

# BRIGHAM AND WOMEN'S HOSPITAL



## Pregnancy and Stroke



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## **Disclosures**

I have no financial relationships with the developers of any of the products discussed.

#### **NINDS**

- SPOTRIAS
- NeuSTART
- IRIS (and Takeda Pharmaceuticals)
- ATACH II
- POINT
- StrokeNET
- DEFUSE-3
- ARCADIA

#### Covidien

• SWIFT PRIME

## **Topics**

- I. Importance of stroke in pregnancy
- II. Pathophysiology of stroke in pregnancy
- III. Cerebral venous thrombosis
- IV. Acute ischemic stroke
- V. Subarachnoid hemorrhage
- VI. PRES and RCVS

#### **Morbidity of Stroke in Pregnancy and the Puerperium**

Healthcare Cost and Utilization Project of the Agency for Healthcare Research and Quality N = 9 million discharges 2000-2001

- Stroke Rate 34 per 100,000
- Mortality Rate 1.4 per 100,000 = 4.1 %

#### **Disability estimates**

- Long-term disability in ~2/3 survivors, greater in women
- Depression in 11-68%
- Major depression in 10-27%

James AA Obstet Gynecol Survey 2006;61:4-5. Bousser M-G Circulation 1999;99:463

### **Leading Causes of Death in Adolescents and Young Adults**

| Cause of Death         | Rate per 10⁵<br>Age 15-24 | Rate per 10⁵<br>Age 25-34 |
|------------------------|---------------------------|---------------------------|
| Accident               | 6.0                       | 37.5                      |
| Homocide               | 0.8                       | 13.0                      |
| Suicide                | 10.0                      | 12.4                      |
| Cancer                 | 4.1                       | 9.0                       |
| Heart disease          | 2.7                       | 8.1                       |
| Diabetes mellitus      | 0.5                       | 1.5                       |
| Cardiovascular disease | 0.5                       | 1.4                       |

## **Summary of Risks**

#### Non-pregnancy women of childbearing age

• The annual risk of stroke in non-pregnant women ages 15-44 is low (10/10<sup>5</sup>). However, the risk may be rising (Ban 25/10<sup>5</sup>). (up to age 49)

#### **Pregnancy-associated**

42% of strokes in women 15-44 y/o are associated with pregnancy.

#### All Strokes

- Considering ALL STROKES (ICH + AIS): There is a small increase in ALL STROKES during pregnancy driven by hemorrhagic stroke.
- There is a marked increase in ALL STROKES during the early postpartum period.

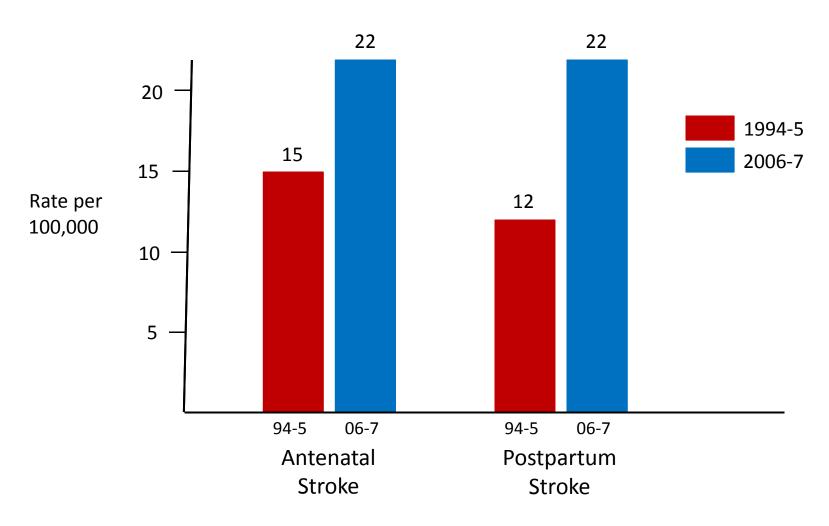
#### **During pregnancy**

- There is no increase in the risk of ISCHEMIC STROKES during pregnancy. (Ban found slight decrease.)
- There is an increase in the risk of HEMORRHAGIC STROKES during pregnancy (Kittner 2.5-fold, Ban increased SAH)

#### During the early postpartum period

- There is an increased risk of ISCHEMIC STROKES in the early postpartum period (up to 6 weeks). (Kittner 8.7-fold)
- There is an increased risk of HEMORRHAGIC STROKES in the early postpartum period (up to 6 weeks). (Kittner 28.5-fold, Ban increased ICH and SAH)

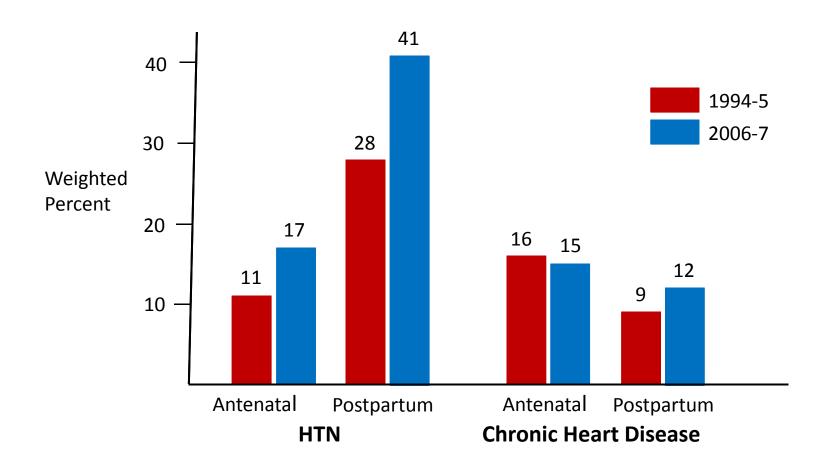
## Rate of Stroke in Pregnancy and the Puerperium Is It Increasing?



Healthcare Cost and Utilization Project of the Agency for Healthcare Research and Quality

Stroke 2011;42:2564

## Increase in Rate of Stroke Follows Increases in Rates of HTN and Chronic Heart Disease



Healthcare Cost and Utilization Project of the Agency for Healthcare Research and Quality

Stroke 2011;42:2564

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### **Relative Risk of Stroke in Pregnancy and the Puerperium**

| Stroke Type         | Relative Risk<br>of Stroke during<br>Pregnancy | Relative Risk of Stroke during the Puerperium (6 wk) |  |  |
|---------------------|--|--|--|--|
| Cerebral infarction | 0.7  | 8.7  |  |  |
| Cerebral hemorrhage | 2.5  | 28.5   |  |  |

Kittner N Engl J Med 1996;335:768-74.

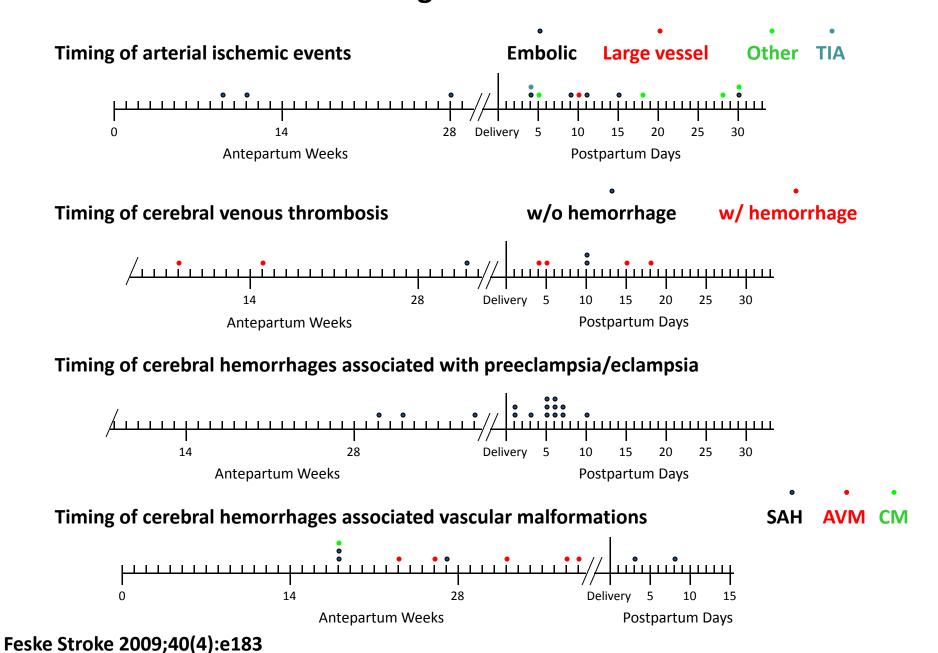
## **Major References: Epidemiology**

Sharshar T, et al. Incidence and causes of strokes associated with pregnancy and puerperium: A study in public hospitals of Ile de France. Stroke 1995;26(6):930-6.

Kittner SJ, et al. Pregnancy and the risk of stroke. N Engl J Med 1996;335(11):768-74.

Ban L, et al. Incidence of first stroke in pregnant and nonpregnant women of childbearing age: A population-based cohort study from England. J Am Heart Assoc 2017;6(4):e004601.

### **Timing of Events**

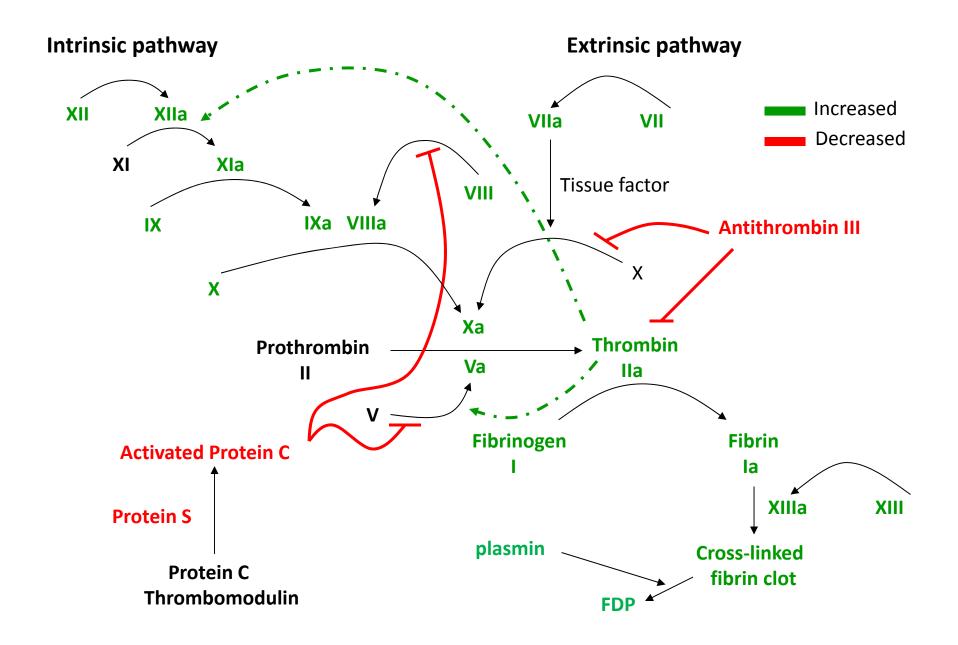


- Physical changes
  - Compression of the IVC
  - Compression of the aorta
  - Compression of uterine arteries and veins
  - Decreased venous compliance
- Increases in procoagulant factors
  - Increase in factors I, VII, VIII, IX, X, XII, and XIII
  - No change in factors II, V, XI
- Decreases in coagulation inhibitors
  - Decreased AT III
  - Decreased protein S
  - Functional protein C resistance
- Thrombin generation and fibrinolysis
  - Increased thrombin generation
  - Increased fibrinogen and fibrinolysis
  - Platelet consumption

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### Acute Hemorrhage is a Thrombophilic State!

- Thrombosis and hemorrhage are well-known complications of trauma
- Increased high molecular weight fibrinogen after delivery as part of the acute phase reaction
- The mild "DIC" state:
  - Increased thrombin generation
  - Increased fibrinolysis
    - increased FDP
    - increased D-dimer
    - Fibrinogen consumption
  - Platelet consumption

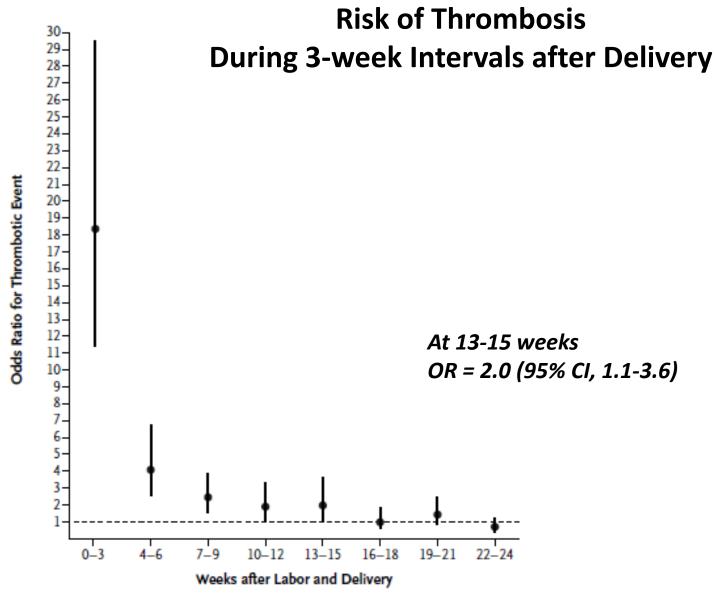
### **Postpartum Thrombophilia**

1,687,930 Californian women hospitalizations for delivery from Jan 2005 to June 2010

Thrombotic events: Stroke, MI, VTE

#### Risk factors

- Older
- White or African American v Hispanic or Asian
- No private insurance
- Other risk factors for thrombosis
  - Age > 35 yr
  - Eclampsia
  - Primary hypercoagulable state
  - Smoking
  - Cesarean delivery



## **Risk of Stroke Based on Time After Delivery**

| Time after<br>Delivery | •   | Crossover Period r 100,000 veries | Absolute Risk<br>Difference | Odds Ratio<br>(95% CI) |
|------------------------|-----|-----------------------------------|-----------------------------|------------------------|
| Weeks 0-6              | 7.1 | 0.8                               | 6.2                         | 8.5 (4.9 - 14.8)       |
| Weeks 7-12             | 0.9 | 0.5                               | 0.4                         | 1.7 (0.7 - 3.8)        |
| Weeks 13-18            | 0.5 | 0.5                               | 0                           | 1.0 (0.4 – 2.5)        |
| Weeks 19-24            | 0.9 | 0.9                               | 0.1                         | 1.1 (0.5- 2.2)         |

## Risk of Thrombotic Event Based on Time After Delivery *Odds Ratios*

| Time after<br>Delivery | Stroke | <b>MI</b><br>Odds | <b>VTE</b><br>Ratio | Composite | All  |
|------------------------|--------|-------------------|---------------------|-----------|------|
| Weeks 0-6              | 8.5    | 13.0              | 12.1                | 10.8      | 22.8 |
| Weeks 7-12             | 1.7    | 4.0               | 2.2                 | 2.2       | 2.1  |
| Weeks 13-18            | 1.0    | 1.0               | 1.6                 | 1.4       | 1.0  |
| Weeks 19-24            | 1.1    | 2.5               | 0.9                 | 1.0       | 0.9  |

## Clear increase in risk for at least 12 weeks; though small after 6 weeks...

Kamel NEJM 2014;370:1307

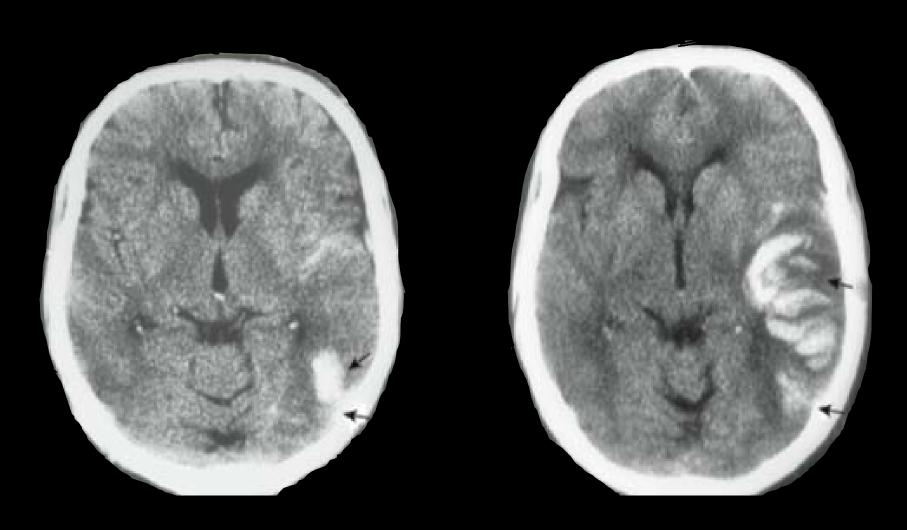
## Risk of Thrombotic Event Based on Time After Delivery \*\*Absolute Risk\*\*

| Time after<br>Delivery | <b>Stroke</b><br>Abs | <b>MI</b><br>solute Risk D | <b>VTE</b><br>Difference | Composite | All   |
|------------------------|----------------------|----------------------------|--------------------------|-----------|-------|
| Weeks 0-6              | 6.2                  | 0.7                        | 15.2                     | 22.1      | 127.6 |
| Weeks 7-12             | 0.4                  | 0.4                        | 2.3                      | 3.0       | 6.1   |
| Weeks 13-18            | 0                    | 0                          | 0.9                      | 0.9       | 0.2   |
| Weeks 19-24            | 0.1                  | 0.2                        | -0.3                     | -0.1      | -0.9  |

Clinical importance: VTE > Stroke > MI

#### Case

A 34-year-old woman began having headaches several days after delivery of her first child. The pregnancy and delivery had been normal, and the baby was healthy. Her headaches were diffuse, worse at night. Four days after onset, her husband witnessed a grand mal seizure. She had no history of prior seizures. On initial examination her pulse was 80 and regular, BP 115/70; she was aphasic and had mild right hemiparesis.



Head CT without contrast

# Causes of Hemorrhagic Stroke in Pregnancy Percent of All Hemorrhages

|               | Preeclampsia<br>Eclampsia | Unknown | AVM | Aneurysm | Other | Cavernous<br>Malformation |
|---------------|---------------------------|---------|-----|----------|-------|---------------------------|
| Feske 2009    | 42                        | 11      | 14  | 14       | 17    | 3                         |
| Liang 2006    | 24                        | 24      | 19  | 10       | 24    |                           |
| Jeng 2004     | 32                        |         | 23  | 14       |       |                           |
| Jaigobin 2000 |                           | 23      | 38  | 23       | 15    |                           |
| Kittner 1996  | 15                        | 31      | 23  |          | 31    |                           |
| Sharshar 1995 | s 44                      | 19      | 13  | 13       |       | 13                        |

# Causes of Hemorrhagic Stroke in Pregnancy Percent of All Hemorrhages

|                   | Preeclampsia<br>Eclampsia | Unknown | AVM | Aneurysm | CVT | Cavernous<br>Malformation |
|-------------------|---------------------------|---------|-----|----------|-----|---------------------------|
| Feske 2009        | 42                        | 11      | 14  | 14       | 17  | 3                         |
| <b>Liang</b> 2006 | 24                        | 24      | 19  | 10       | 24  |                           |
| Jeng 2004         | 32                        |         | 23  | 14       |     |                           |
| Jaigobin 2000     |                           | 23      | 38  | 23       | 15  |                           |
| Kittner 1996      | 15                        | 31      | 23  |          | 31  |                           |
| Sharshar 1999     | 5 44                      | 19      | 13  | 13       |     | 13                        |

### **Goals of Neuroimaging in Pregnancy**

- 1. Provide standard of care imaging able to answer important diagnostic questions.
- 2. Minimize risks to the fetus.
- 3. Use radiation doses as low as reasonably achievable for potential stochastic effects.
- 4. Use doses below exposure thresholds for deterministic effects.

- Stochastic effects May occur after any dose of radiation; higher doses increase risk.
  - Mutagenesis
  - Childhood malignancy
- Deterministic effects Predictably occur above specific exposure thresholds.
  - Cataract formation
  - Infertility

### **Considerations in Neuroimaging in Pregnancy**

- 1. Radiation dose and rate absorbed
  - 1. E.g. Estimate 6% increase in risk of childhood cancer per 100 rad.
  - 2. Fetal exposure to indirect radiation from CT is to be < 0.01 rad.
  - 3. Fetal exposure to direct radiation from pelvic CT may reach 3 rad.
- 2. Fetal gestational age
  - 1. 0-4 weeks Increase risk of miscarriage with doses > 10 rad.
  - 2. 5-10 weeks Fetal malformation, growth retardation, and death possible with doses > 5 -10 rad.
  - 3. 6 weeks birth Mental retardation with doses > 5-10 rad.
    - 1. Very low risk after 15 weeks
- 3. Urgency of diagnostic need

## For potential stroke in pregnancy:

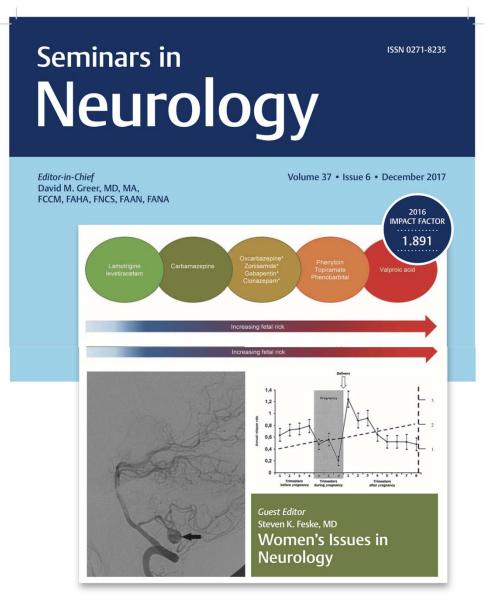
- Degree of urgency is high.
- Exposure is indirect and doses are low.
- Events occur late in pregnancy when fetal risks are minimal.

## **Further Considerations in Neuroimaging in Pregnancy**

- 1. MRI
- 2. Iodinated Contrast Agents
- 3. Gadolinium

### **Further Considerations in Neuroimaging in Pregnancy**

- 1. MRI is felt to be safe.
  - 1. No conclusive evidence of fetal harm from exposure up to 3 T.
  - 2. Theoretical concerns
    - 1. Noise exposure
    - 2. Strong magnetic fields
    - 3. Increase in body temperature
- 2. Iodinated contrast agents should be avoided, except when no alternative.
  - 1. Theoretical concerns
    - 1. Neonatal hypothyroidism
    - 2. Renal injury
- 3. Gadolinium should be avoided.
  - 1. Theoretical concerns
    - 1. Miscarriage
    - 2. Developmental abnormalities





Chansakul T, Young GS.
Neuroimaging in pregnant women.
Semin Neurol 2017;37(6):712-23.

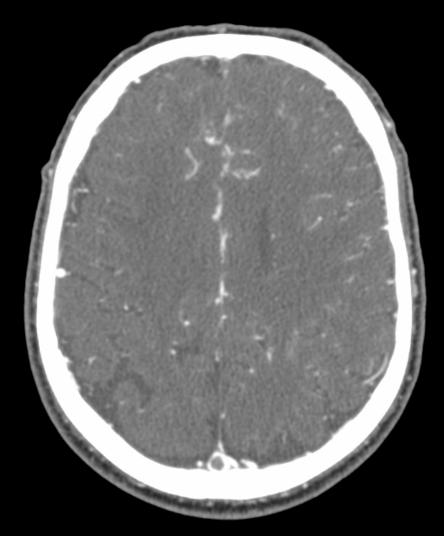
Sells CM, Feske SK. Stroke in pregnancy. Semin Neurol 2017;37(6):669-78.

Can A, Du R. Neurosurgical issues in pregnancy. Semin Neurol 2017;37(6):689-93.



CT without contrast

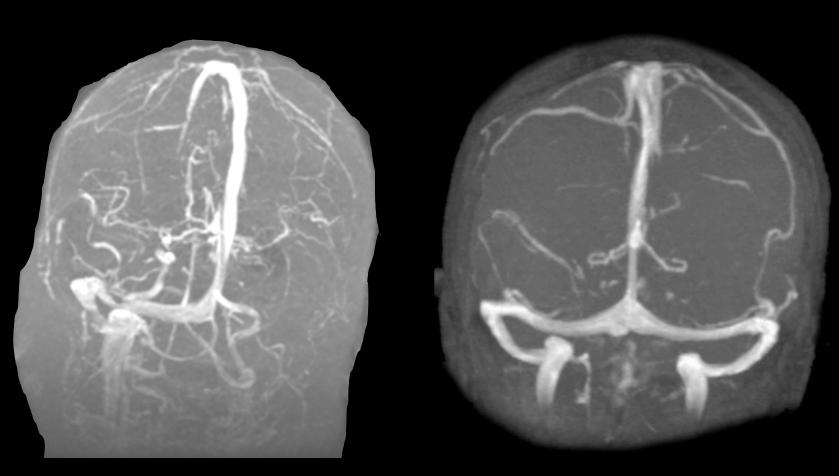
## Empty Delta Sign



CT with contrast



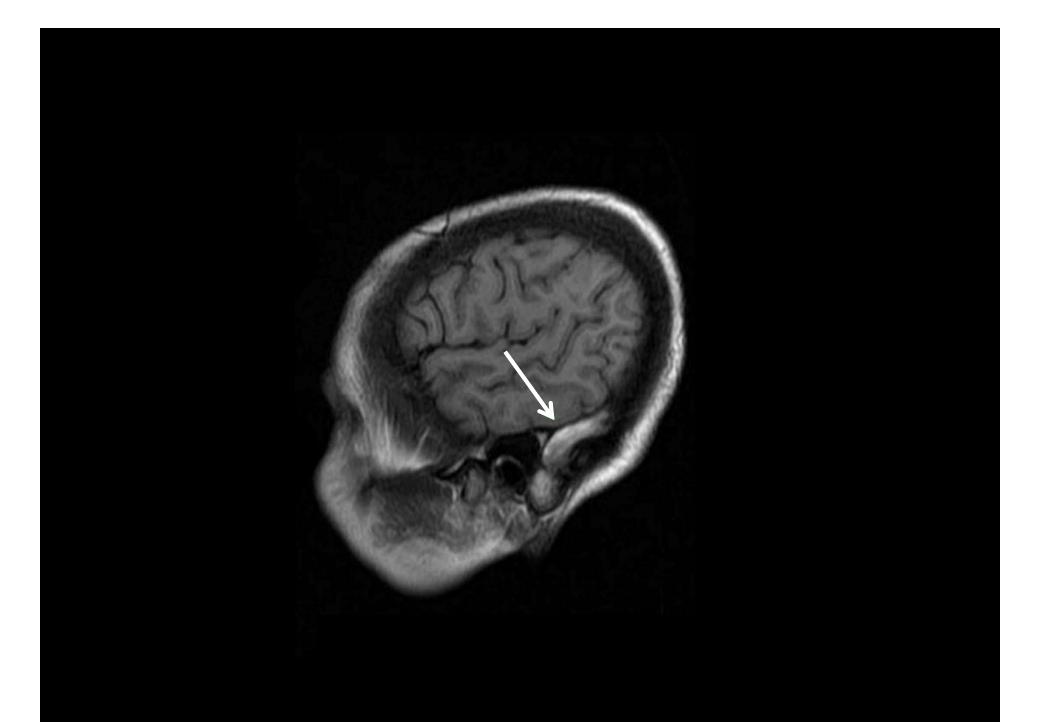
MR Venogram



Left Transverse Sinus Thrombosis

Normal MR Venogram



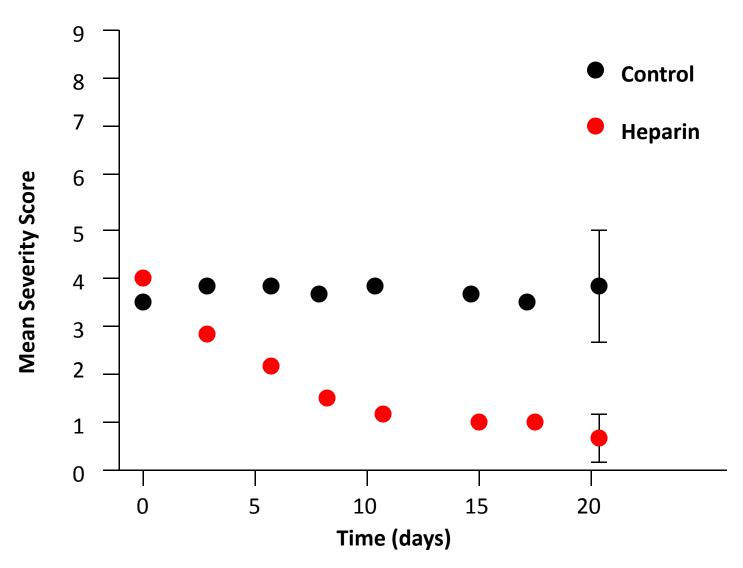




#### **Treatment of CVT**

- 1. Einhäupl KM et al. Heparin treatment in sinus venous thrombosis. Lancet 1991;338:597-600.
- 2. de Bruijn SF et al. Randomized, placebo-controlled trial of anticoagulation treatment with low-molecular-weight heparin for cerebral sinus thrombosis. Stroke 1999;30:484-8.
- 3. Stam J, et al. Anticoagulation for cerebral sinus thrombosis. Cochrane Database Syst Rev 2002;(4):CD002005.
- 4. Ferro JM, et al for the ISCVT Investigators. Prognosis of cerebral vein and dural sinus thrombosis: Results of the International Study on Cerebral Vein and Dural Sinus Thrombosis (ISCVT) Stroke 2004;35:664-70.

#### Heparin for Venous Sinus Thrombosis

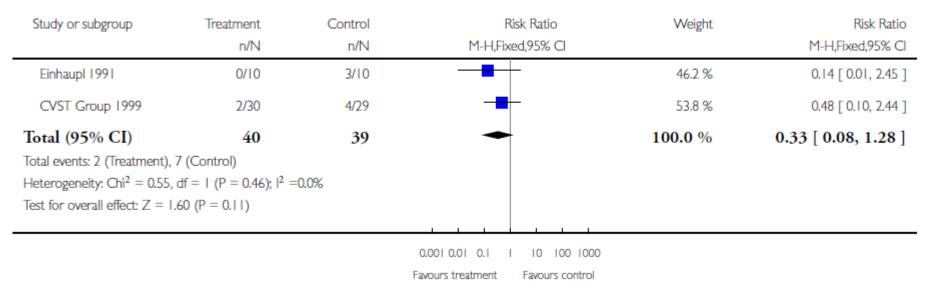


Einhäupl Lancet 1991;338:597

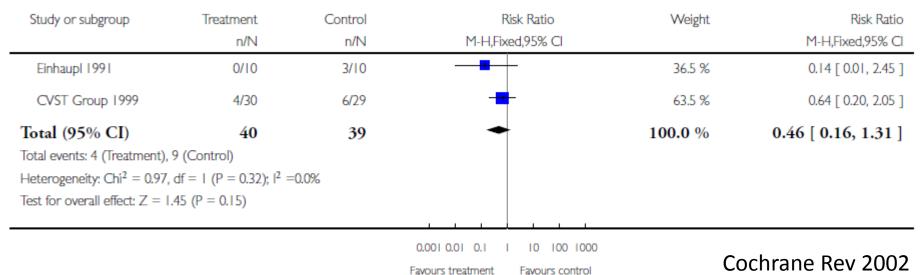
# Heparin for Venous Sinus Thrombosis 3-month Outcomes

| Outcome                   | Control (N=10) | Heparin (N=10) |  |
|---------------------------|----------------|----------------|--|
| Complete recovery         | 1              | 8              |  |
| Slight neurologic deficit | 6              | 2              |  |
| Death                     | 3              | 0              |  |

#### Overall Benefit or Harm of Heparin, Outcome: Death



#### Overall Benefit or Harm of Heparin, Outcome: Death and Dependency



## **AHA/ASA Guidelines 2011**

- 6. For patients with CVT, initial anticoagulation with UFH or LMWH in full anticoagulant doses is reasonable, followed by warfarin, regardless of the presence of ICH (Class IIa; Level B)
- 8. In patients with CVT and increased ICP it is reasonable to initiate treatment with acetazolamide (Class IIa; Level C)
- 9. Endovascular intervention may be considered if deterioration occurs despite intensive anticoagulation treatment (Class IIb; Level C)
- 10. In patients with neurological deterioration due to severe mass effect or ICH causing intractable intracranial hypertension, decompressive hemicraniectomy may be considered (Class IIb; Level C)

#### Case

#### 28-year-old RH woman 30-weeks pregnant without prior complications

- 10:00 AM last seen well
- 10:50 AM found on ground by her husband, eyes open, mute, weak on R
- Brought to a local hospital
  - Alert without gaze deviation
  - Dense motor aphasia, mute
  - Dense right hemiplegia
  - No signs of trauma
  - Normal CBC, platelets, INR, PTT
  - Head CT normal (or subtle change of acute stroke; no hemorrhage)
  - MRI early acute stroke left basal ganglia

#### Case

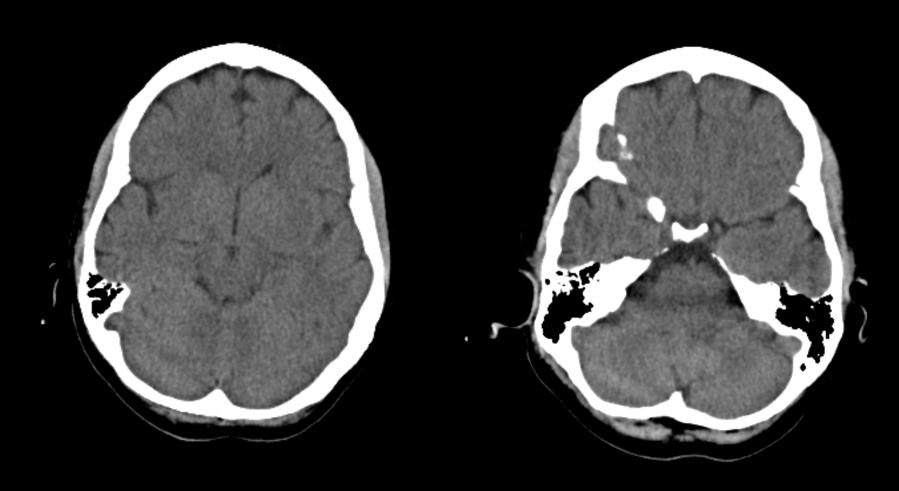
#### **PMH**

- G2 P1
- G1 2009; stat C-section at term for concerning fetal heart tracing
- G2 current @ 30 weeks;
  - Rh<sup>-</sup> received Rhogam at 28 weeks
  - Observed briefly for preterm contractions at 28 weeks
- No miscarriages
- No pre-eclampsia-eclampsia
- No prior abnormal thrombosis
- No trauma
- Nonsmoker; no alcohol or drug abuse

#### FH

- 2 maternal uncles and one aunt with DVT/PE; ?FVIII excess
- No family history of arterial dissection, aneurysm, or AVM
- Father estranged and history unknown

## On Arrival at Local Hospital



CT without contrast

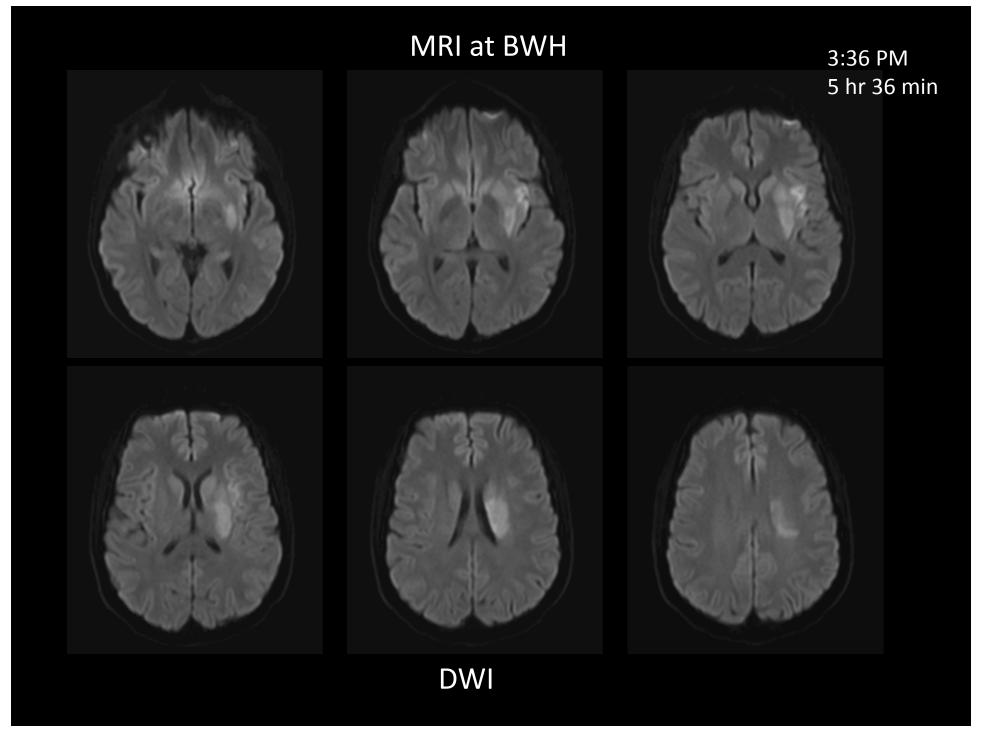
## On Arrival at Local Hospital



CT without contrast

What would you do now?

# What additional imaging studies would you get?



#### MRA at BWH

3:36 PM 5 hr 36 min





Left M1 occlusion





3:36 PM 5 hr 36 min

Open left ICA without evidence of dissection

#### **Mechanisms of Ischemic Stroke in Pregnancy**

| Author/Year                    | % Cardio-<br>embolism | %<br>PEE | % Peripartum<br>Angiopathy | % %<br>CVT | Unknown | %<br>Other |
|--------------------------------|-----------------------|----------|----------------------------|------------|---------|------------|
| Awada 1995<br>Saudi Arabia     | 33                    | 11       |                            |            | 44      | 11         |
| Sharshar 1995<br>Ile de France |                       | 47       | 7                          |            | 27      | 20         |
| Kittner 1996<br>Md/Wash DC     | (                     | 25       | 13                         | 6          | 38      | 19         |
| Witlin 1997<br>Memphis         |                       |          |                            | 64         |         |            |
| Jiagobin 2000<br>Toronto       | 20                    | 20       |                            | 40         | 20      | 15         |
| Jeng 2004<br>Taiwan            | 44                    |          |                            | 22         | 22      |            |
| Liang 2006<br>Taiwan           | 36                    | 18       |                            | 27         |         |            |
| Feske 2009<br>Boston           | 35                    | 26       |                            | 39         |         | 22         |

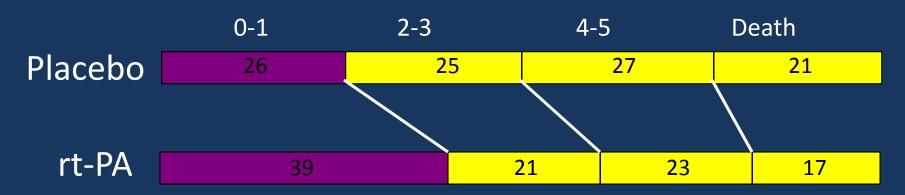


# Treatment of Acute Ischemic Stroke



# NINDS Study of IV rt-PA for Acute Ischemic Stroke Outcome

#### **Modified Rankin Scale**



ARR = 13 %

NNT = 8

NNH = 20 (sICH 6 v 0.6 %)

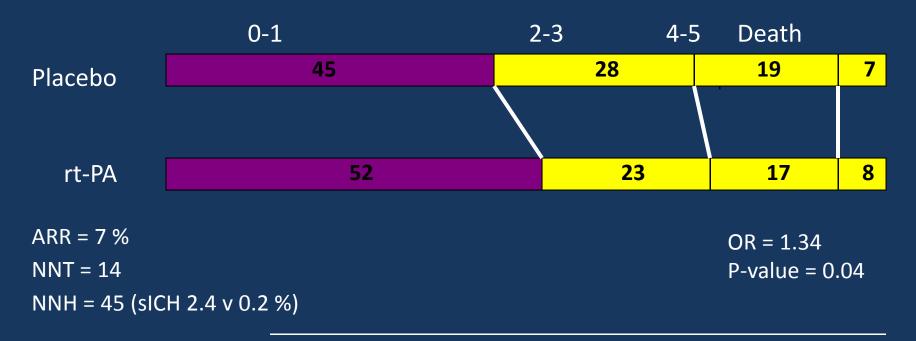
(Harm as sICH)

The NINDS Stroke Study Group NEJM 1995;333:1581

#### IV tPA in 3-4.5 hr Window

#### **ECASS III Results: Primary Endpoint**

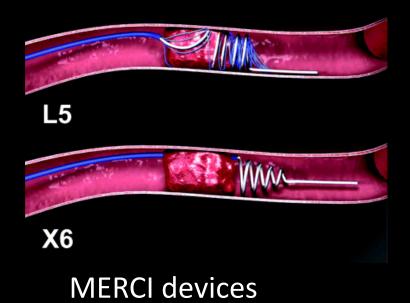
#### Modified Rankin Scale at 90 days



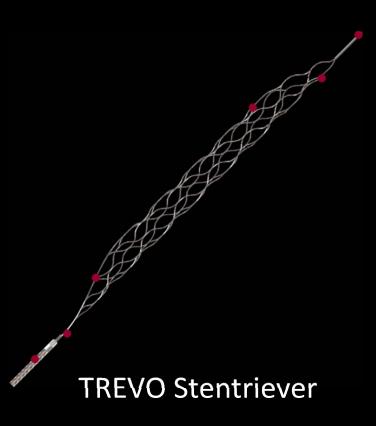
ECASS III NEJM 2008;359:1317

# Usable length Usable length Push wire Proximal marker

#### **SOLITAIRE Flow Restoration**



### Mechanical Clot Retrieval Devices



# Dense Left MCA Sign

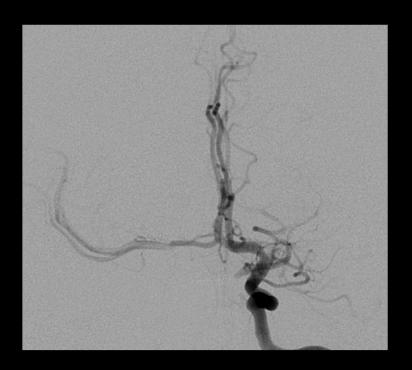


CT without contrast

# Occlusion of the Terminal LICA



CTA



Before clot extraction



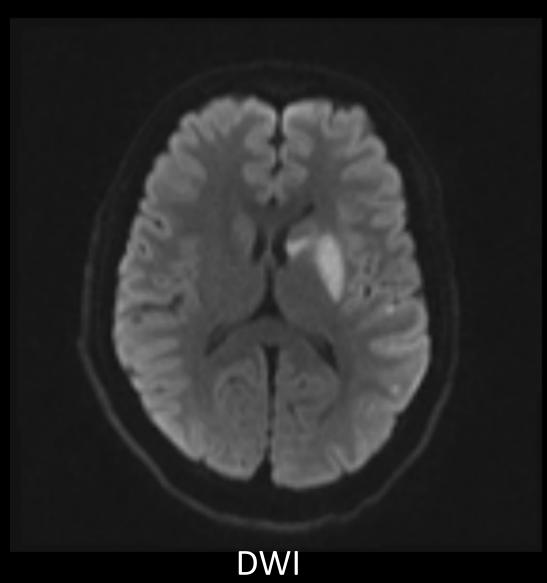
After clot extraction

Angiogram

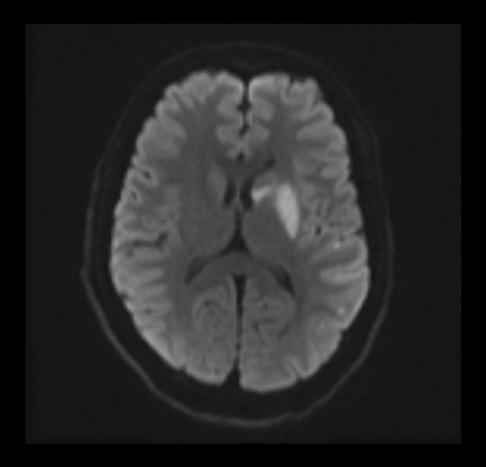


Stent Retriever with Extracted Clot

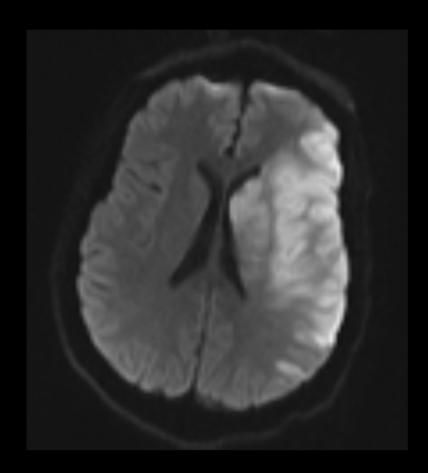
# Final Stroke



## Final Stroke

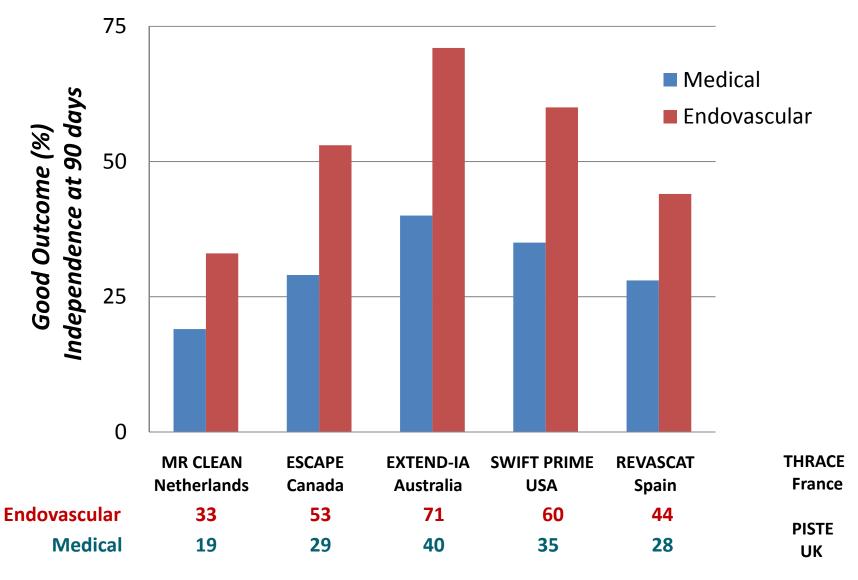


This...

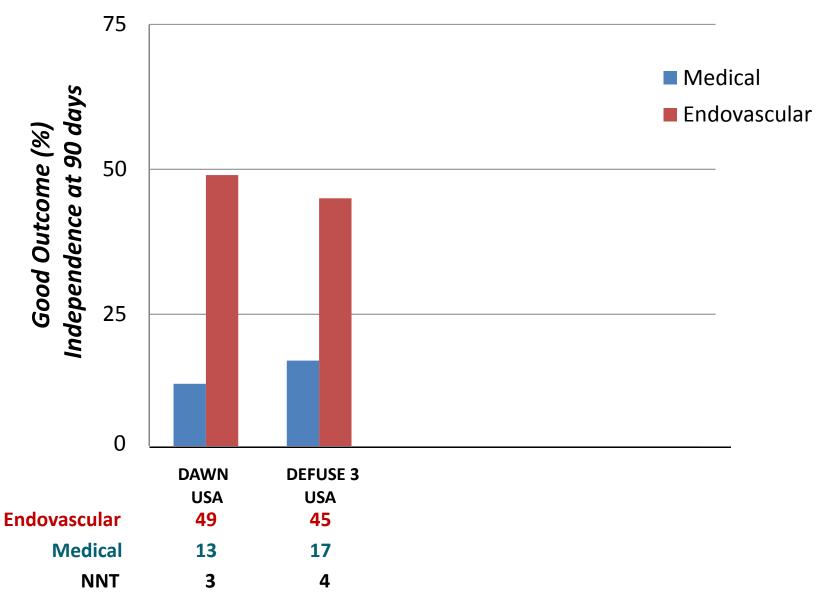


Not This!

# Randomized Clinical Trials of Endovascular Therapy for Acute Ischemic Stroke 2015



# Randomized Clinical Trials of Endovascular Therapy for Acute Ischemic Stroke in Late Window 2018



#### tPA use in Pregnancy

| Study/Year Wk   | Gestation | Indication<br>No.treated | Outcome<br>Mother                  | Outcome<br>Fetus     |
|-----------------|-----------|--------------------------|------------------------------------|----------------------|
| Baudo 1990      | 35        | PE 1                     | No cx                              | No cx                |
| Flossdorf 1990  | 31        | PE 1                     | No cx                              | No cx                |
| Azzano 1995     | 16        | Valve 1                  | Severe<br>bleeding<br>rethrombosis | Fetal death<br>after |
| Schumacher 1996 | 21        | MI 1                     | No cx                              | No cx                |
| Fleyfe 1997     | 28        | Valve 1                  | No cx                              | No cx                |
| Total           |           | 5                        | 1 bleeding                         | 1 death              |

From Ahearn Arch Intern Med 2002;162:1221

#### **IV tPA Use for Stroke in Pregnancy**

|                 | Maternal/Gestational | Outcome                |       |
|-----------------|----------------------|------------------------|-------|
| Study/Year      | Age                  | Maternal               | Fetal |
| Dapprich 2002   | / 12 wk              | minor ICH              | No Cx |
| Weise 2006      | 33 yr / 13 wk        | No Cx                  | No Cx |
| Leonhardt 2006  | 26 yr / 23 wk        | No Cx                  | No Cx |
| Murugappan 2006 | 37 yr / 12 wk        | minor uterine hematoma | MTP*  |
| Murugappan 2006 | 31 yr / 4 wk         | No Cx                  | MTP*  |
| Murugappan 2006 | 29 yr / 6 wk         | died**                 | died  |
| Yamaguchi 2010  | 36 yr / 18 wk        | No Cx                  | No Cx |
| Hori 2013       | 35 yr / 4 mos        | no Cx                  | No Cx |
| Tassi 2013      | 28 yr / 16 wk        | no Cx                  | No Cx |
| Ritter 2014     | 32 yr / 36 wk        | no Cx                  | No Cx |

<sup>\*</sup>MTP = medical termination of pregnancy

Dapprich Cerebrovasc Dis 2002;13:290
Wiese Stroke 2006;37:2168
Leonhardt J Throm Thrombolys 2006;21:271
Murugappan Neurology 2006;66:768
Yamaguchi Rinsho Shinkeigaku 2010;50:315
Hori Rinsho Shinkeigaku 2013;53:212
Tassi Am J Emerg Med 2013;31:448
Ritter J Neurol 2014;261:632

<sup>\*\*</sup> died from arterial dissection complicating angioplasty

#### **IA tPA Use for Stroke in Pregnancy**

|                 | Maternal/Gestational              | Outcome          |              |  |
|-----------------|-----------------------------------|------------------|--------------|--|
| Study/Year      | Age                               | Maternal         | Fetal        |  |
| Elfort 2002     | 28 / 1wk (after IVF)              | minor ICH        | No Cx        |  |
| Johnson 2005    | 39 yr / 37 wk                     | No Cx            | No Cx        |  |
| Murugappan 2006 | 43 yr / 37 wk                     | No Cx            | No Cx        |  |
| Murugappan 2006 | 28 yr / 6 wk                      | buttock hematoma | No Cx        |  |
| Murugappan 2006 | 25 yr / 1 <sup>st</sup> trimester | minor ICH        | Miscarriage* |  |
| Li 2012         | 24 yr / 11 wk                     | No Cx            | No Cx        |  |

Elfort Neurology 2002;59:1270 Johnson Stroke 2005;36:e53

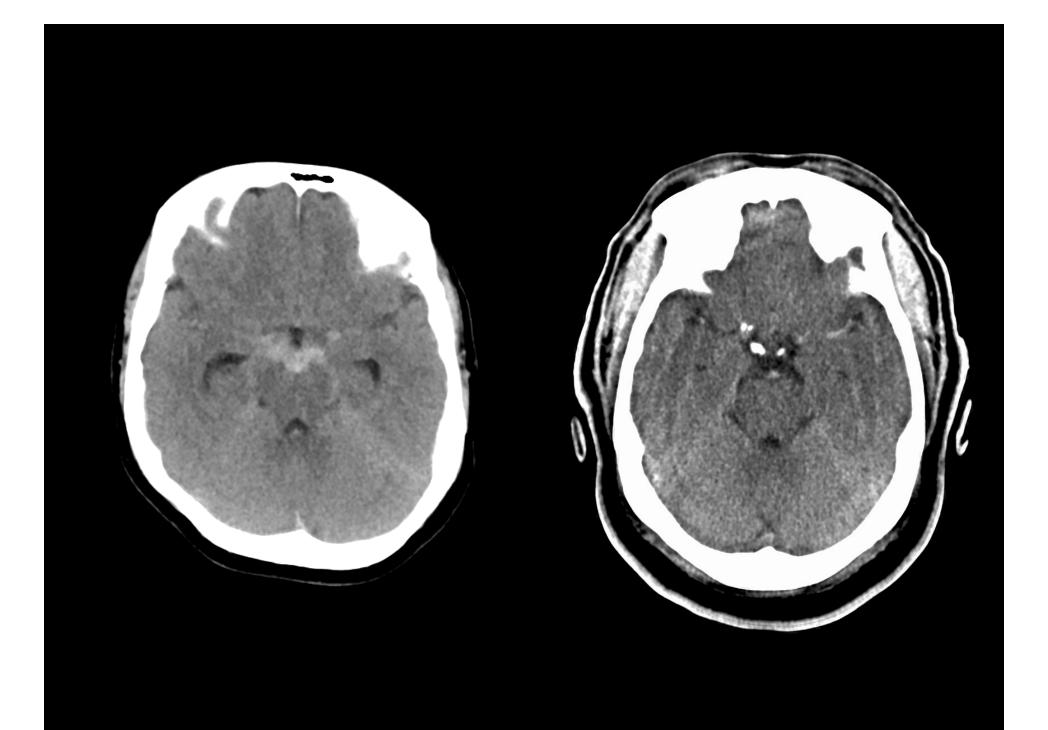
Murugappan Neurology 2006;66:768

Li Neurologist 2012;18:44

<sup>\*</sup> Mother had bacterial endocarditis

#### Case

A 37-year-old woman 27 weeks pregnant developed a **sudden**, **severe headache** and nausea and vomiting and **neck stiffness**. On initial examination her pulse was 100 and regular, BP 145/70; she was initially alert and then slightly drowsy. Otherwise mental state and the rest of the neurologic examination were normal.



# Conventional Angiogram



L Vertebral Injection AP

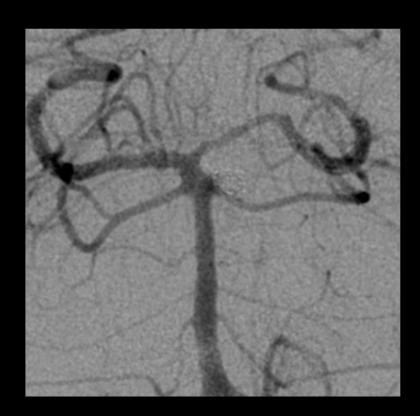
# Conventional Angiogram



3D Reconstruction

## Conventional Angiogram





Before After Coiling

L Vertebral Injection AP

## **Importance of Hemorrhagic Stroke in Pregnancy**

| <b>Absolute risk</b> | 0.006 |
|----------------------|-------|
| Relative risk        | 28.5  |
|                      |       |
| Mortality            | 5-12% |
| •                    |       |

## **Mechanisms of Hemorrhagic Stroke in Pregnancy**

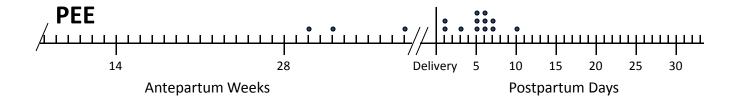
| Author/Year                           | %<br>AVM | %<br>Aneurysm | %<br>CM | %<br>PEE | %<br>Unknown | %<br>Other |
|---------------------------------------|----------|---------------|---------|----------|--------------|------------|
| Sharshar 1995<br>Ile de France N = 16 | 13       | 13            | 13      | 44       | 19           |            |
| Kittner 1996<br>Md/Wash DC N = 13     | 23       |               |         | 15       | 31           | 31         |
| Witlin 1997<br>Memphis N = 6          | 5        | 60            |         |          | 50           |            |
| Jiagobin 2000<br>Toronto N = 13       | 38       | 23            |         |          | 23           | 15         |
| Jeng 2004<br>Taiwan N = 22            | 23       | 14            |         | 32       |              |            |
| Liang 2006<br>Taiwan N = 21           | 19       | 10            |         | 24       | 24           | 24         |
| Feske 2009*<br>Boston N = 30          | 17       | 17            | 3       | 50       | 13           |            |

<sup>\* 6</sup> CVT not included

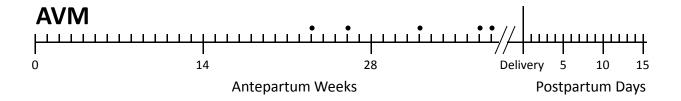
## **Mechanisms of Hemorrhagic Stroke in Pregnancy**

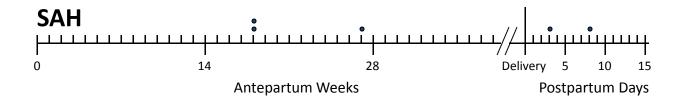
| Author/Year                           | %<br>AVM | %<br>Aneurysm | %<br>CM | %<br>PEE | %<br>Unknown | %<br>Other |
|---------------------------------------|----------|---------------|---------|----------|--------------|------------|
| Sharshar 1995<br>Ile de France N = 16 | 13       | 13            | 13      | 44       | 19           |            |
| Kittner 1996<br>Md/Wash DC N = 13     | 23       |               |         | 15       | 31           | 31         |
| Witlin 1997<br>Memphis N = 6          |          | 50            |         |          | 50           |            |
| Jiagobin 2000<br>Toronto N = 13       | 38       | 23            |         |          | 23           | 15         |
| Jeng 2004<br>Taiwan N = 22            | 23       | 14            |         | 32       |              |            |
| Liang 2006<br>Taiwan N = 21           | 19       | 10            |         | 24       | 24           | 24         |
| Feske 2009*<br>Boston N = 30          | 17       | 17            | 3       | 50       | 13           |            |
| 6 CVT not included                    |          |               |         |          |              |            |

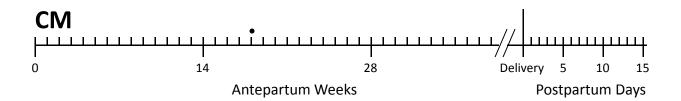
# Timing of cerebral hemorrhages associated with preeclampsia/eclampsia



# Timing of cerebral hemorrhages associated with vascular malformations







## **Treatment of aneurysms**

### Risks

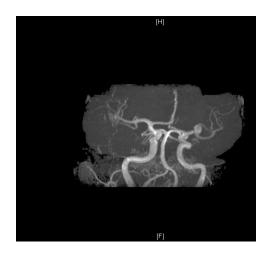
- Increasing risk of recurrent hemorrhage with progression of pregnancy; peaks at 30-34 weeks
- High risk of recurrent hemorrhage if an initial bleeding aneurysm goes unsecured: 33-50%



Overall: mother 35%; fetus 17% With no surgery: mother 63%; fetus 27% With surgery: mother 11%; fetus 5%



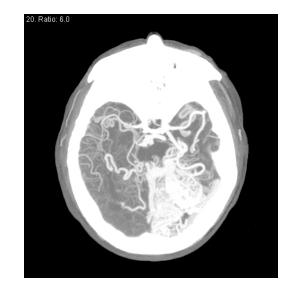
- Secure aneurysm as soon as possible after rupture by open or endovascular surgery.
- If cannot, because urgent obstetrical issues prevent it, then proceed to C-section and then secure aneurysm.



## **Treatment of AVMs**

### Risks

- Some authors have found increased risk of AVM hemorrhage during pregnancy, others have not.
- Analysis of risk of rupture per day shows many-fold increase of risk on day of delivery.



• Risk is greatly increased after hemorrhage during pregnancy; to about 26% (vs 6% risk if hemorrhage before pregnancy).

### **Recommendations**

- If known AVM, address before pregnancy.
- If AVM found during pregnancy without hemorrhage, "controlled delivery" with plan to treat AVM after delivery.
- If AVM bleeds during pregnancy, treat definitively based on neurosurgical principles (based on grading of AVM).

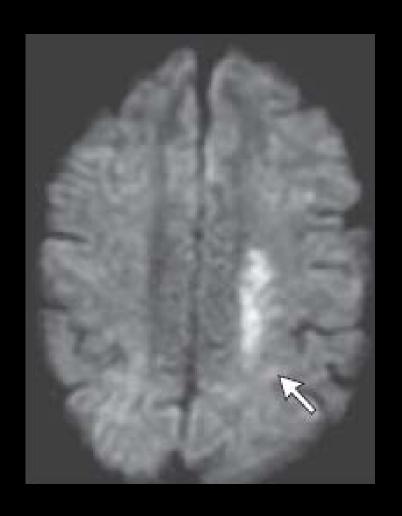
## Case

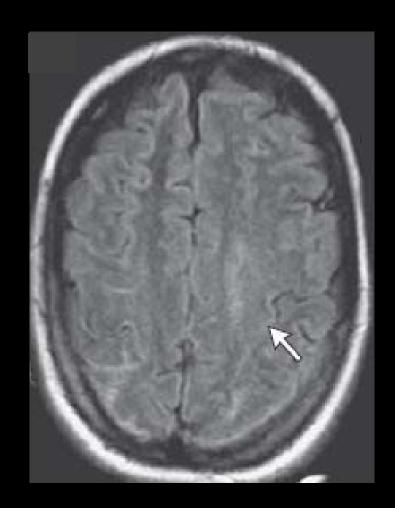
A 36-year-old woman complained of headaches and was found to have new HTN 10 days after delivery of twins by C-section. Initial head CT and MRI were normal. Headaches persisted, and she had a grand mal seizure and developed aphasia and right hemiparesis.





Singhal AB NEJM 2009;360:1126





Singhal AB NEJM 2009;360:1126

What is the diagnosis?



# MgSO4 versus Phenytoin for Eclampsia

