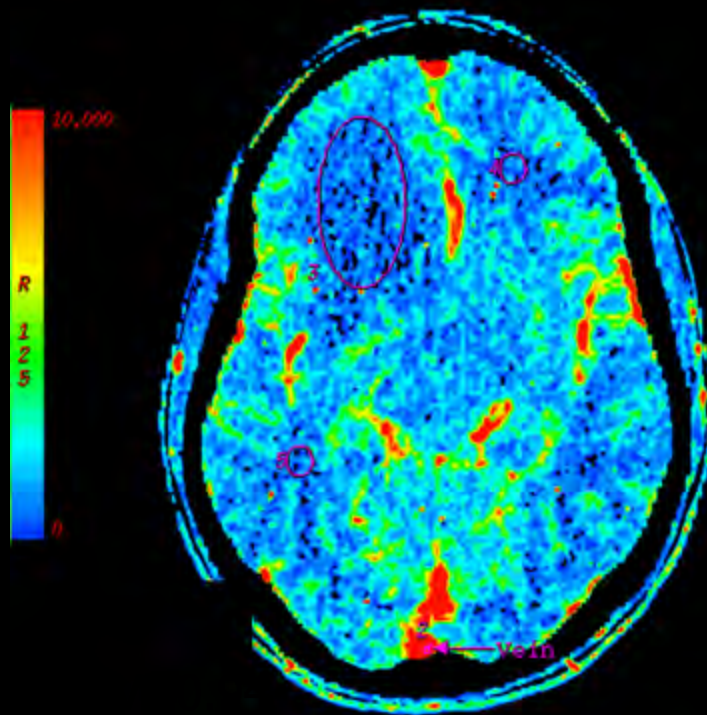


# WHO Grade II (Low Grade Glioma)

34 yo man with WHO grade II glioma.  
CBV map shows low blood volume (nCBV=0.94).



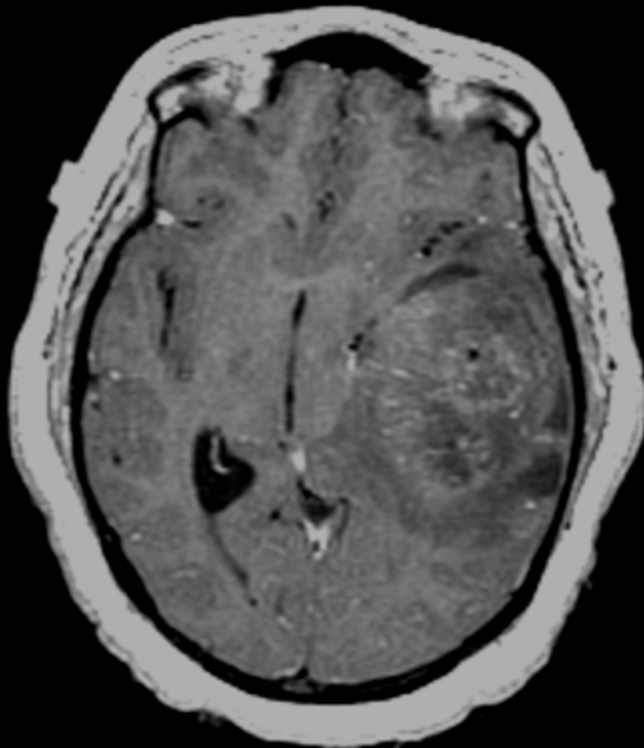
Post-gad T1WI



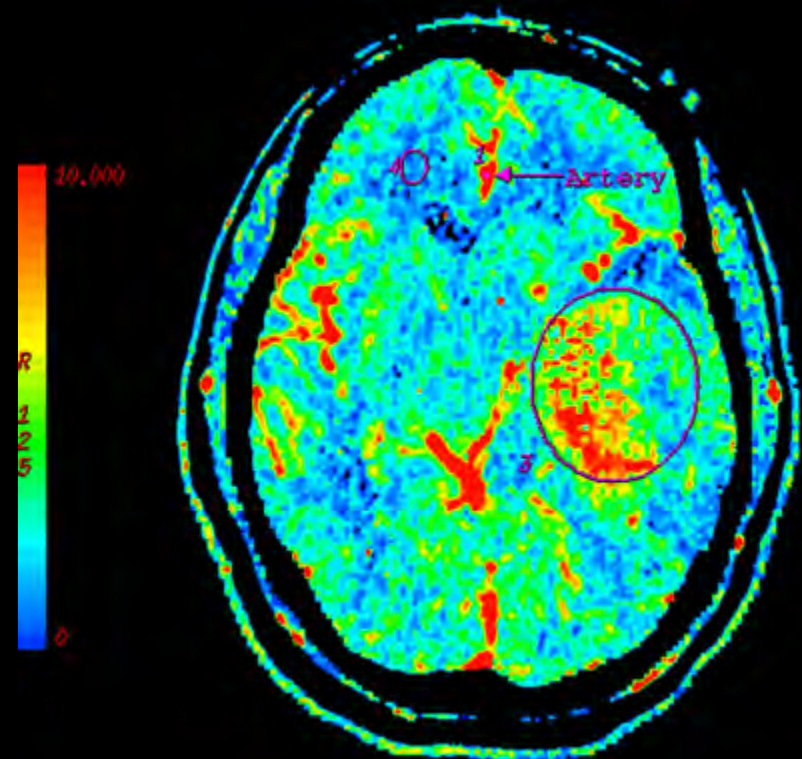
CBV map

# WHO Grade III (Anaplastic Astrocytoma)

WHO grade III glioma in a 39 yo woman who presented with seizure.  
CBV map shows higher CBV (nCBV=2.61).



Post-gad T1WI



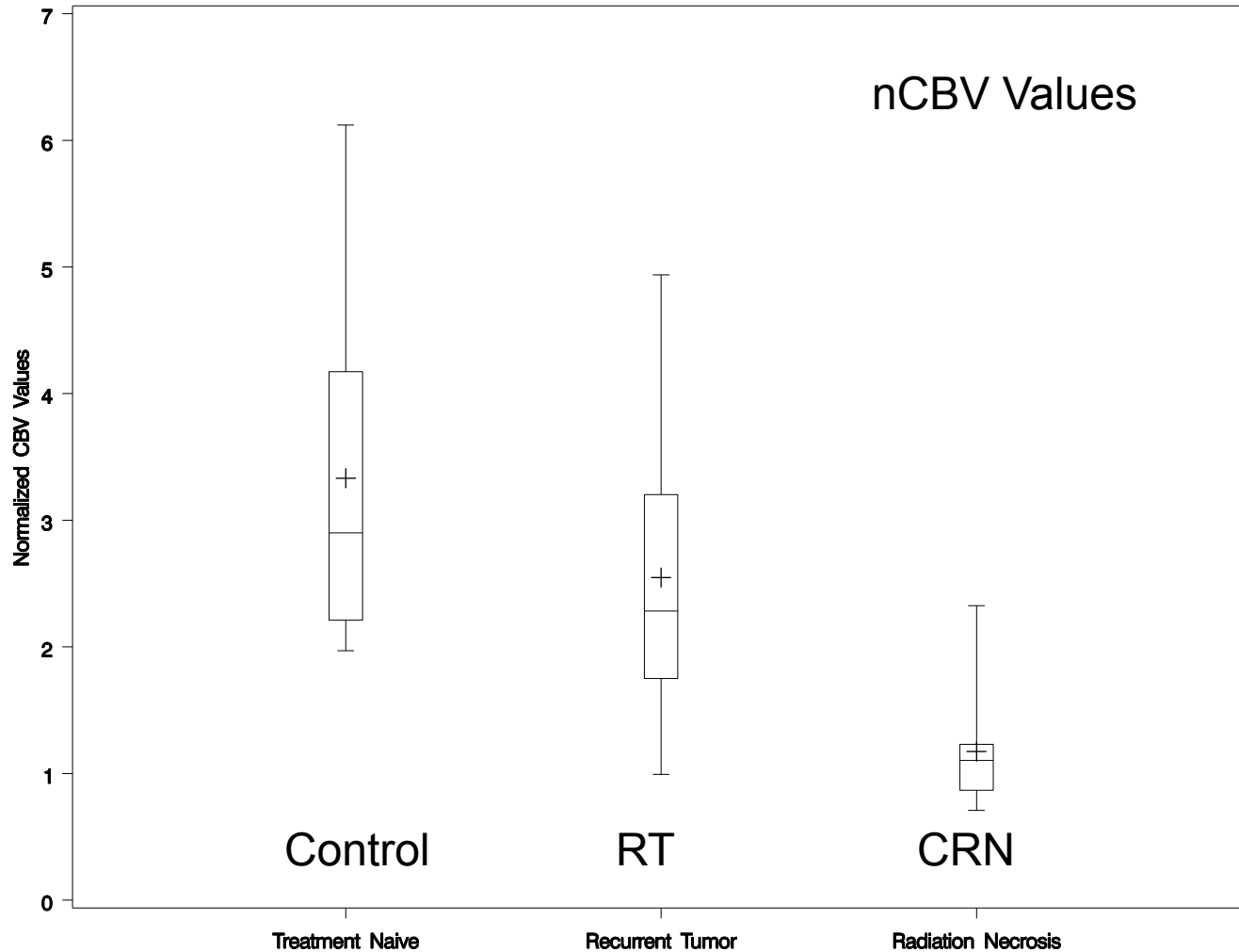
CBV map

# Perfusion CT :

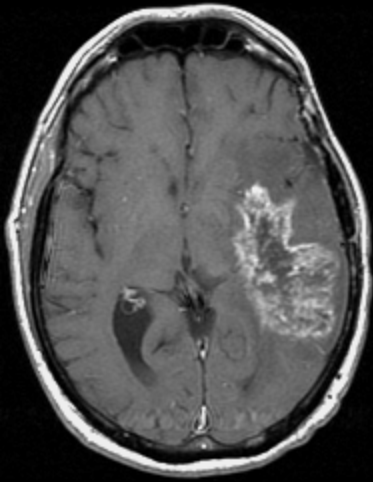
## Recurrent Tumor vs. Radiation Necrosis

	nCBV Mean (SD)	nCBF Mean (SD)	nMTT Mean (SD)
Recurrent Tumor (RT)	<b>2.54 (0.22)</b>	<b>2.63 (0.34)</b>	<b>1.02 (0.09)</b>
Cerebral Radiation Necrosis (CRN)	<b>1.17 (0.15)</b>	<b>0.97 (0.08)</b>	<b>1.41 (0.09)</b>
<b>p-values RT vs. CRN</b>	<b>&lt;0.0001</b>	<b>&lt;0.0001</b>	<b>&lt;0.0042</b>

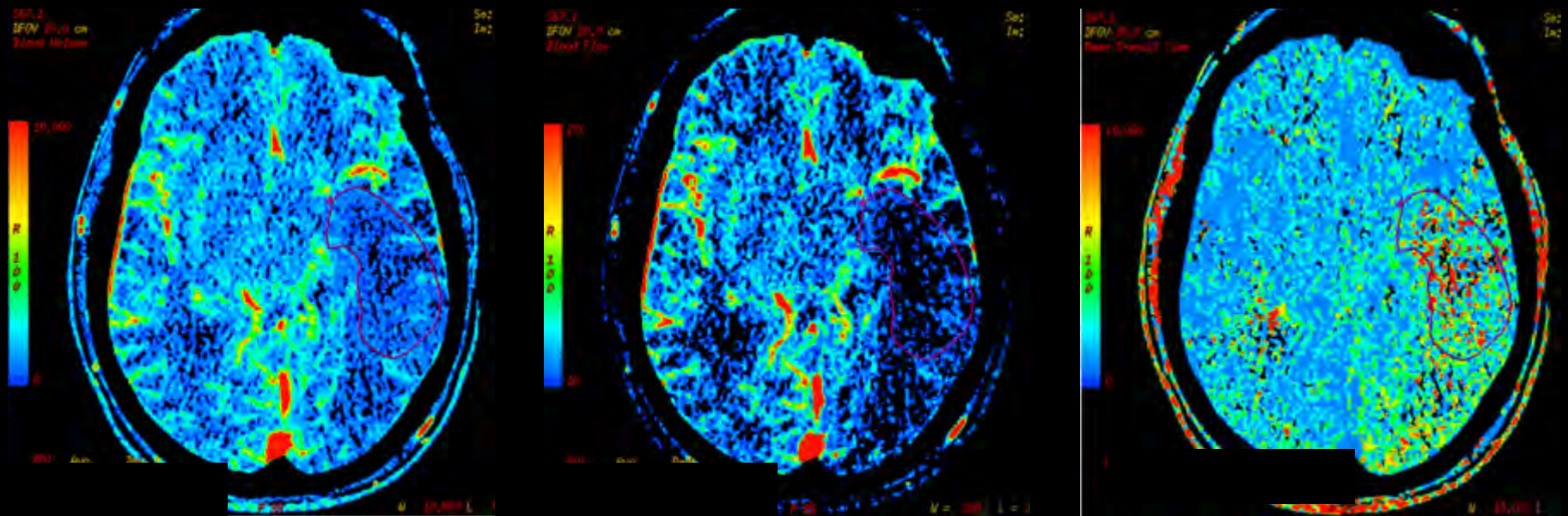
# Perfusion CT Imaging: Recurrent Tumor vs. Radiation Necrosis



# Cerebral Radiation Necrosis



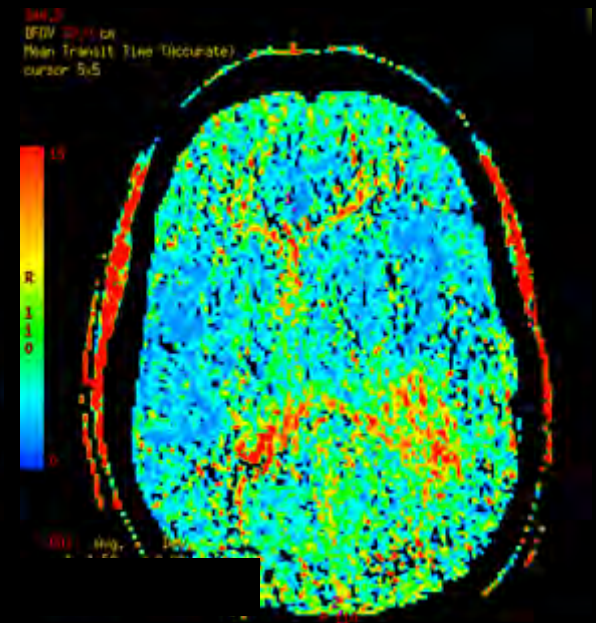
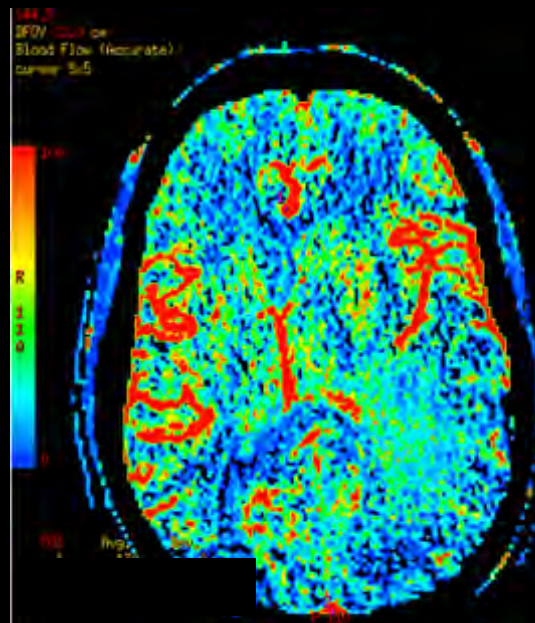
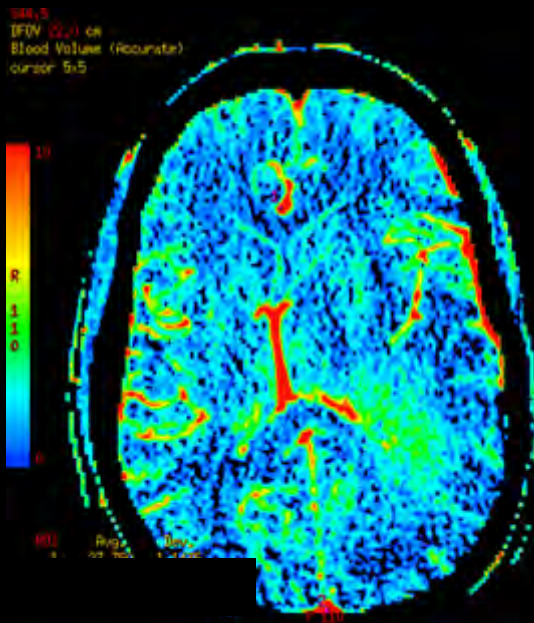
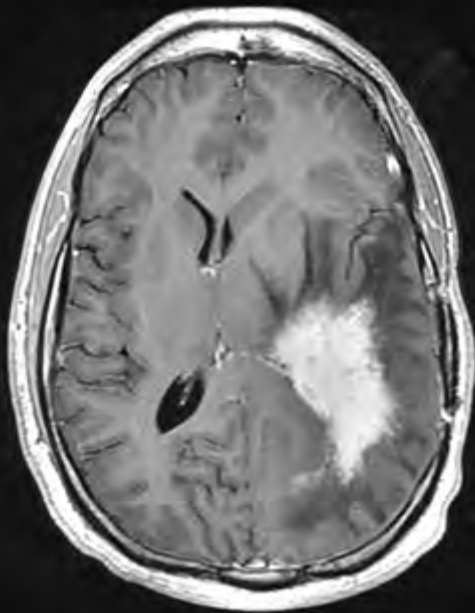
49 yo male with left temporal lobe anaplastic astrocytoma presented with a recurrent enhancing lesion 13 months after radiation therapy.



**Low CBV, low CBF and high MTT consistent with cerebral radiation necrosis.**

# Recurrent Tumor

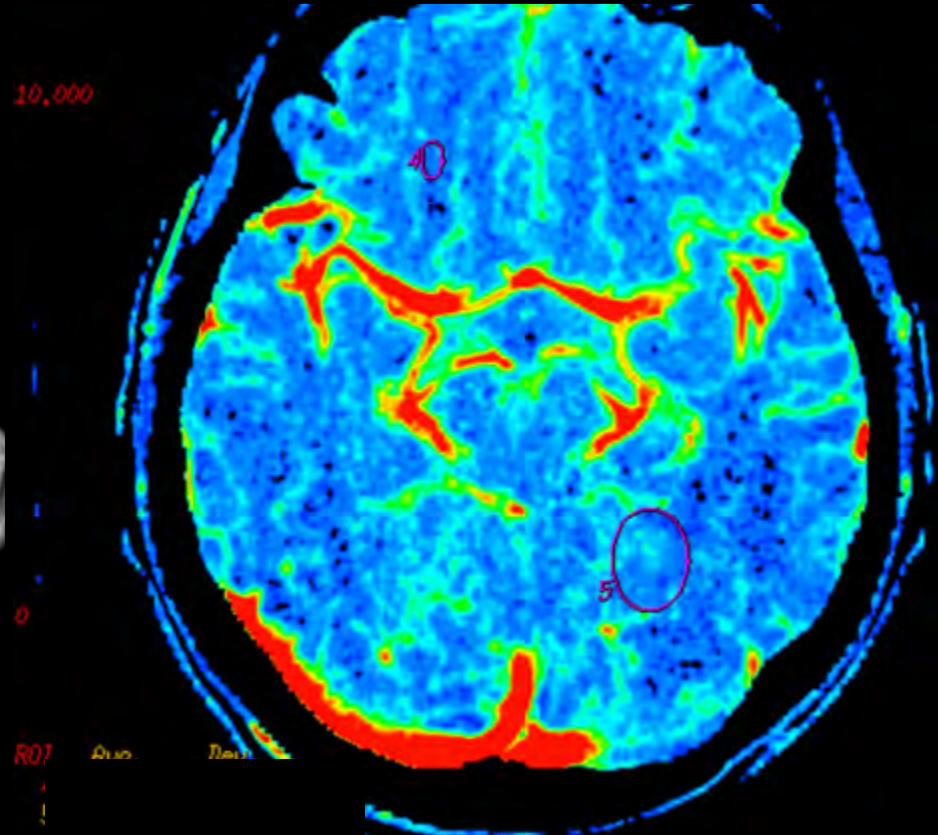
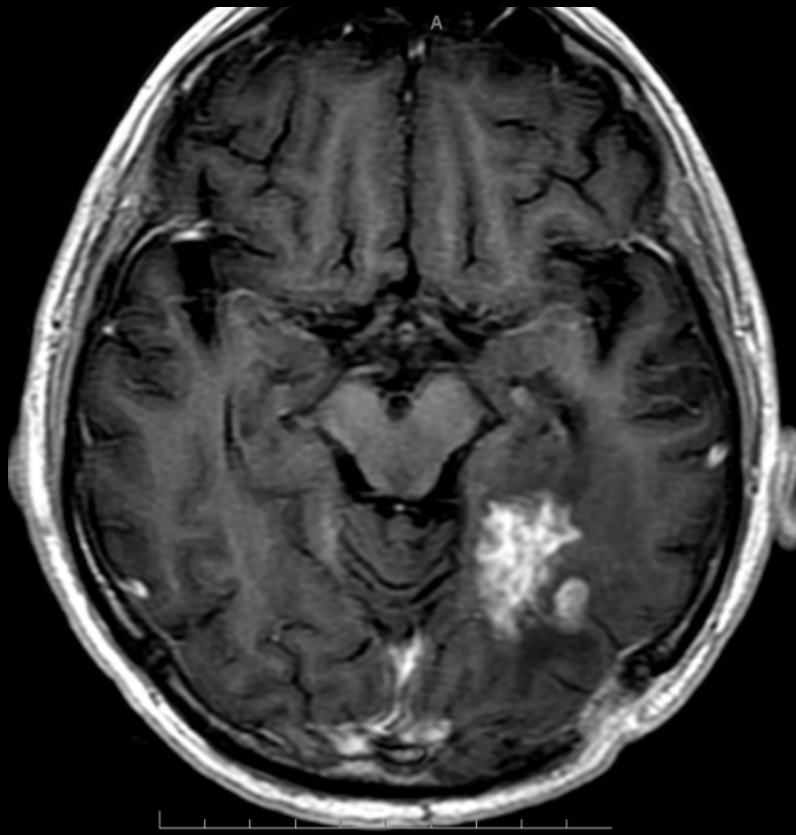
21 yo male with a left posterior temporal lobe astrocytoma 24 months after radiation therapy presenting with a recurrent enhancing lesion



High nCBV, high nCBF and lower nMTT suggestive of recurrent tumor

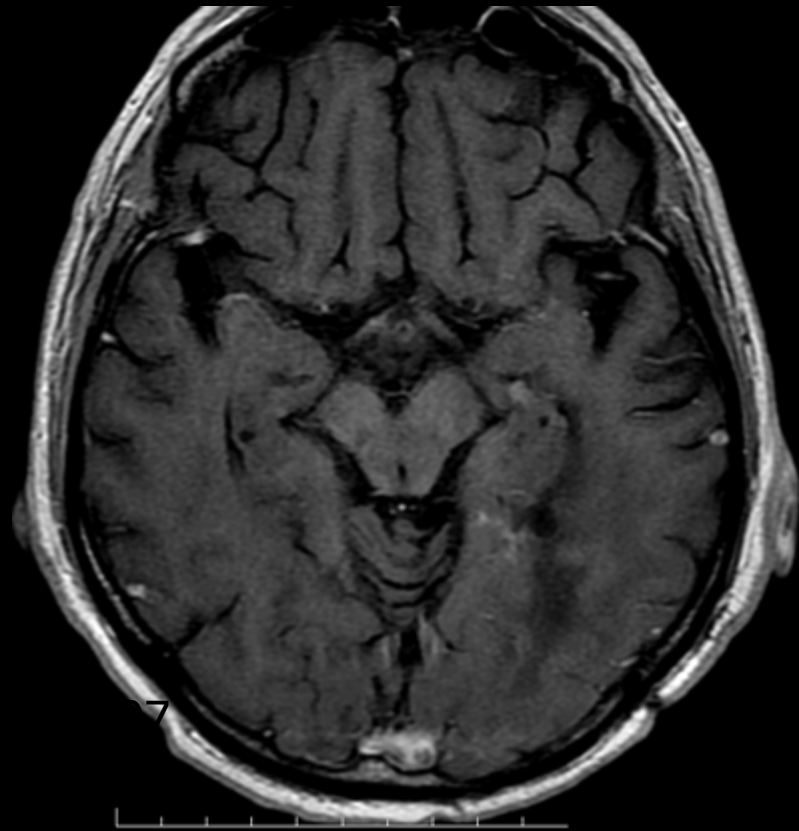
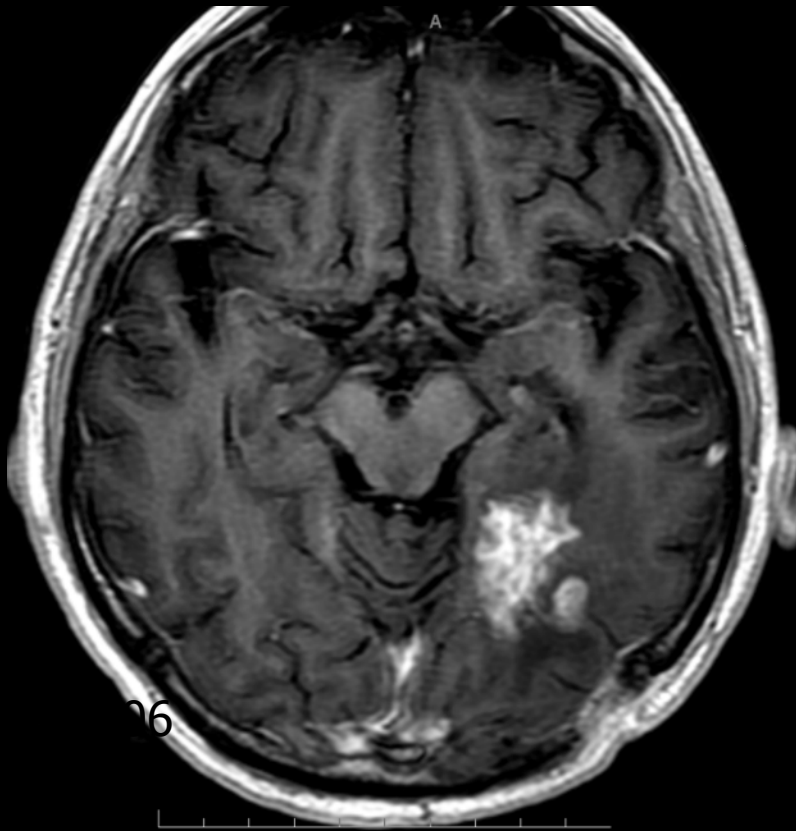
# Cerebral Radiation Necrosis

50 yo male post radiation therapy for lung carcinoma metastases.  
Lesion has low CBV suggesting cerebral radiation necrosis.



# Cerebral Radiation Necrosis

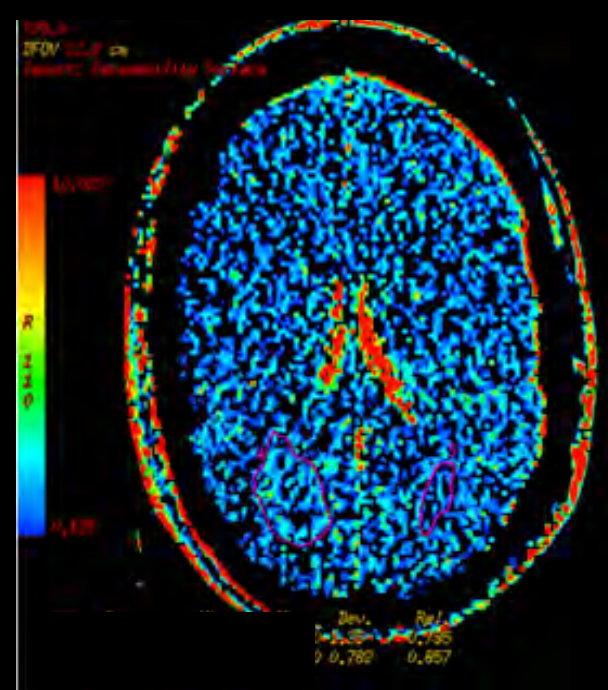
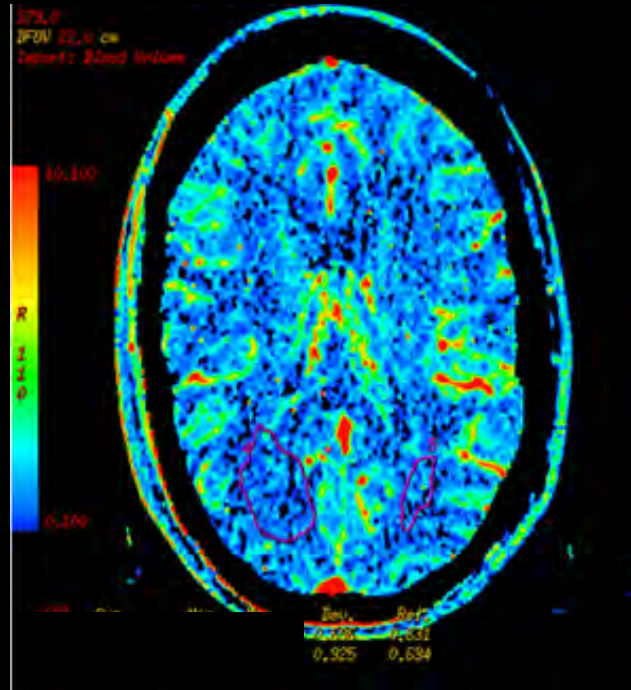
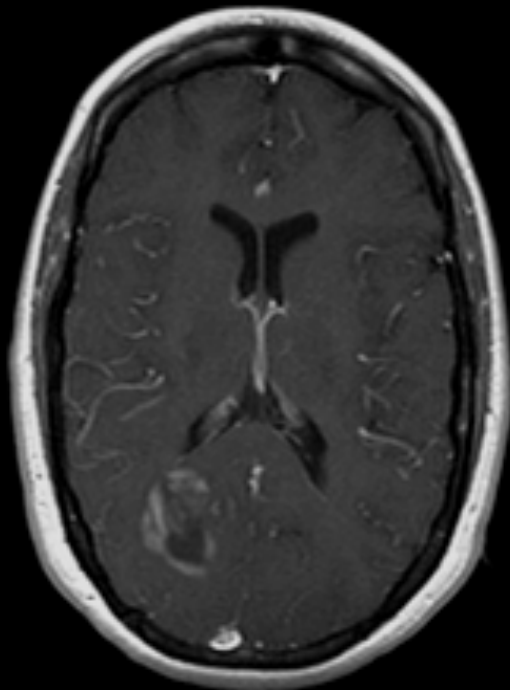
**Patient was treated supportively with Vitamin E and Trental, and no anti-neoplastic treatment. 8 month follow-up MR shows resolution of the lesion confirming radiation necrosis.**





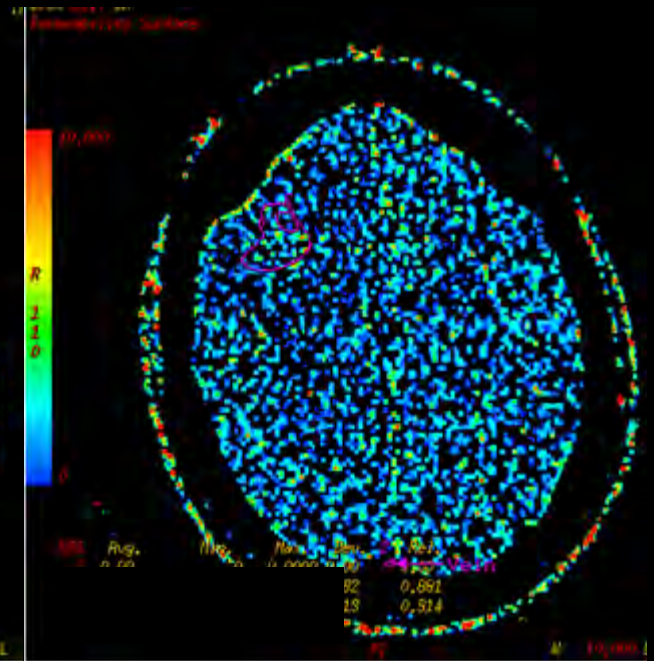
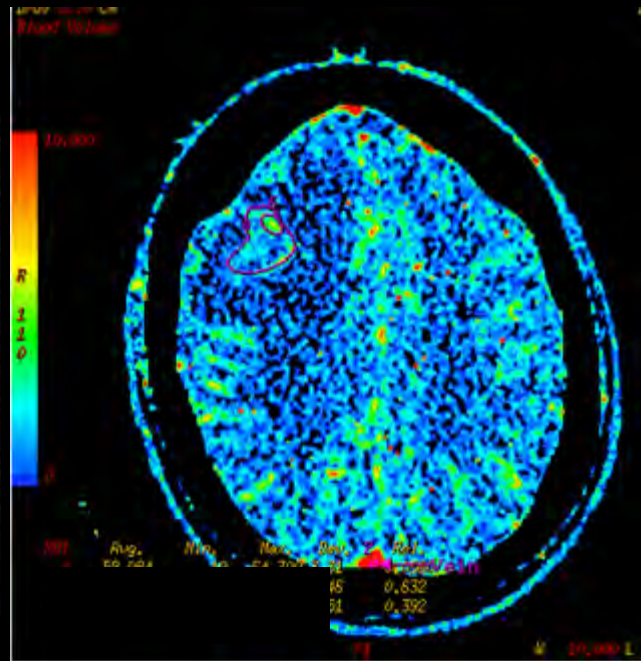
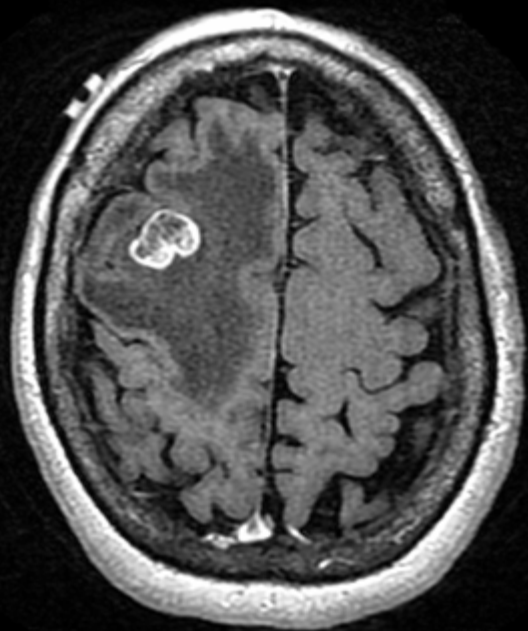
# 38 yo female with dizziness and headache

Low CBV suggested tumefactive MS rather than a glioma. Biopsy revealed this to be a demyelinating lesion.



# 63 yo female with history of multiple sclerosis and a lung mass

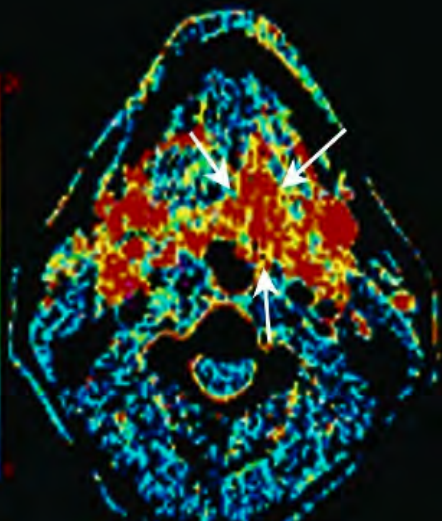
High CBV and low PS suggested neoplasm rather than TDL. Biopsy showed metastatic adenocarcinoma.



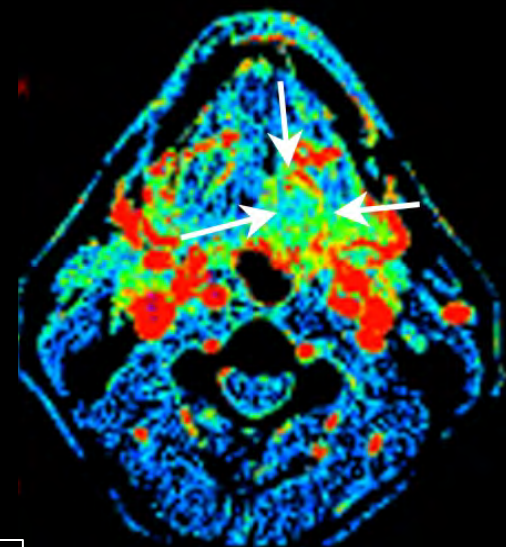
# Outline

- **Technique**
- **Clinical Applications**
  - **Stroke**
  - **Brain Tumors**
  - **Head & Neck**

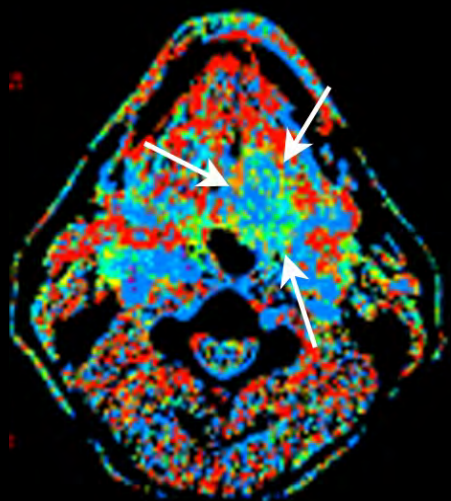
# CT Perfusion



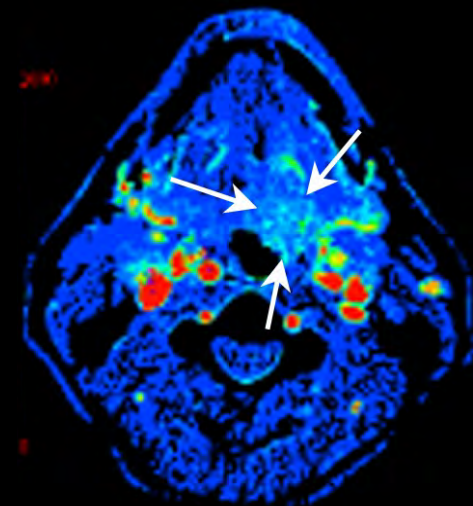
Capillary Perm



Blood Volume



Mean Transit Time



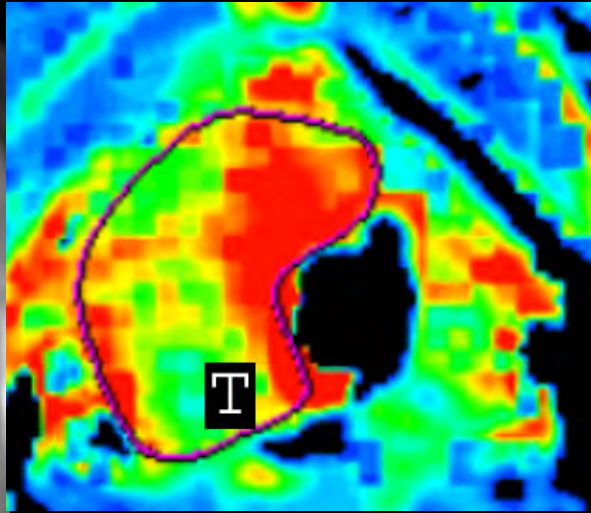
Blood Flow

HNSCCA vs Normal Muscle

- ↑ CP
- ↑ BF
- ↑ BV
- ↓ MTT

# CTP Perfusion vs Microvascular Density

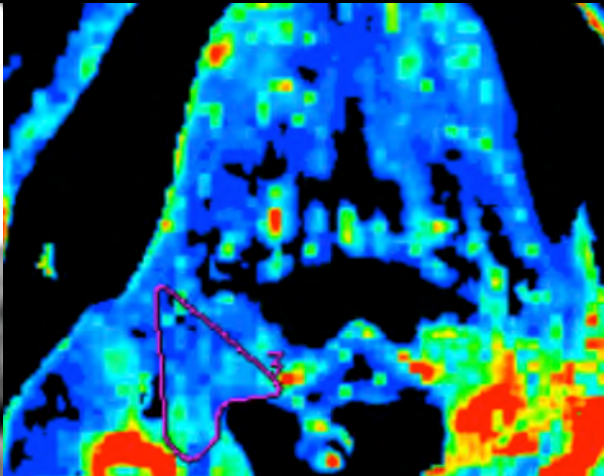
- Intratumoral microvessel density (MVD)
  - Marker of tumor angiogenesis
  - Prognostic indicator in head and neck squamous cell carcinoma (HNSCCA)
- Increased MVD
  - Advanced tumor stage
  - Locoregional and distant metastases
  - Reduced disease-free survival
  - Higher tumor oxygenation
- Requires endoscopic biopsy/tissue specimen



**45 yo male with stage IV**

**Tongue Base SCCA**

- Increased BF = 190.15 ml/100g/min
- Increased MVD = 47.2 vessels/mm<sup>2</sup>



**57 yo female with stage III**

**Tongue Base SCCA**

- Decreased BF = 39.13 ml/100g/min
- Decreased MVD = 19.2 vessels/mm<sup>2</sup>

*Ash et al. Radiology 2009;251:422-428*

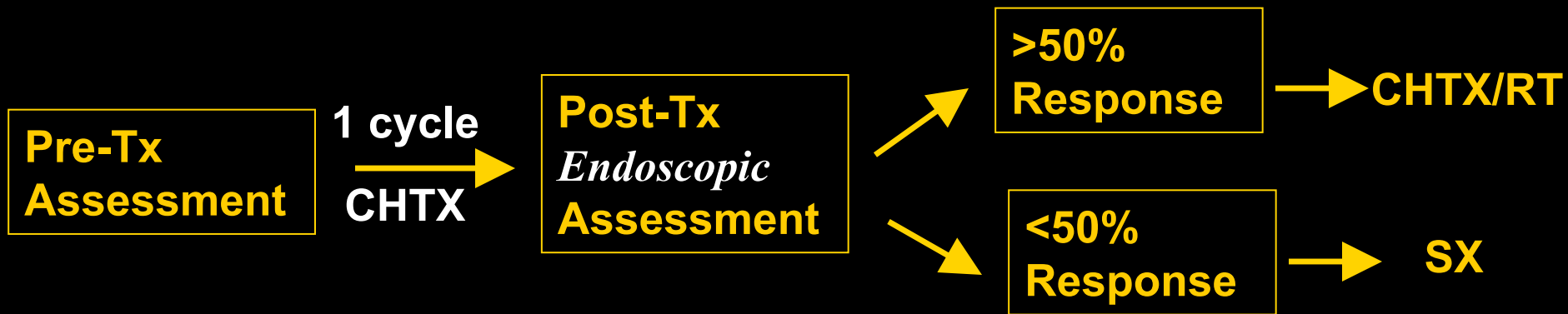
# CT Perfusion

## Microvascular Density

- Positive correlation
  - MVD & BF
  - MVD & BV
- No correlation
  - MVD & MTT
  - MVD & CP

# Clinical Applications

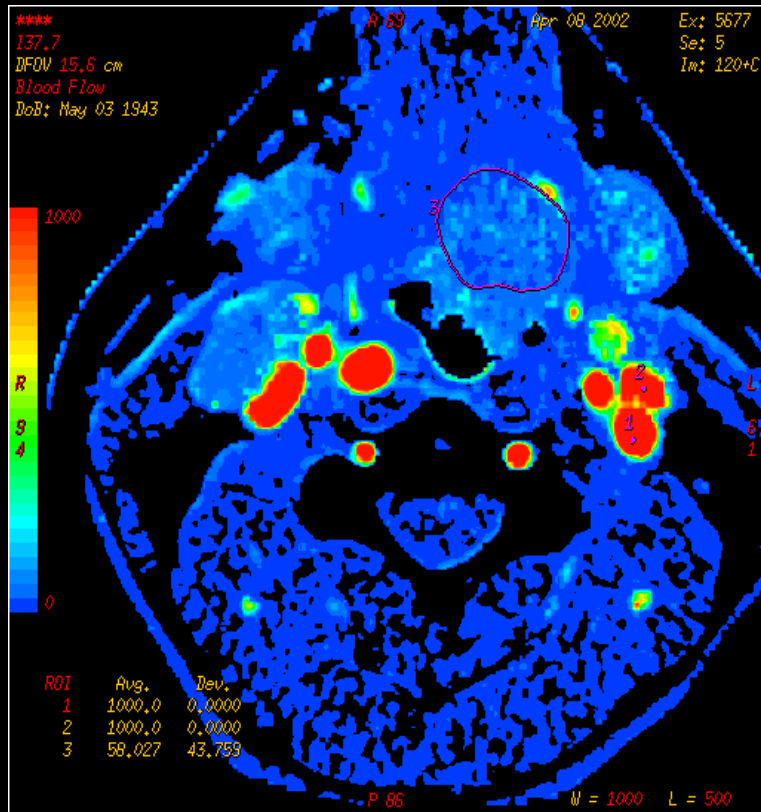
## *Neoadjuvant Protocol*



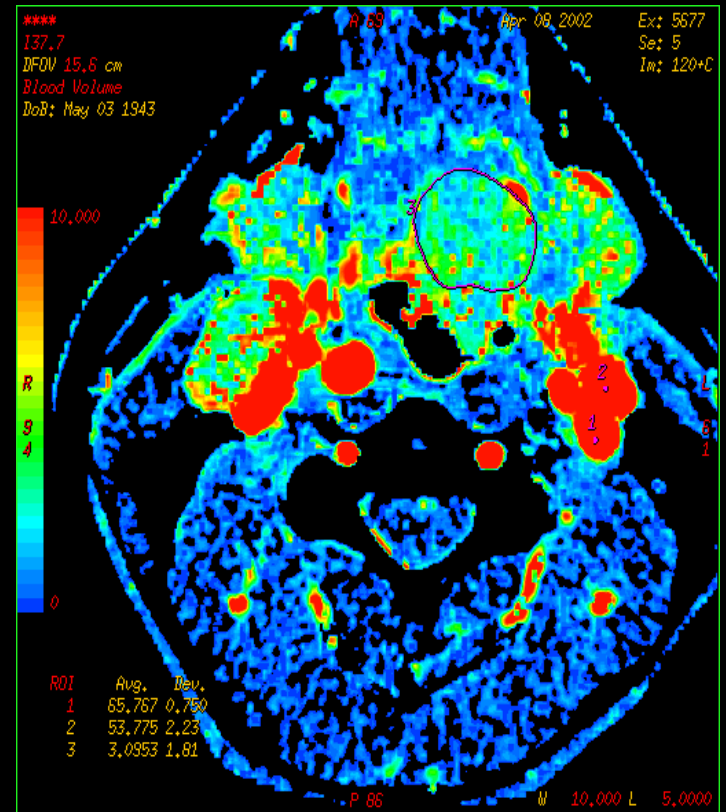


# CT Perfusion

## *Pre-treatment Parameters*



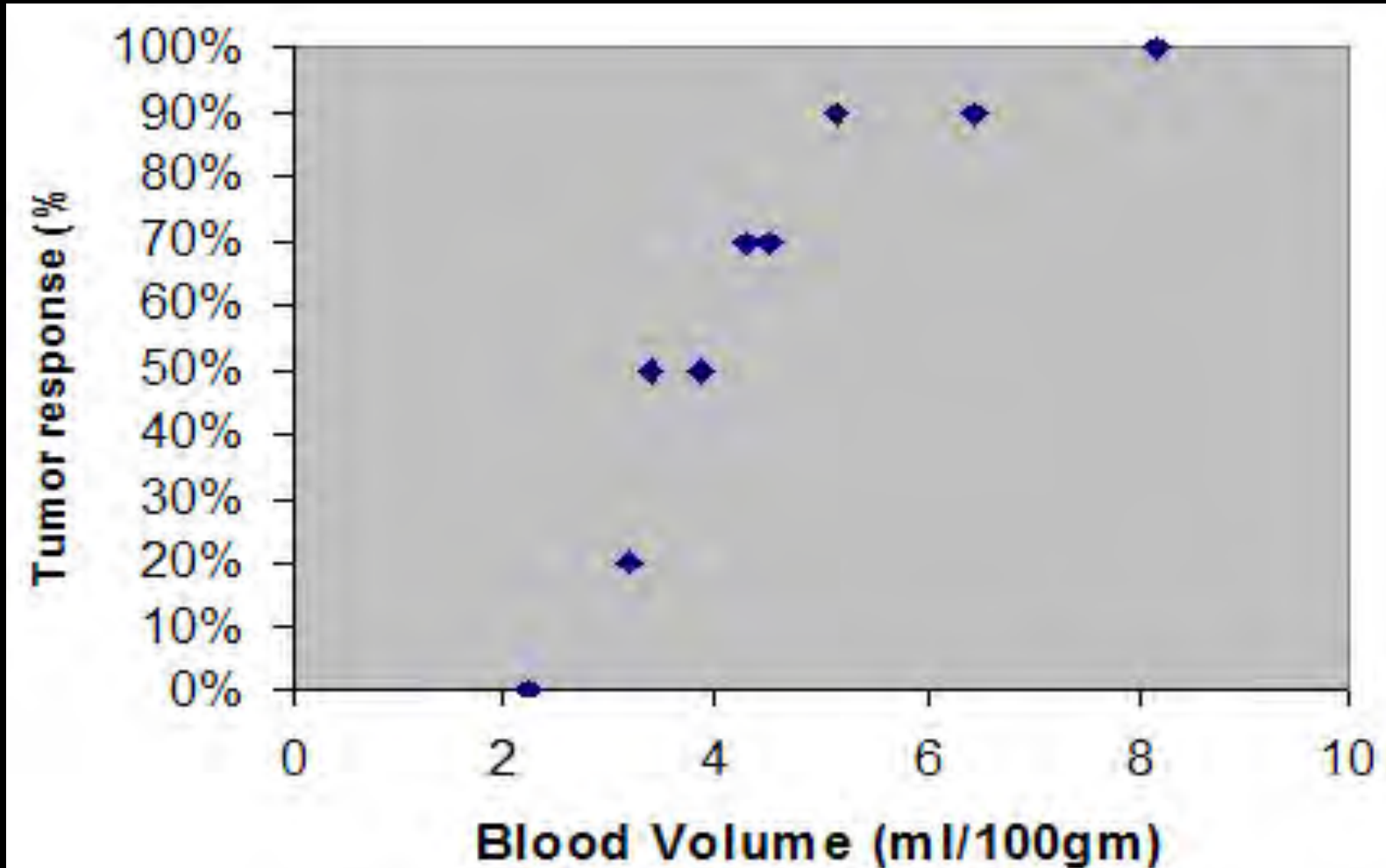
Blood Flow



Blood Volume

# CT Perfusion

*Pre-treatment Parameters: Blood Volume*



# CT Perfusion

## *Pre-treatment Parameters*

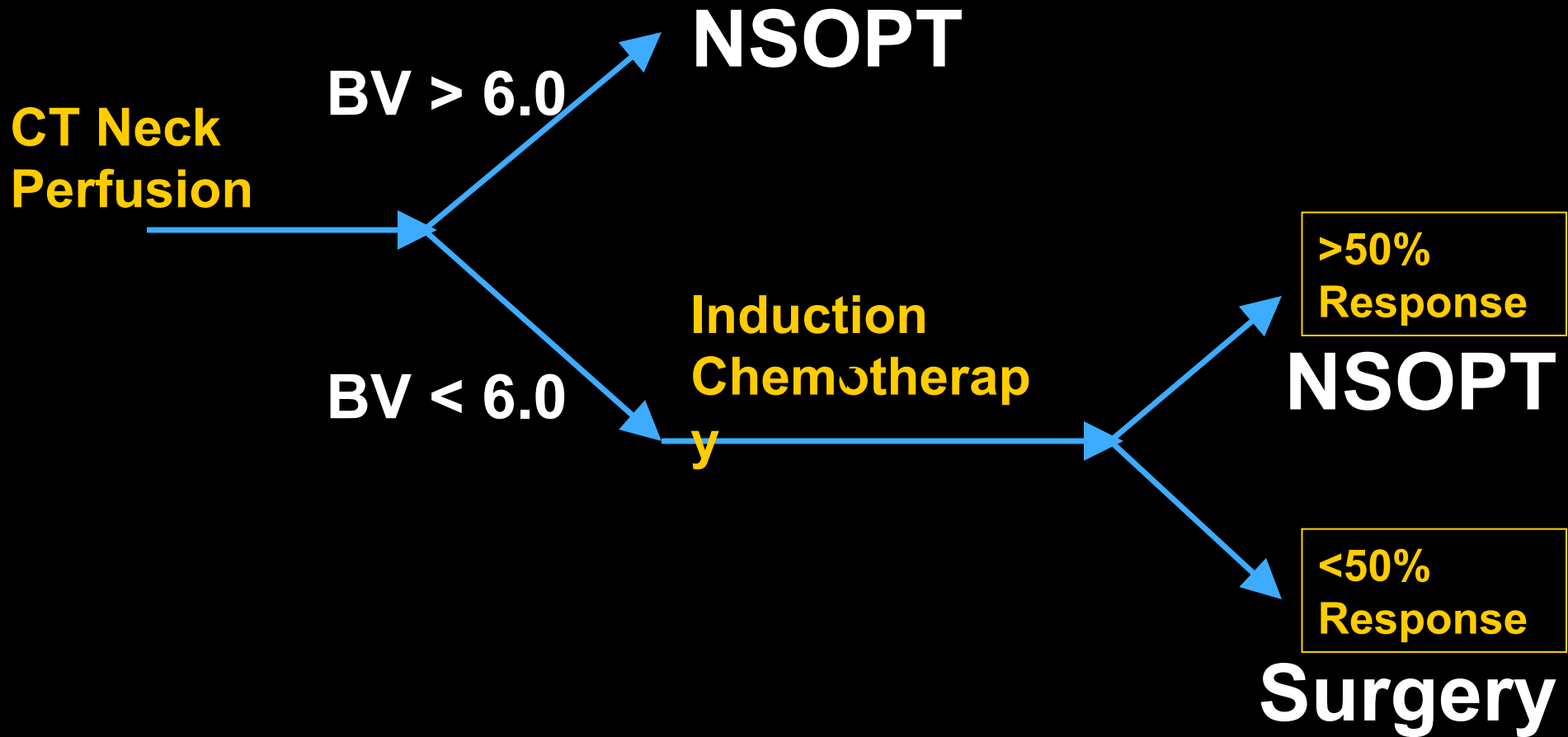
- Pretreatment values of BV & BF were significantly correlated to >50% reduction in tumor size following induction therapy.
- All patient with blood volume greater than 6 mg/dl successfully responded to induction therapy.

	p-value
BF	0.03
BV	0.004
MTT	0.29
CP	0.07

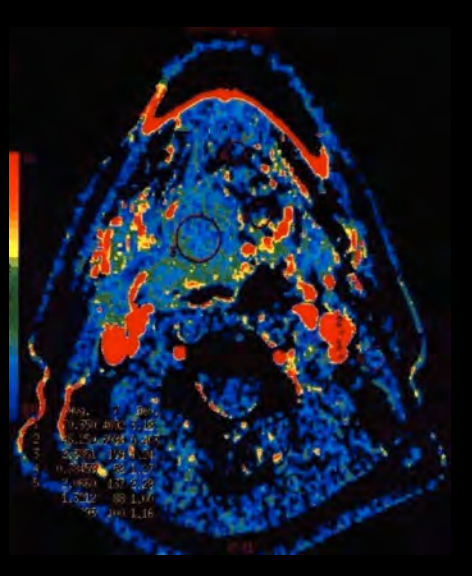
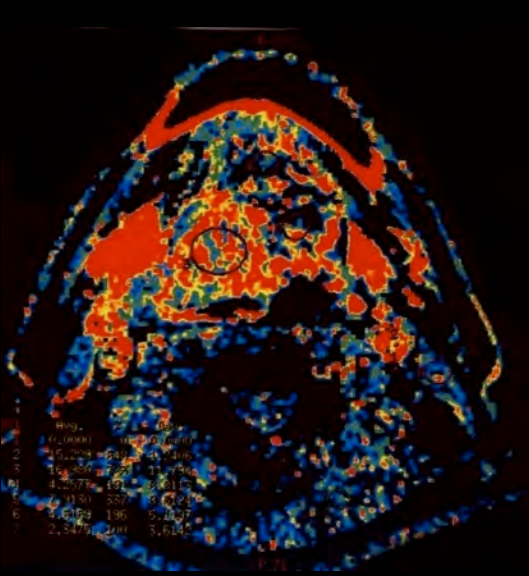
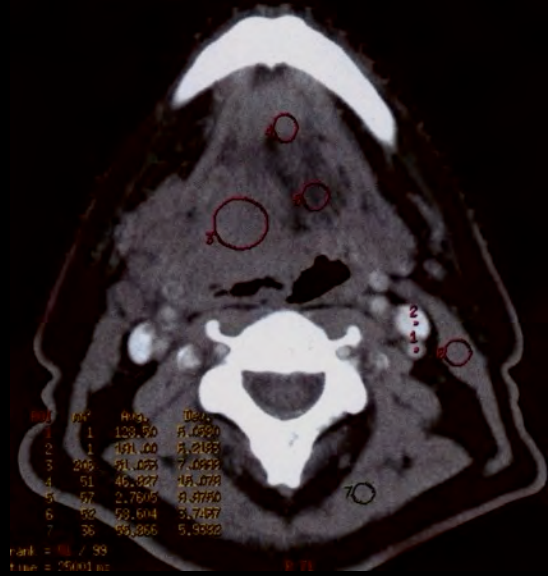
*Zima et al. AJNR 2007;28:328-334*

*Bisdas et al. AJNR 2009;30:793-799*

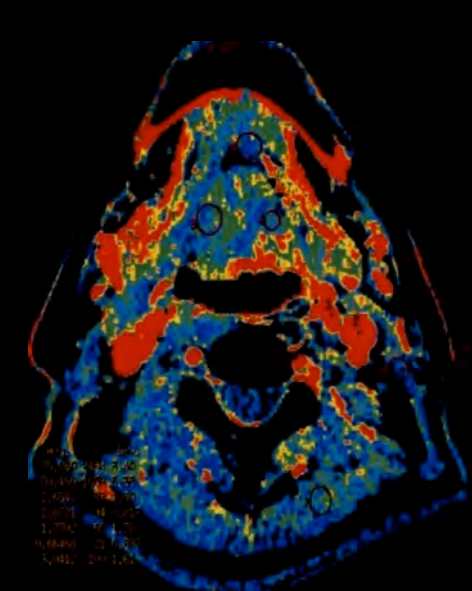
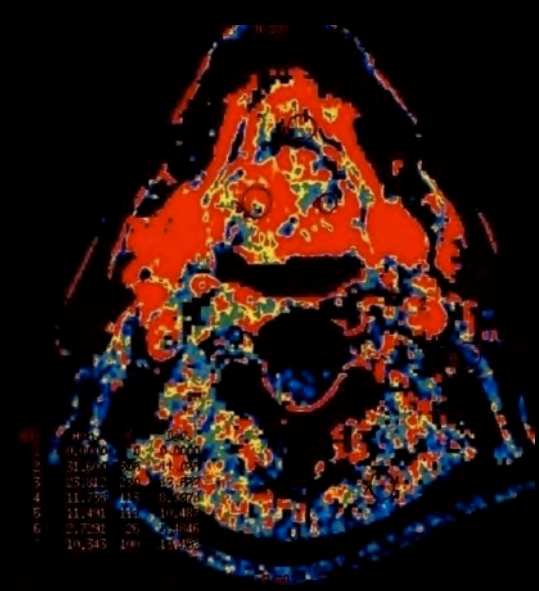
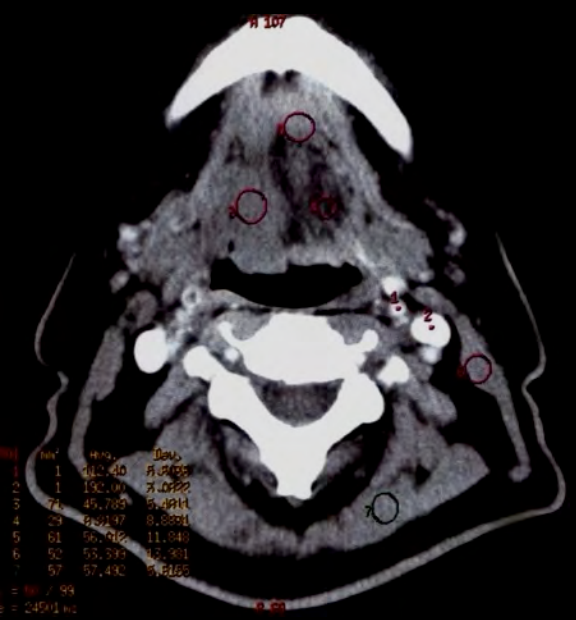
# Proposed Organ Preservation Therapy Treatment Plan



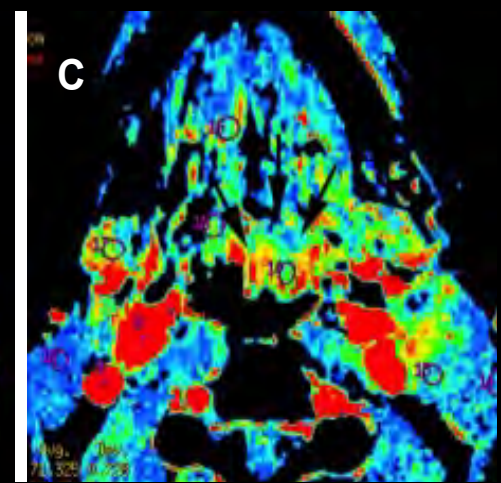
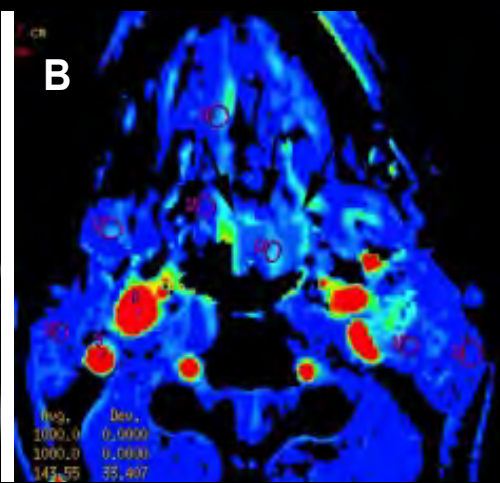
Pre-Tx



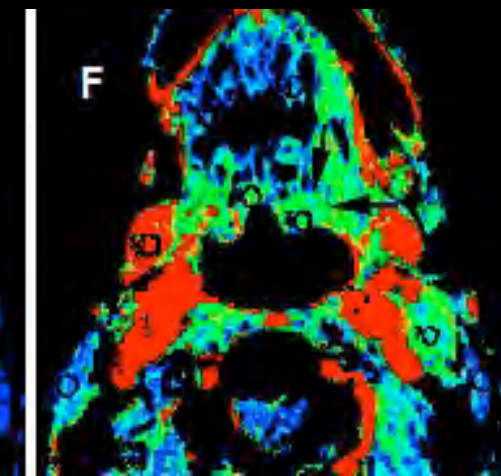
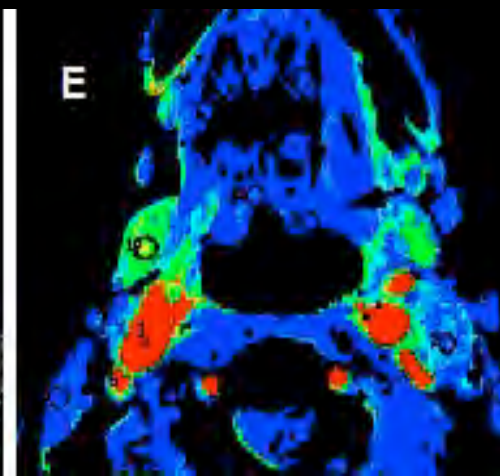
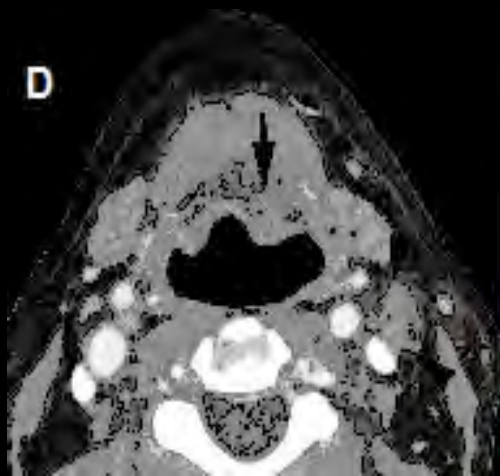
1 cycle



Pre-Tx



1  
cycle



# CTP vs Clinical Response

## Neoadjuvant Therapy

<u>CTP Parameter</u>	<u>Kappa Value</u>
<b>Blood Volume</b>	<b>0.73</b>
Blood Flow	0.37
Capillary Perm	0.37
MTT	0.37

*Gandhi et al. AJNR 2006;27:101-106*

# CTP vs Clinical Response

## Concomitant Therapy Monitoring

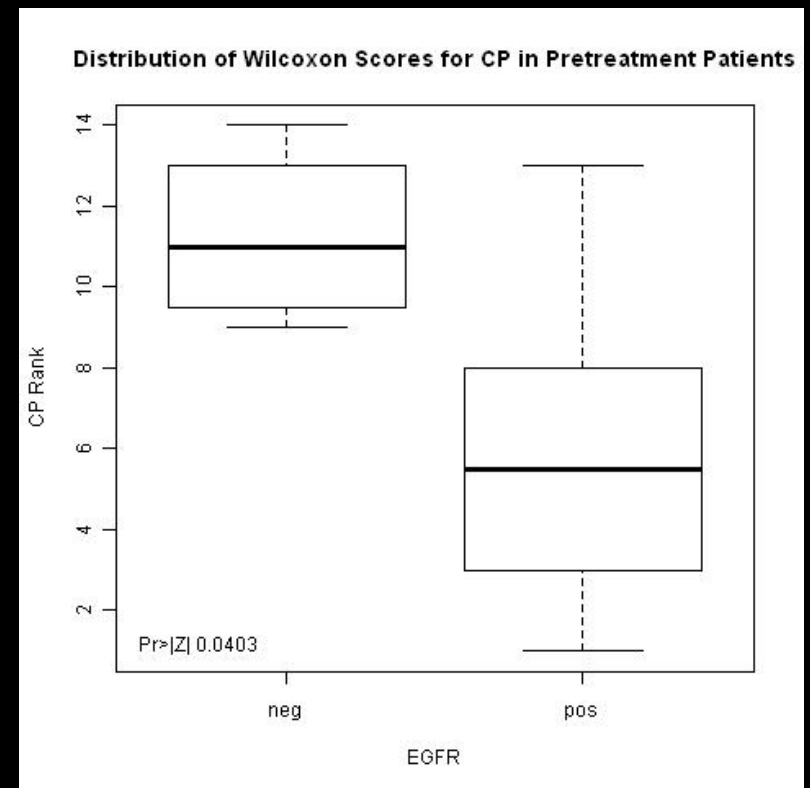
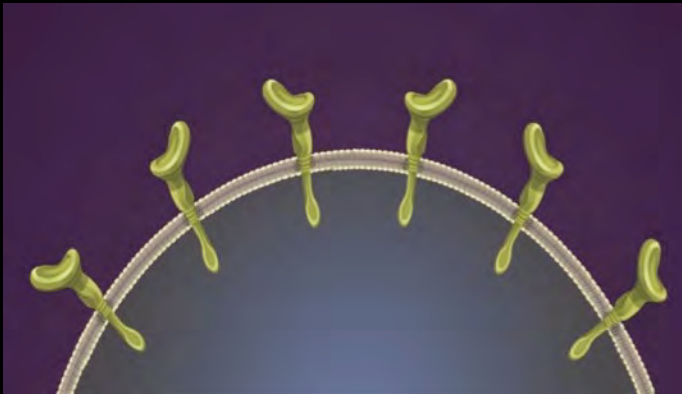
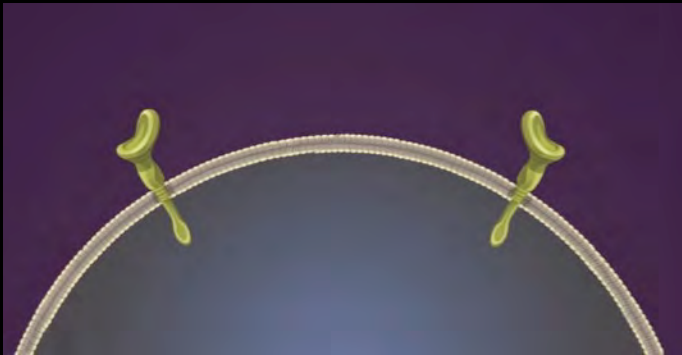
### 40y & 70gy

- Decreased BV suggests responders (40Gy)
- No change or increase BV indicates non-responders

*Surlan-Popovic et al. AJNR 2010;31:570-575*



# Correlation with EGFR Biomarker



# Outline

- **Technique**
- **Clinical Applications**
  - **Stroke**
  - **Brain Tumors**
  - **Head & Neck**
- **Radiation Dose Update**

# **FDA Alert: 10/8/09**

**At least 206 patients in an 18-month period received extremely high radiation doses during perfusion CT imaging. Patients were expected to receive a dose of 0.5 Gy (max) to their head but instead received 3-4 Gy.**

**Resulted in hair loss and skin erythema.  
Possibility of long term effects**

**CT unit had been set at incorrect levels for 18 months, after the hospital made an error while reconfiguring the scanner.**

# **FDA Alert: 12/8/09**

- **FDA has identified at least 50 additional patients who were exposed to excess radiation during CT perfusion scans**
- **Cases involved more than one CT vendor**
- **If patient doses are higher than the expected level, but not high enough to produce obvious signs of radiation injury, the problem may go undetected and unreported, putting patients at increased risk for long-term radiation effects including cataracts.**

# **FDA Alert: 12/8/09**

- 1. Assess whether patients who underwent CT perfusion scans received excess radiation**
- 2. Review radiation dosing protocols for all CT perfusion studies to ensure that the correct dosing is planned for each study**
- 3. Implement quality control procedures to ensure that dosing protocols are followed every time and the planned amount of radiation is administered.**
- 4. Technologists check the CT scanner display panel before performing a study to make sure that amount of radiation to be delivered is at the appropriate level for the individual patient.**
- 5. If more than one study is performed on a patient during one imaging session, practitioners should adjust the dose of radiation so it is appropriate for each study**



level, causing a **CT radiation overdose**. Now, patients who were affected by the excess CT scan radiation have filed lawsuits against Cedars-Sinai and the makers of the CT machine, GE Healthcare Inc. and GE Healthcare Technologies.

#### FREE CASE EVALUATION

Send your CT Brain Scan Radiation claim to a Lawyer who will review your case at **NO COST** or obligation.

[CLICK HERE](#)

#### CT Brain Perfusion Scan Radiation Overdose

According to an FDA alert, at least 206 patients in an 18-month period received extremely high radiation doses when they were undergoing perfusion CT imaging. Specifically, the FDA says that the patients were expected to receive a dose of 0.5 Gy (maximum) to their head but instead received 3-4 Gy.



An article in *The New York Times* (October 15, 2009) notes that Cedars-Sinai Hospital disclosed that it had accidentally exposed patients to high doses of radiation. Meanwhile, in an incident unrelated to the FDA CT scan alert at Cedars-Sinai, at Mad River Community Hospital in Arcata, California, an x-ray technician has lost her state license for reportedly putting a 2½ year old through more than an hour of CT scans, according to the same article in *The New York Times*.

The FDA alert notes that some patients experienced hair loss and erythema (redness of the skin). However, also of concern is that overexposure to radiation that is not at radiation effects. Furthermore, there is a concern that the problem may be more widespread than just one facility.

\*While this event involved a single kind of diagnostic test at one facility, the magnitude of these

#### Exposure, Increases Oversight



Washington, DC: In the wake of a radiation scare where more than 300 patients undergoing **CT scans** and other medical procedures in four hospitals were given too much radiation, the US Food and Drug Administration (FDA) has announced it will step up oversight for the potentially dangerous procedure. [ [Read More](#) ]

#### Health scans seen as risky in wake of ct radiation overdose case



Miami, FL: Medical science has provided modern man with many tests to better facilitate preventative health care, yet some of these tests - notably the **CT brain scan** - may potentially be more harmful to the patient in the long run. [ [Read More](#) ]

#### CT Scan Overdose Victims Have Rights



Los Angeles, CA: It seems that somebody at Cedars-Sinai Medical Center made a serious error when the hospital's CT Scanner was recalibrated in February 2008. Dozens of patients booked for diagnostic brain scans were blasted with **megadoses of radiation** - doses usually reserved for treating brain tumors. [ [Read More](#) ]

#### MORE CT BRAIN SCAN RADIATION

## CT Brain Scan Radiation Overdose Legal Help

If you or a loved one has suffered an adverse health event resulting from CT brain perfusion scanning, please click the link below and your complaint will be sent to a lawyer who may evaluate your claim at no cost or obligation.

[Please click here for a free evaluation of your CT Brain Scan Radiation case](#)

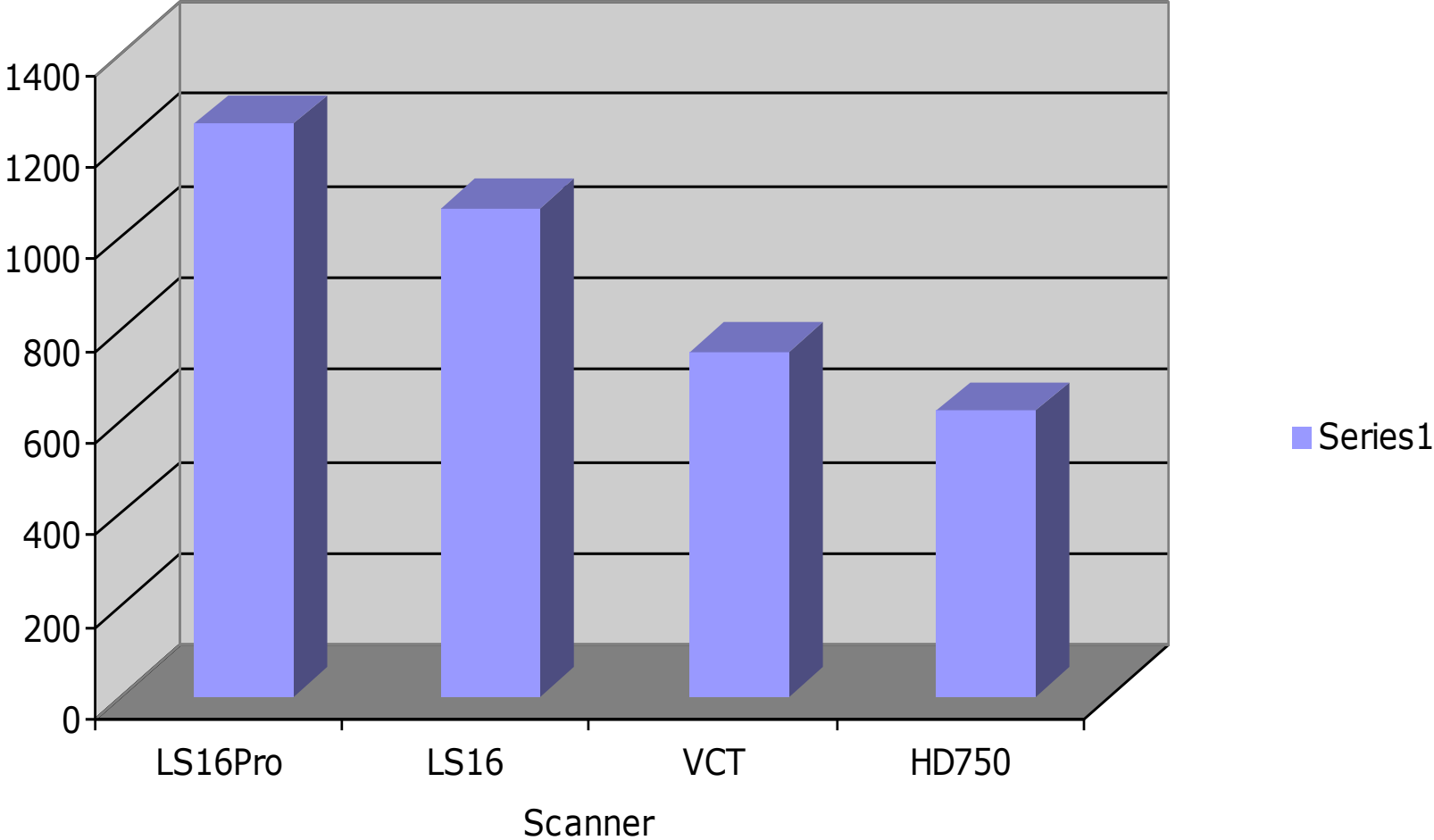
# **Lawsuit in Alabama CT Perfusion Case:**

## **12/15/09**

**Attorneys in Huntsville, AL, have filed suit in federal court on behalf of a patient who allegedly received excessive radiation during a CT perfusion head scan for suspected stroke.**

**The lawsuit represents more than 300 patients, including many of the 260 patients who allegedly received CT overdoses at Cedars-Sinai Medical Center in Los Angeles.**

**Mean Skull Base + Neck DLP (mG)**





<b>Scanner #</b>	<b>Measured head phantom (mGy)</b>	<b>Randon patient head dose from scanner display (mGy)</b>	<b>Acceptable dose range for brain perfusion according to FDA (mGy)</b>
<b>CT 2</b>	<b>387</b>	<b>309</b>	<b>&lt;500</b>
<b>CT 4</b>	<b>355</b>	<b>317</b>	<b>&lt;500</b>
<b>CT 6</b>	<b>389</b>	<b>311</b>	<b>&lt;500</b>

# FDA requirement for perfusion study: less than 500 mGy

## 120 mm acute stroke volume shuttle

0.4 s rotation  
40 mm detector coverage  
5 mm thickness  
pitch 0.984:1, 39.37 mm/rot  
coverage time 45.7 s  
27 shuttle passes  
80 kV  
490 mA

CTDI= 617.30 perfusion

CTDI= 45.92 non-contrast head

Total CTDI **663.22** mGy

300 mA

CTDI=378 perfusion

Total CTDI 424 mGy

350 mA

CTDI=441 perfusion

Total CTDI 487 mGy

## 80 mm shuttle axial

0.4 s rotation  
40 mm detector coverage  
5 mm, 8i  
coverage time=46.6s  
17 passes  
80 kV  
500 mA

CTDI= 222.57 perfusion

CTDI= 45.92 non-contrast  
head

Total CTDI 268.49 mGy

## 40 mm cine

1.0 s rotation  
40 mm detector coverage  
5 mm, 8i  
coverage time=50s  
80 kV  
200 mA

CTDI= 654.76 perfusion

CTDI= 45.92 non-contrast  
head

Total CTDI **700.68** mGy

# Outline

- **Technique**
- **Clinical Applications**
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  - **Brain Tumors**
  - **Head & Neck**
- **Radiation Dose Update**