

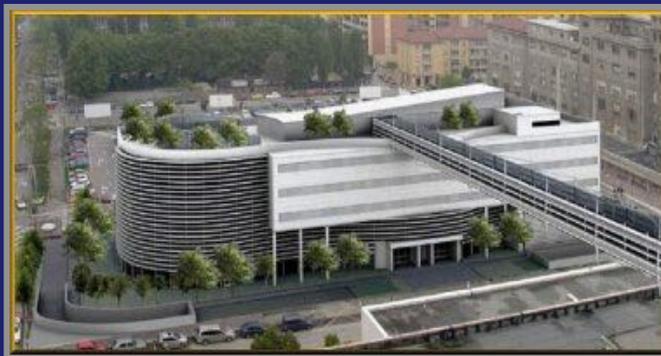


# Orthopaedic – Trauma Hospital and Sport Medicine Institute

## Department of Radiology and Diagnostic Imaging

Director: Dott. Carlo Faletti

### TORINO



ORTHOPAEDIC HOSPITAL  
and TRAUMA CENTER  
TURIN - ITALY



**The world  
of  
soft-tissue masses**

## WHO classification of soft tissue tumours

### ADIPOCYTIC TUMOURS

#### Benign

Lipoma	8850/0*
Lipomatosis	8850/0
Lipomatosis of nerve	8850/0
Lipoblastoma / Lipoblastomatosis	8881/0
Angiolipoma	8861/0
Myolipoma	8890/0
Chondroid lipoma	8862/0
Extrarenal angiomyolipoma	8860/0
Extra-adrenal myelolipoma	8870/0
Spindle cell/ Pleomorphic lipoma	8857/0 8854/0
Hibernoma	8880/0

#### Intermediate (locally aggressive)

Atypical lipomatous tumour/ Well differentiated liposarcoma	8851/3
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#### Malignant

Dedifferentiated liposarcoma	8858/3
Myxoid liposarcoma	8852/3
Round cell liposarcoma	8853/3
Pleomorphic liposarcoma	8854/3
Mixed-type liposarcoma	8855/3
Liposarcoma, not otherwise specified	8850/3

### FIBROBLASTIC / MYOFIBROBLASTIC TUMOURS

#### Benign

Nodular fasciitis	
Proliferative fasciitis	
Proliferative myositis	
Myositis ossificans fibro-osseous pseudotumour of digits	
Ischaemic fasciitis	
Elastofibroma	8820/0
Fibrous hamartoma of infancy	
Myofibroma / Myofibromatosis	8824/0
Fibromatosis colli	
Juvenile hyaline fibromatosis	
Inclusion body fibromatosis	
Fibroma of tendon sheath	8810/0
Desmoplastic fibroblastoma	8810/0
Mammary-type myofibroblastoma	8825/0

\* Morphology code of the International Classification of Diseases for Oncology (ICD-O) (726) and the Systematized Nomenclature of Medicine (<http://snomed.org>).

Calcifying aponeurotic fibroma	8810/0
Angiomyofibroblastoma	8826/0
Cellular angiofibroma	9160/0
Nuchal-type fibroma	8810/0
Gardner fibroma	8810/0
Calcifying fibrous tumour	
Giant cell angiofibroma	9160/0

#### Intermediate (locally aggressive)

Superficial fibromatoses (palmar / plantar)	
Desmoid-type fibromatoses	8821/1
Lipofibromatosis	

#### Intermediate (rarely metastasizing)

Solitary fibrous tumour and haemangiopericytoma (incl. lipomatous haemangiopericytoma)	8815/1 9150/1
Inflammatory myofibroblastic tumour	8825/1
Low grade myofibroblastic sarcoma	8825/3
Myxoinflammatory fibroblastic sarcoma	8811/3
Infantile fibrosarcoma	8814/3

#### Malignant

Adult fibrosarcoma	8810/3
Myxofibrosarcoma	8811/3
Low grade fibromyxoid sarcoma hyalinizing spindle cell tumour	8811/3
Sclerosing epithelioid fibrosarcoma	8810/3

### SO-CALLED FIBROHISTIOCYTIC TUMOURS

#### Benign

Giant cell tumour of tendon sheath	9252/0
Diffuse-type giant cell tumour	9251/0
Deep benign fibrous histiocytoma	8830/0

#### Intermediate (rarely metastasizing)

Plexiform fibrohistiocytic tumour	8835/1
Giant cell tumour of soft tissues	9251/1

#### Malignant

Pleomorphic 'MFH' / Undifferentiated pleomorphic sarcoma	8830/3
Giant cell 'MFH' / Undifferentiated pleomorphic sarcoma with giant cells	8830/3
Inflammatory 'MFH' / Undifferentiated pleomorphic sarcoma with prominent inflammation	8830/3

### SMOOTH MUSCLE TUMOURS

Angioleiomyoma	8894/0
Deep leiomyoma	8890/0
Genital leiomyoma	8890/0
Leiomyosarcoma (excluding skin)	8890/3

### PERICYTIC (PERIVASCULAR) TUMOURS

Glomus tumour (and variants) malignant glomus tumour	8711/0 8711/3
Myopericytoma	8713/1

### SKELETAL MUSCLE TUMOURS

#### Benign

Rhabdomyoma adult type fetal type genital type	8900/0 8904/0 8903/0 8905/0
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#### Malignant

Embryonal rhabdomyosarcoma (incl. spindle cell, botryoid, anaplastic)	8910/3 8912/3 8910/3
Alveolar rhabdomyosarcoma (incl. solid, anaplastic)	8920/3
Pleomorphic rhabdomyosarcoma	8901/3

### VASCULAR TUMOURS

#### Benign

Haemangiomas of subcut/deep soft tissue: capillary cavernous arteriovenous venous intramuscular synovial	9120/0 9131/0 9121/0 9123/0 9122/0 9132/0 9120/0
Epithelioid haemangioma	9125/0
Angiomatosis	
Lymphangioma	9170/0

#### Intermediate (locally aggressive)

Kaposiform haemangiioendothelioma	9130/1
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#### Intermediate (rarely metastasizing)

Retiform haemangiioendothelioma	9135/1
Papillary intralymphatic angioendothelioma	9135/1

Composite haemangiioendothelioma	9130/1
Kaposi sarcoma	9140/3

#### Malignant

Epithelioid haemangiioendothelioma	9133/3
Angiosarcoma of soft tissue	9120/3

### CHONDRO-OSSEOUS TUMOURS

Soft tissue chondroma	9220/0
Mesenchymal chondrosarcoma	9240/3
Extraskeletal osteosarcoma	9180/3

### TUMOURS OF UNCERTAIN DIFFERENTIATION

#### Benign

Intramuscular myxoma (incl. cellular variant)	8840/0
Juxta-articular myxoma	8840/0
Deep ('aggressive') angiofibroma	8841/0
Pleomorphic hyalinizing angiectatic tumour	
Ectopic hamartomatous thymoma	8587/0

#### Intermediate (rarely metastasizing)

Angiomatoid fibrous histiocytoma	8836/1
Ossifying fibromyxoid tumour (incl. atypical / malignant)	8842/0
Mixed tumour/ Myoepithelioma/ Parachordoma	8940/1 8982/1 9373/1

#### Malignant

Synovial sarcoma	9040/3
Epithelioid sarcoma	8804/3
Alveolar soft part sarcoma	9581/3
Clear cell sarcoma of soft tissue	9044/3
Extraskeletal myxoid chondrosarcoma ("chordoid" type)	9231/3
PNET / Extraskeletal Ewing tumour pPNET extraskeletal Ewing tumour	9364/3 9260/3
Desmoplastic small round cell tumour	8806/3
Extra-renal rhabdoid tumour	8963/3
Malignant mesenchymoma	8990/3
Neoplasms with perivascular epithelioid cell differentiation (PEComa) clear cell myomelanocytic tumour	
Intimal sarcoma	8800/3



# FOCUS POINTS

1. Site of the lesion and anatomic compartment
2. Morphology
3. Relationship with the adjacent structures
4. Response of the lesion to treatment
5. Relapse

# Quali quesiti? Quali metodiche?

Diagnosi differenziale  
Stadiazione  
Ristadiazione (neoadiuvante)  
Follow up

## Imaging morfologico

US, TC, angio TC, RM,  
angio-RM , mdc blood-  
pool

## Imaging funzionale

Perfusione (CEUS, TC,  
RM dinamica),  
RM Diffusion  
spettroscopia MR

## Targeted Imaging

Imaging molecolare

# Imaging

1. Conventional Radiology
2. US
3. CT
4. MRI
5. Angiography
6. Nuclear Medicine

# Literature

Only few lesions have typical aspect

lipoma, sub-acute haematoma, neurinoma,  
villonodular pigmentous synovitis

*Sundaram et al: MR imaging of tumor and tumorlike lesions of bone and soft tissue. AJR 1990; 155: 817-824*

# Conventional Radiology

## A “Must” in the Study of Tumours

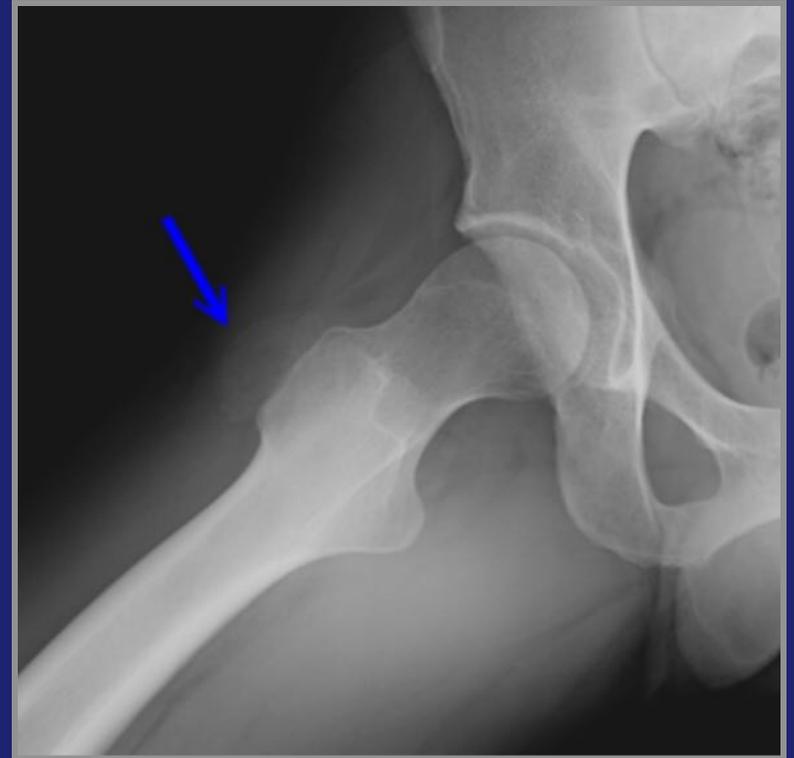
Soft  
tissue:

Differences in the density of soft-tissues  
Presence of calcifications and/or ossifications

Bone:

Cortical erosions  
Periosteal reactions  
Bone involvement

# Conventional Radiology



Different soft-tissues density

# Conventional Radiology



Presence of calcifications and/or ossifications

# Conventional Radiology



Cortical Erosion



Periosteal reaction



Bone alteration

First of all!

According to the Piedmont *Sarcoma Group Guide Lines*

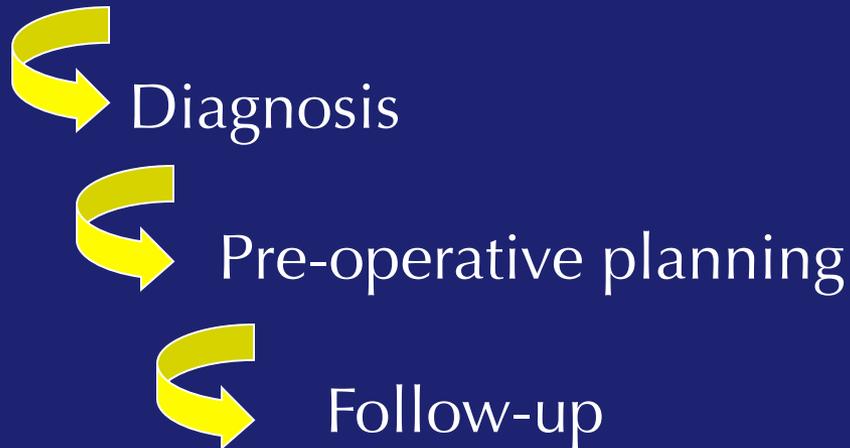


**A soft-tissue mass  
( less than 10 cm)  
is studied by ...**

**US**

# The aim

To establish a diagnostic US protocol  
for the study of soft-tissue masses



It is often difficult to identify sarcomatous lesions  
with quick and inexpensive techniques

cost-effective  
easy to perform  
well tolerated  
non invasive



immediate identification of aggressive lesions

swelling  
tenderness  
pain



# Method and materials

November 2002 - May 2008

The CTO/Maria Adelaide Hospital  
has investigated 1.164 subjects with soft-tissues  
masses.

Age

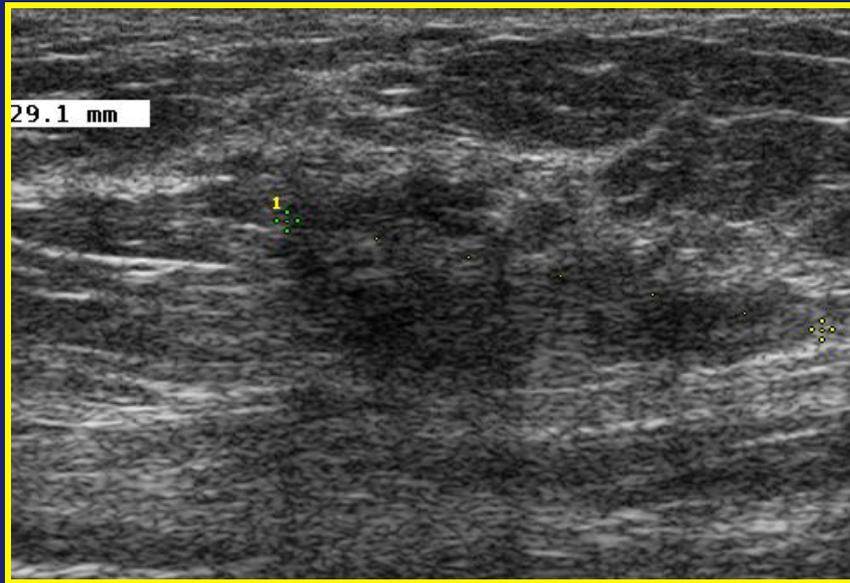
Gender

Site

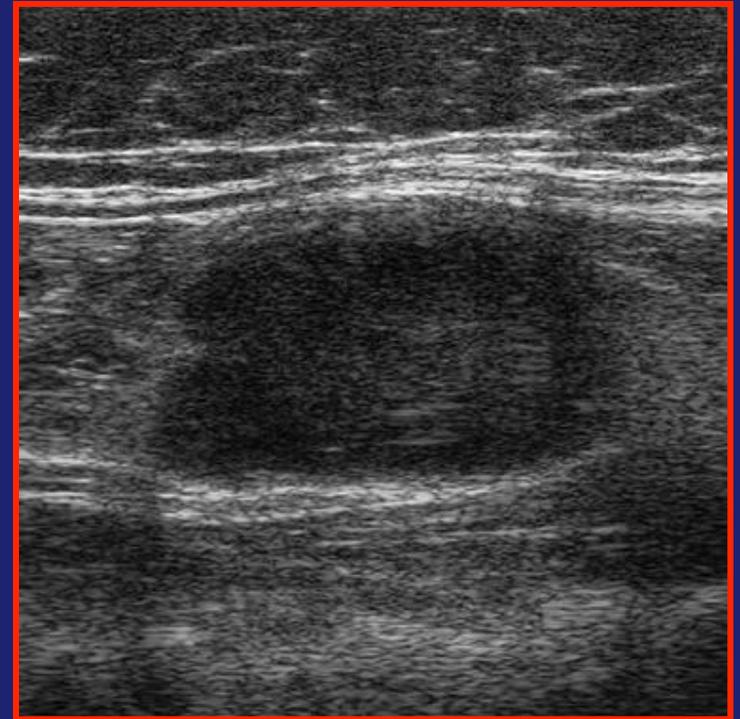
Morphology

Size

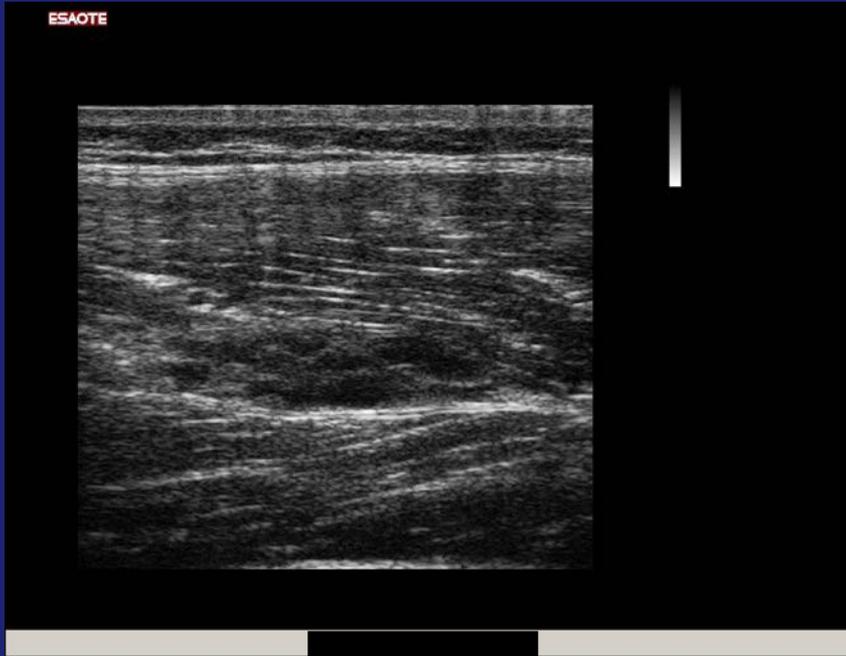
Vascularization



ENDOMETRIOSI



MIXOFIBROSARCOMA



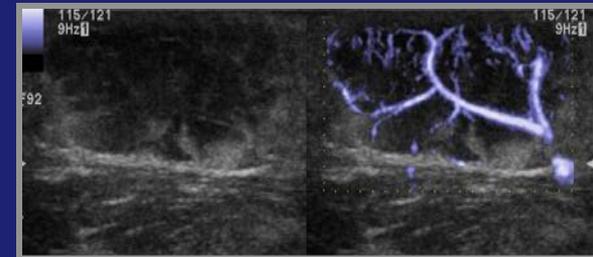
linfoangioma

# CD-US

Helpful in assessing the  
vascularity of soft tissue masses

Adler RS.1999  
Boddner G. 2002  
Griffith JF . 2004  
Hakan I 2006

A combination  
of vascular  
characteristics  
Specificity 93%



**PD**

sensibile ai flussi lenti rilevati  
solo nel macrocircolo  
rappresenta un passo avanti  
nella caratterizzazione

# CEUS

Microbolle gassose che rimangono confinate nei vasi, più piccole di un globulo rosso, atossiche, persistenti in circolo ed eliminate con il respiro

con macchine dedicate a basso indice meccanico (**MI**) le microbolle emettono buona risposta armonica senza andare incontro a rottura



durano per lungo tempo in circolo permettendo uno studio in real-time

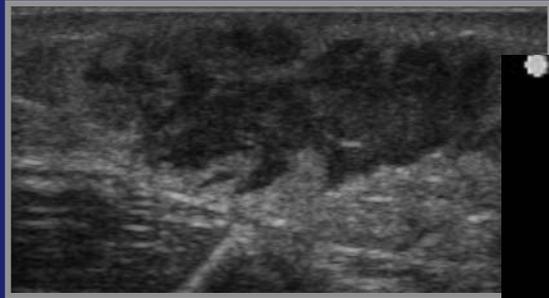
vera “**mappa vascolare**” del macro e microcircolo senza passare nello spazio interstiziale

**valore aggiunto**

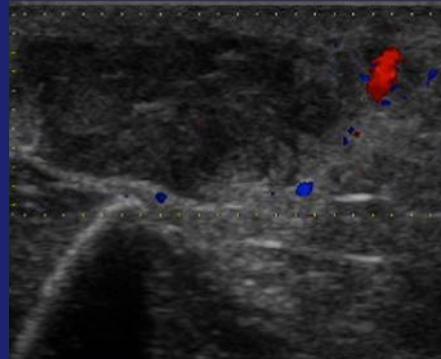
# Quando ?

- In tutte le lesioni  $>$  di cm5
- In tutte le lesioni cresciute rapidamente, anche  $<$  di cm5
- In tutte le lesioni dubbie per caratteristiche ecografiche e cliniche
- Controllo post-oper lesioni maligne per D.D fra cicatrice e recidiva
- Per la biopsia al fine di mirare la zona più significativa della lesione
  - Controllo lesioni benigne operate plurirecidivate
  - Controllo di lesioni benigne trattate con terapia conservativa
  - Quando il risultato di altre metodiche è in discordanza con la clinica

US



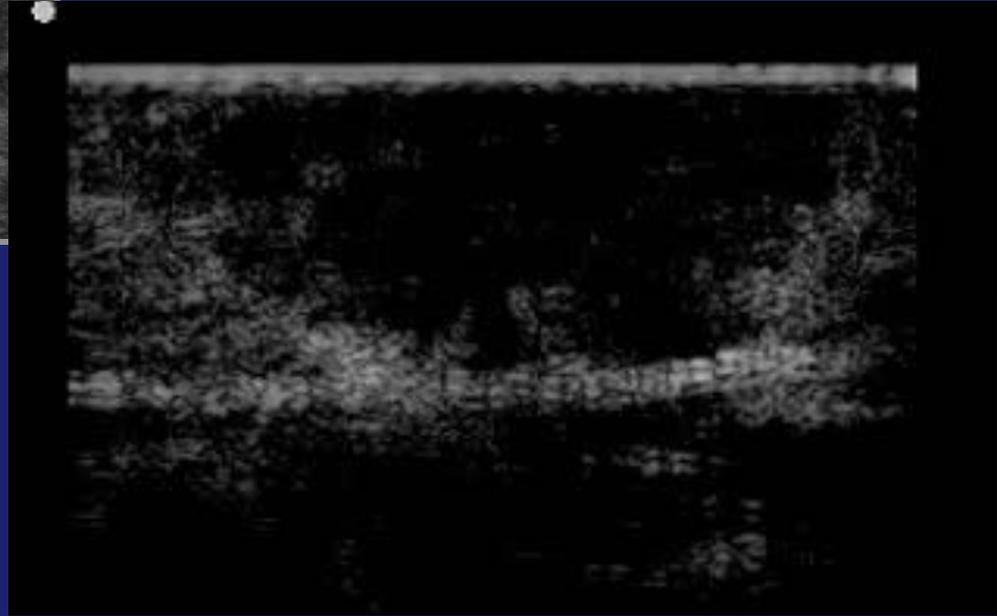
Longitudinale



Assiale

CD-US

CEUS



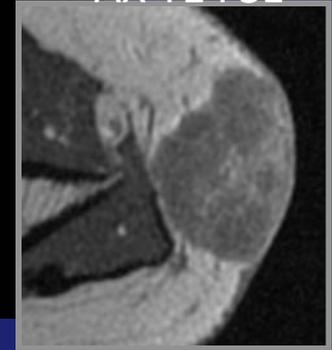
Vascular abnormalities may be missing or not detectable with current technique

Hwart L et al. J Radiol 2007  
Goh V et al. Abdom Imaging 2006

MRI



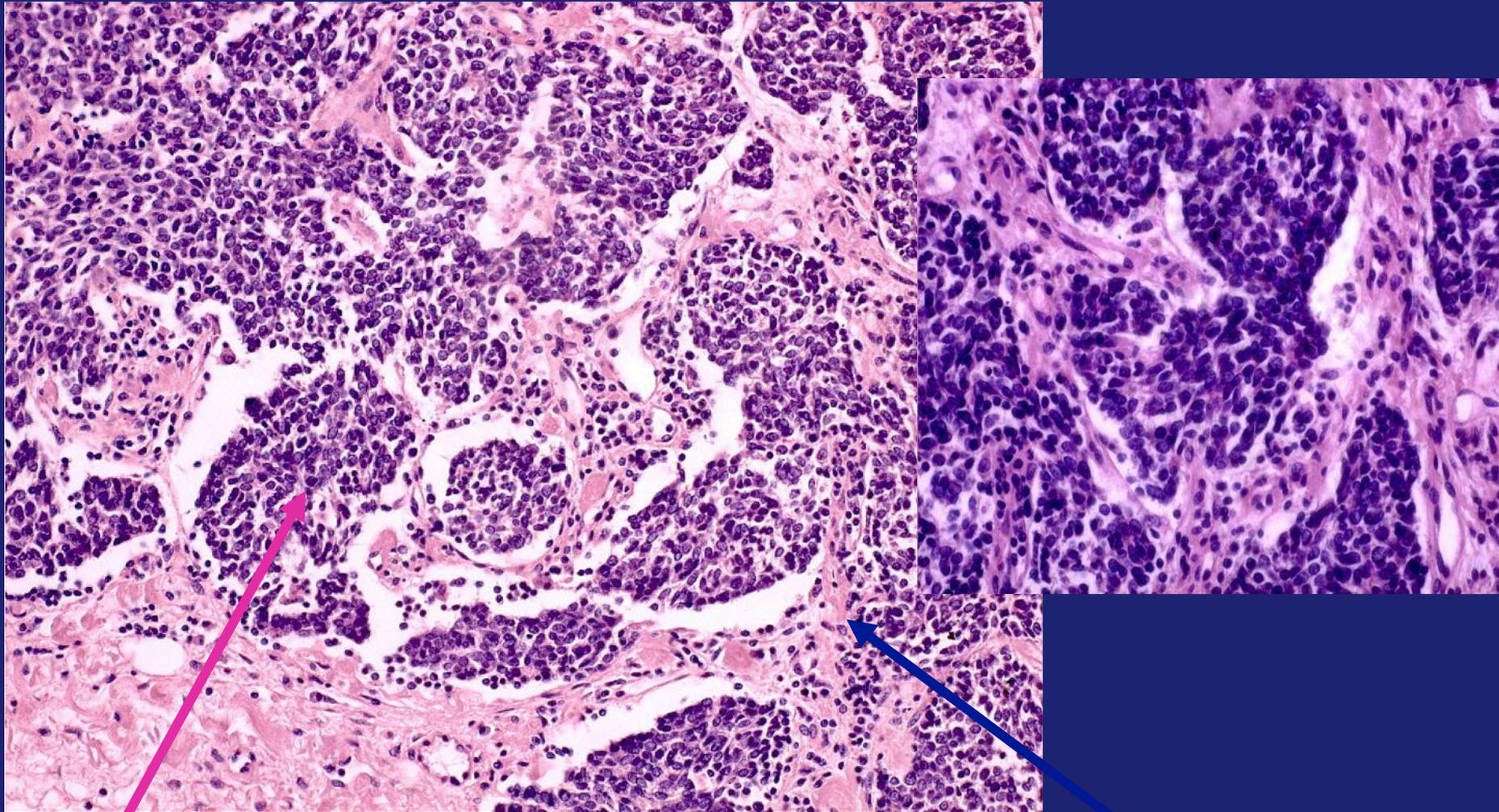
AX T1 FSE



AX T1 FSE MdC

CT and MR  
tumour enhancement  
is due to the presence  
of the contrast product  
in the interstitial  
space

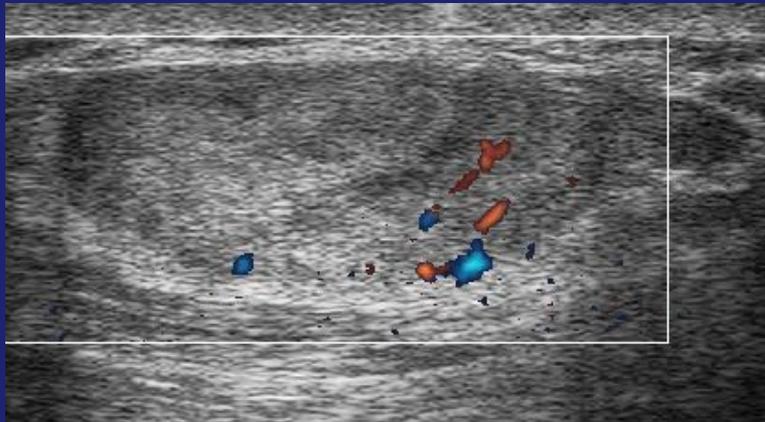
# Shwannoma maligno



Lacune vascolari malformate  $\rightleftharpoons$  neoangiogenesi  
Densità cellulare altissima: un tappeto  
di cellule

# CEUS first stage

Qualitative visual assessment “by eye”



CD-US



CEUS

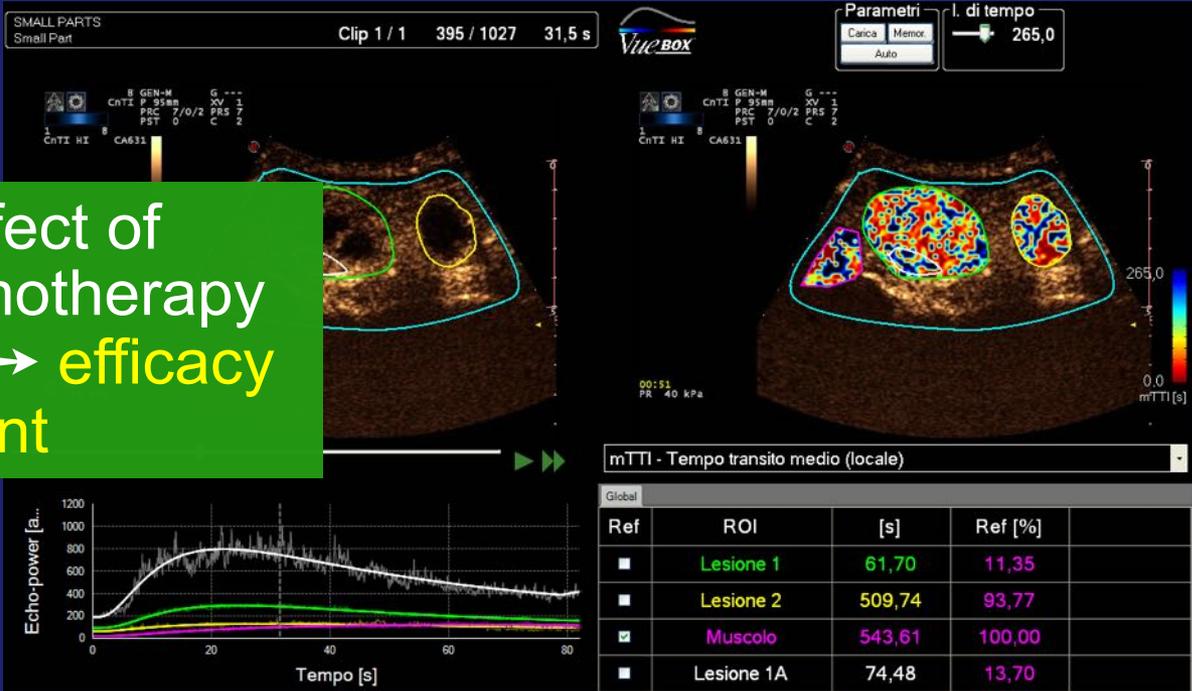
myxofibrosarcoma

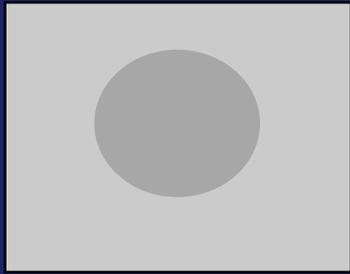
# CEUS second stage

## Quantitative assessment with a scanner software

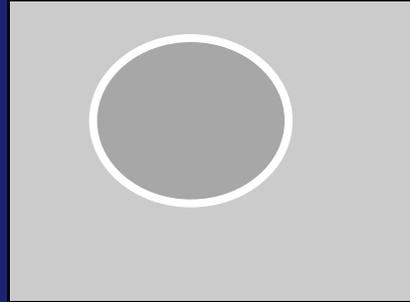
microbubble time-intensity curves

to monitor the effect of neoadjuvant chemotherapy and radiotherapy → efficacy of treatment

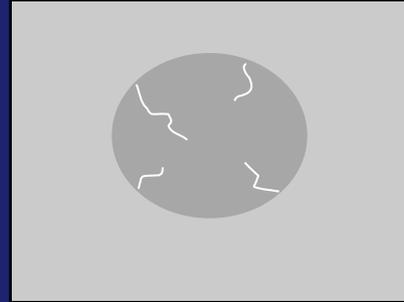




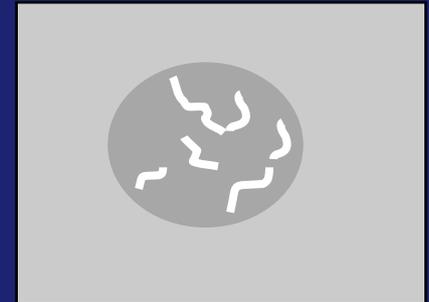
Pattern 1



Pattern 2

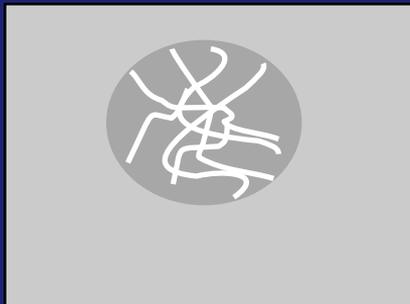


Pattern 3



Pattern 4

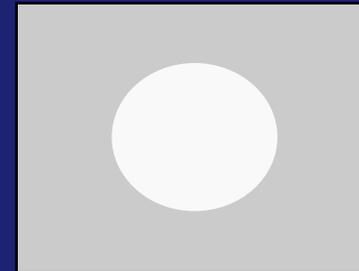
White=vessell



Pattern 5



Pattern 6



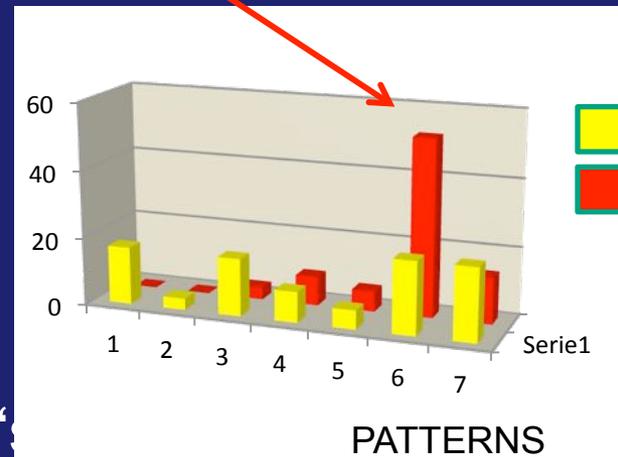
Pattern 7

54 patients: perfusion pattern with size and site is a new diagnostic tool to distinguish

Sensitivity 89% <sup>B/M</sup>  
 Specificity 85%  
 PPV 86% NPV 88%

Personal casistic:  
 205 patients:  
 117 m 88 b

Loizides A et all.  
 Eur Raiol 2012



♀ L.D 41 years old

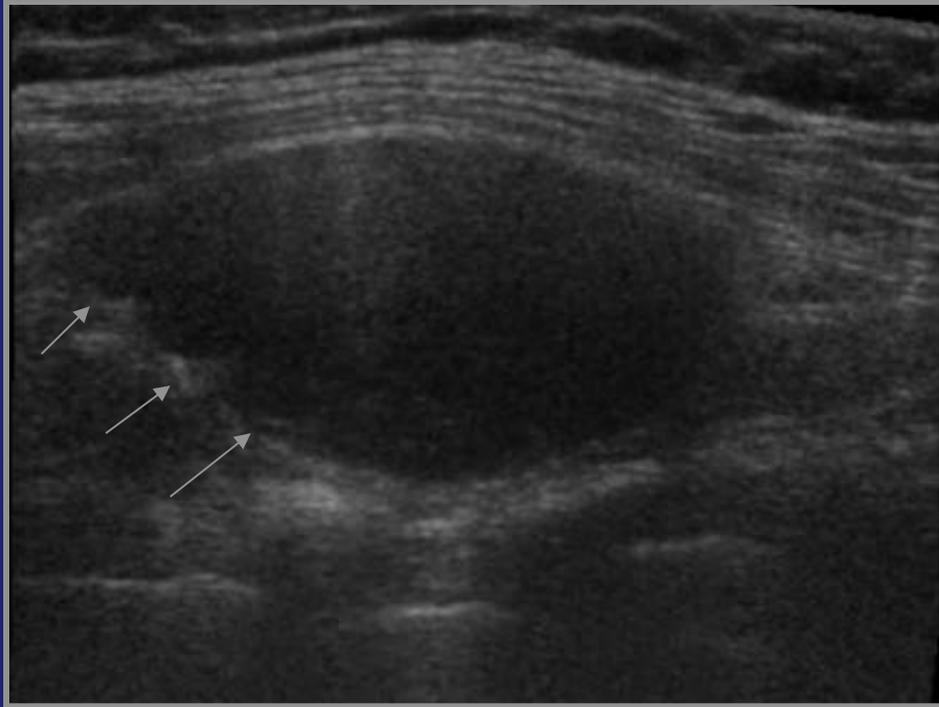


A painless mass, mobile on both superficial and deep planes present in the subscapular region for one year.

Based on site and clinical behaviour preliminary diagnosis:

Elastofibroma or Lipoma

♀ L.D 41 years old



Ext. field

Original Report

Elastofibroma Dorsi: Sonographic Findings

Stefano Bianchi<sup>1</sup>  
Carlo Martinoli<sup>2</sup>  
Ibrahim F. Abdelwahab<sup>3</sup>  
Nicola Gandolfo<sup>2</sup>  
Lorenzo E. Derchi<sup>2</sup>  
Sandro Damiani<sup>1</sup>

**OBJECTIVE.** Elastofibroma dorsi is a rare, slow-growing reactive pseudotumor of connective tissue typically located in the subscapular region. We report the sonographic characteristics of three cases of elastofibroma dorsi.

**CONCLUSION.** The sonographic appearance of elastofibroma dorsi consists of arrays of interspersed linear or curvilinear hypoechoic strands against an echogenic background. These findings reflect the alternating pattern of fibroelastic streaks and fat that is also detected with other imaging techniques and found on pathologic specimens. In the proper clinical setting, a prospective sonographic diagnosis can obviate the need for further imaging assessment and biopsy.

J Radiol 2005;96:1712-6  
© Editions Françaises de Radiologie, Paris, 2005

fait clinique ostéo-articulaire

Imagerie de l'élastofibrome dorsal

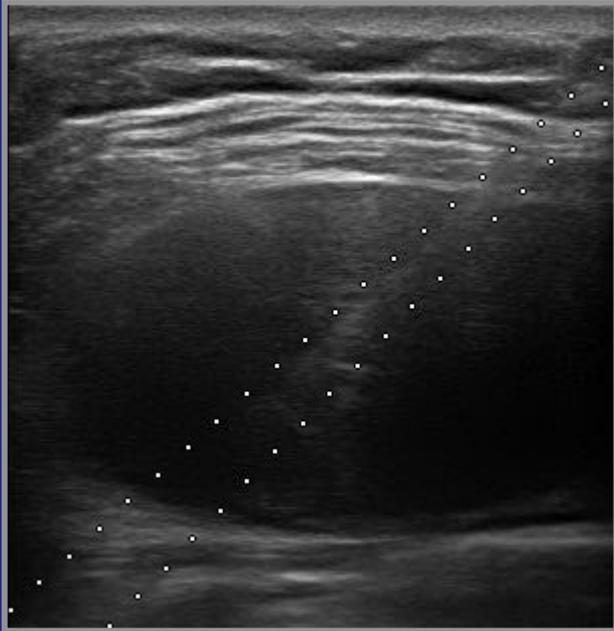
R. Alouini (1), M Allani (1), L Harzallah (2), M Bahri (3), C Kraiem (2) et K Tlili-Graies (4)



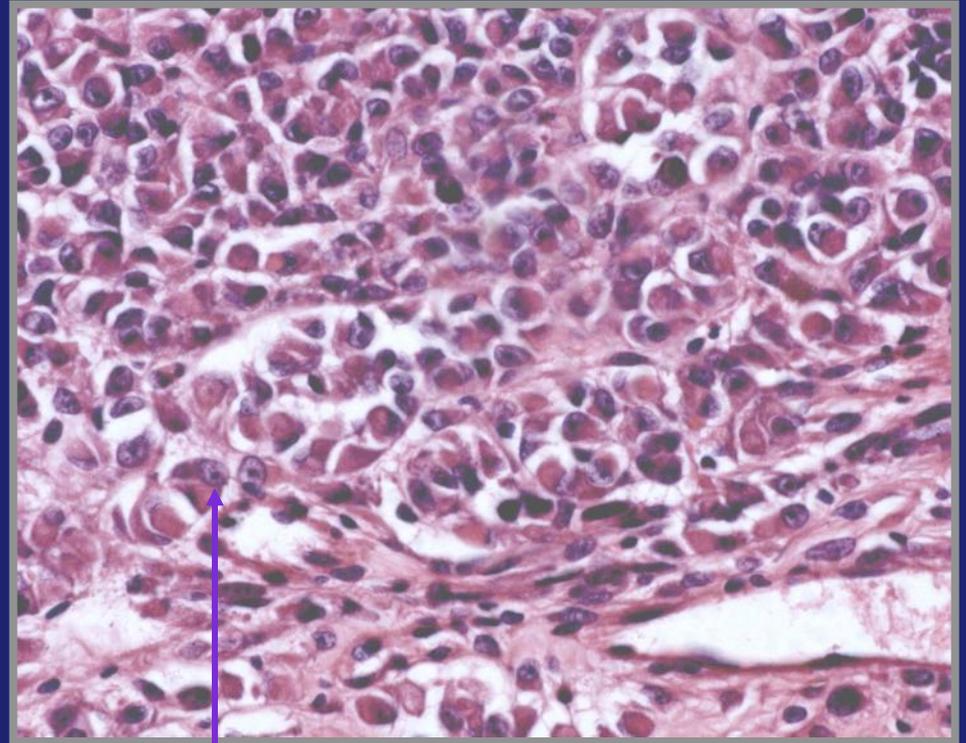
*De Marchi A, De Petro P, Faletti C e coll. (2003) Echocolor PowerDoppler with contrast medium to evaluate vascularization in lesions of the soft-tissues of the limbs. Chir Organi Mov.2003 Apr-Jun;88(2):225-31*

♀ L.D 41 years old

## Biopsy



## Histology

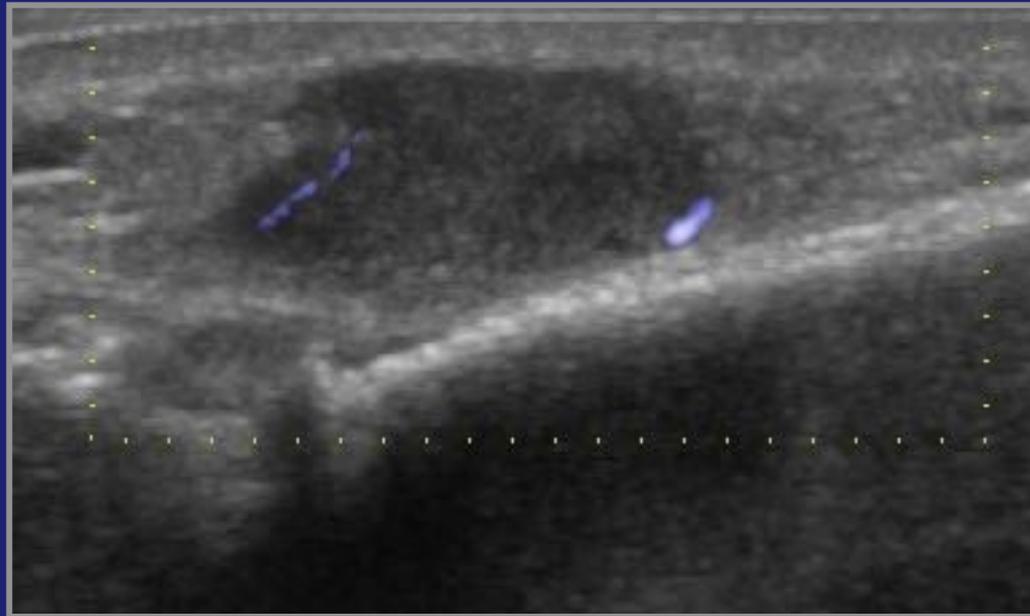


Cellular cannibalism

High-grade Sarcoma

# Follow-up

Post-excision outcome  
an anaplastic leiomyosarcoma excised by inappropriate  
surgical access



An extremely allergic subject  
no c.m.

e-Flow help us to evidence vascularization

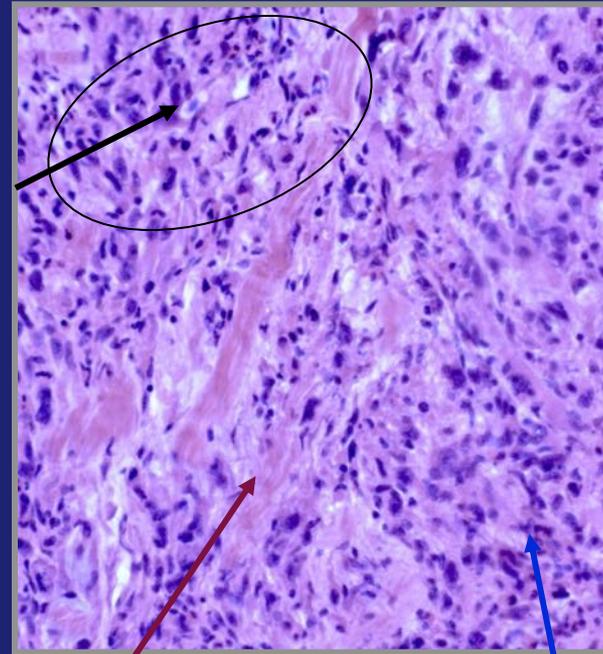
# Follow-up



# Follow-up

## Cellule neoplastiche Histology

### Biopsy



Fibrosis

Inflammatory Cells

## Relapse of the leiomyosarcoma

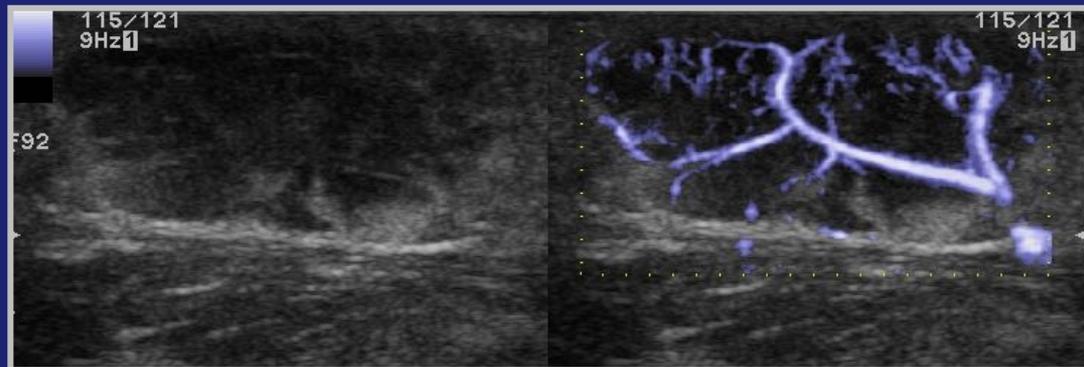
Liu JC, ChiouHJ, Chen WM, Chou YH, Chen TH, Chen W, Yen CC, Chiu SY, Chang CY. 2004

Sonographically guided core needle biopsy

of soft tissue neoplasms. J Clin Ultrasound. 2004 Jul-Aug; 32(6):294-8.

## Our **protocol** is based on the following procedure:

1. B-mode and colour Doppler sonography in basal condition  
( present on all the units!)
2. Contrast medium (SonoVue®-Bracco) bolus injection with  
dedicated software ( low MI)
3. Ultrasound-guided Needle Biopsy  
(sterile device on the probe)



**“Current imaging techniques provide some non invasive techniques to diagnose and stage suspected soft tissue sarcomas”**

**but....**

**“NO single approach is ideal for every tumor”**

**Knapp ER., Kransdorf MS., Letson G.D. : Diagnostic imaging update: soft tissue sarcomas- 2005**

# CT - MR

“The bundle, the periostium, the vascular wall areas are barriers against tumour invasion, therefore, a correct evaluation of the surgical margins is essential in the soft-tissue surgery ”

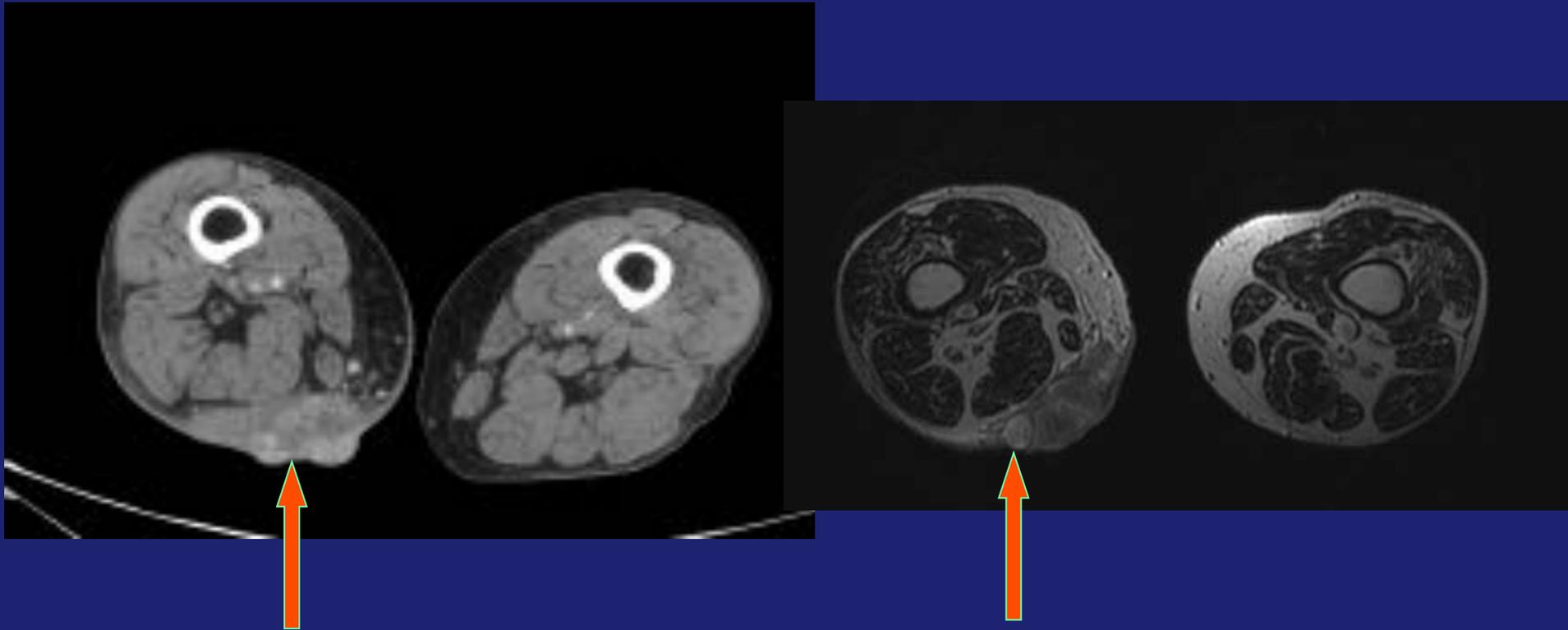
Enneking.W.F.



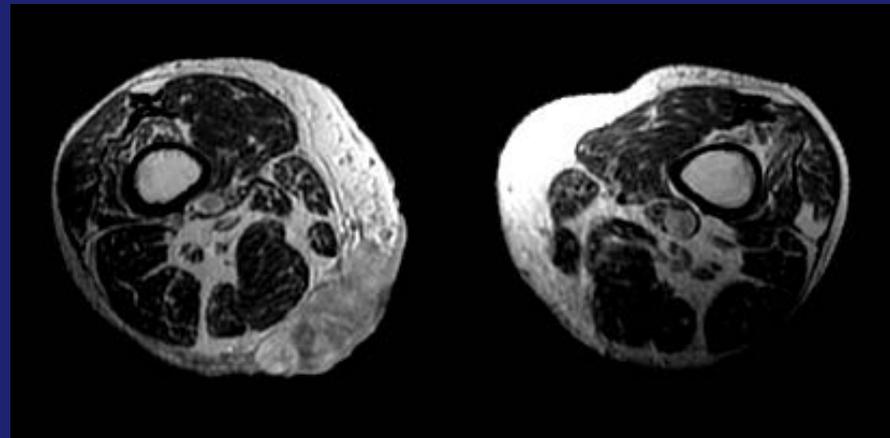
Compartmentality

Extra-compartmentality

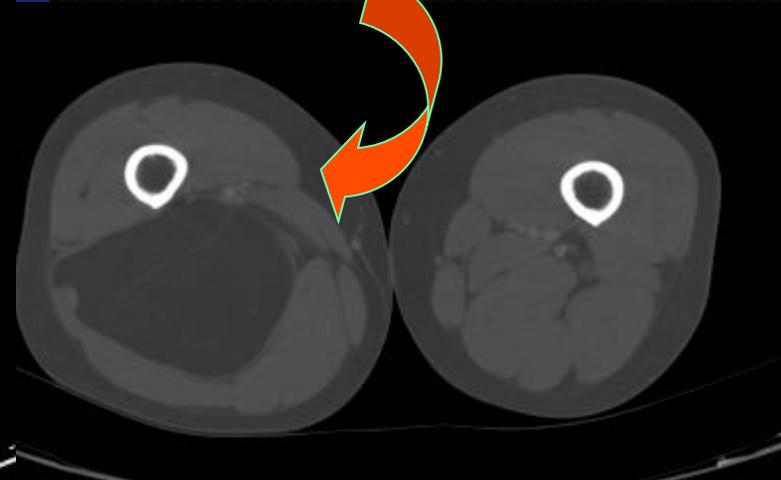
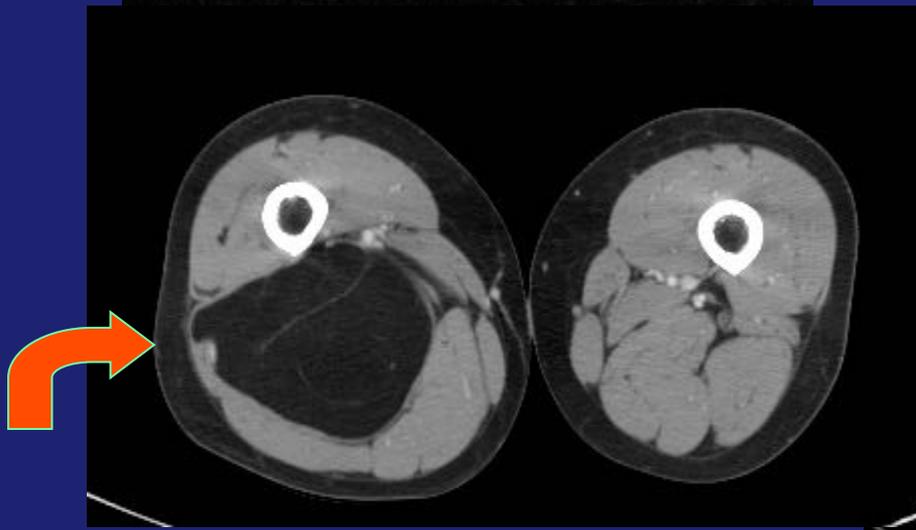
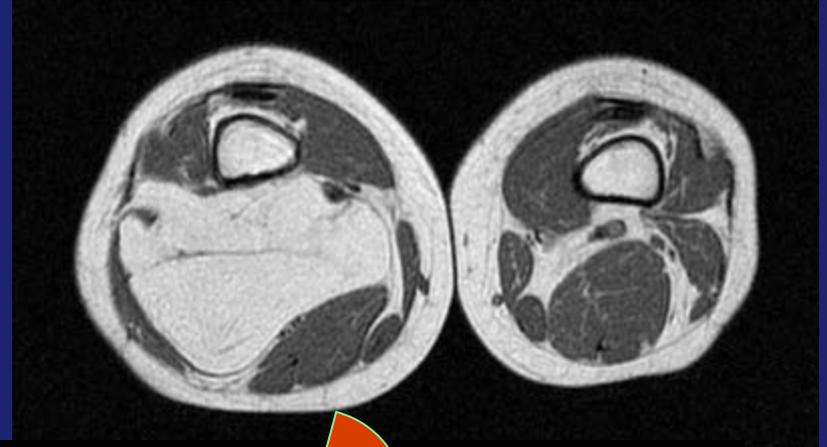
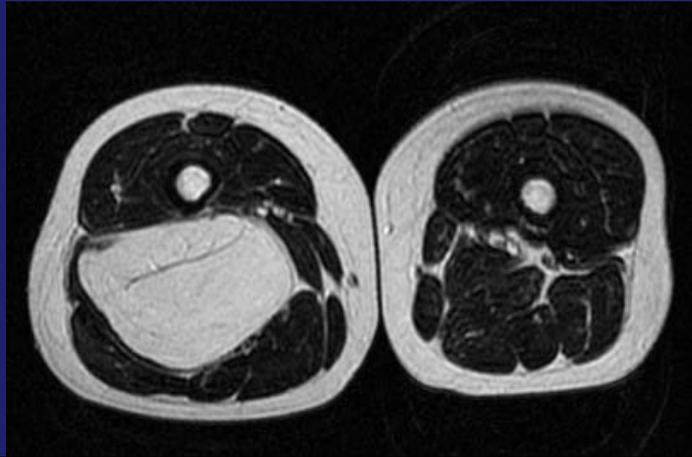
# EXTENSION...



High grade sarcoma

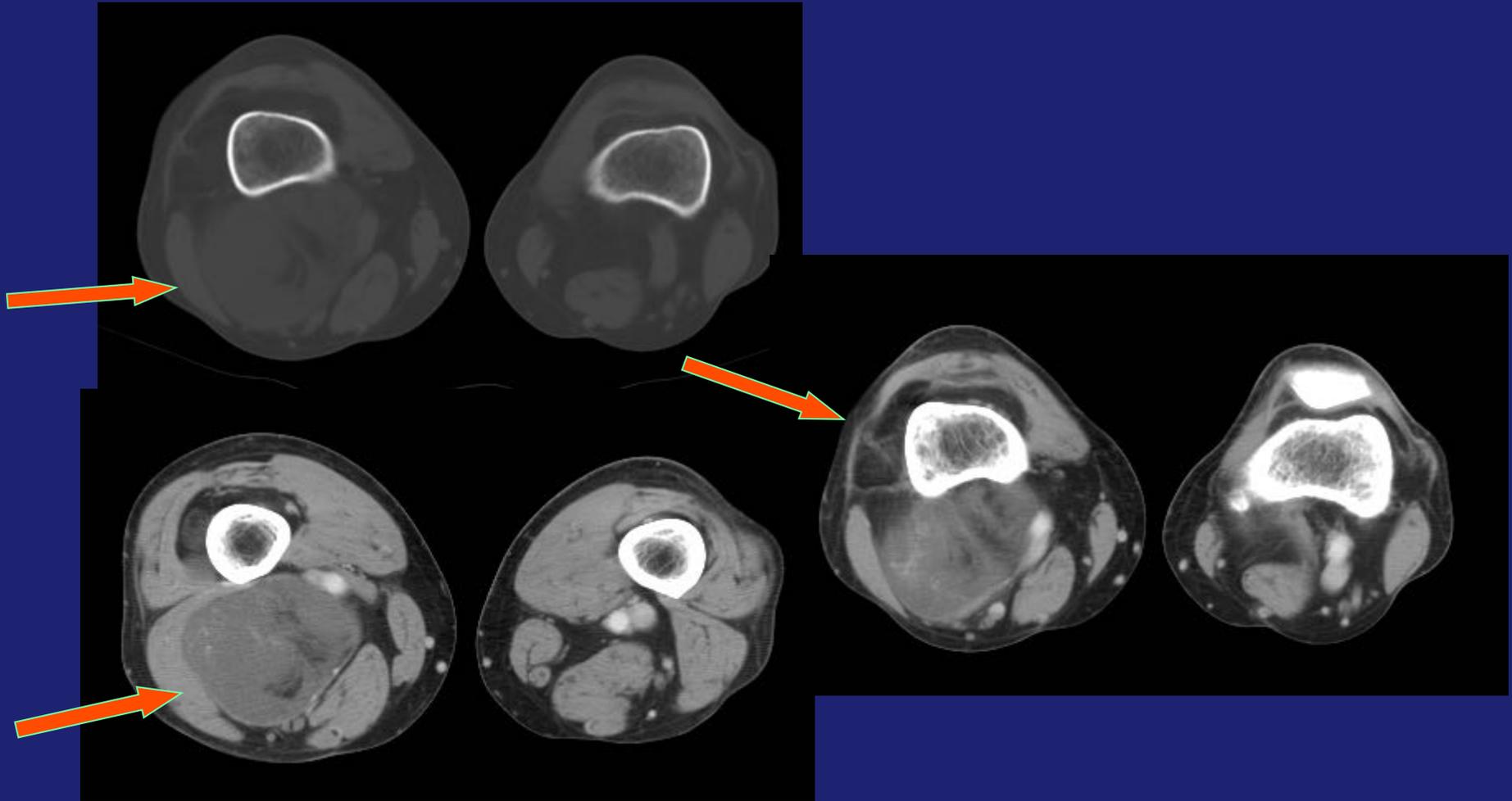


INTRA....



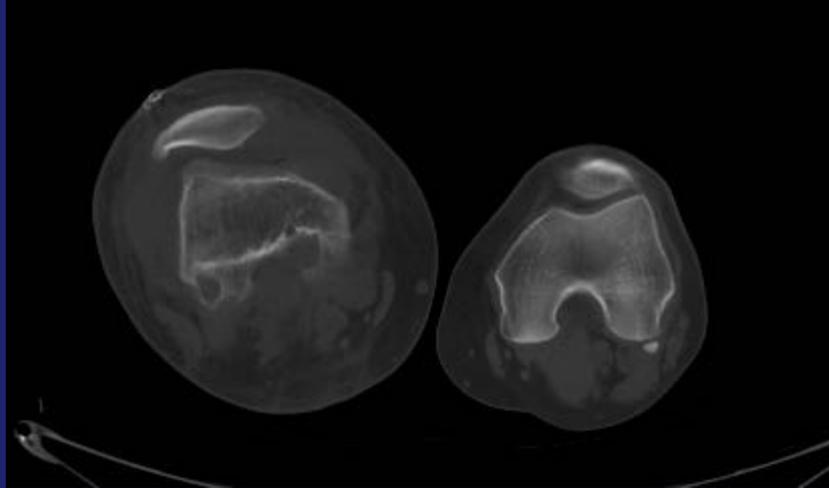
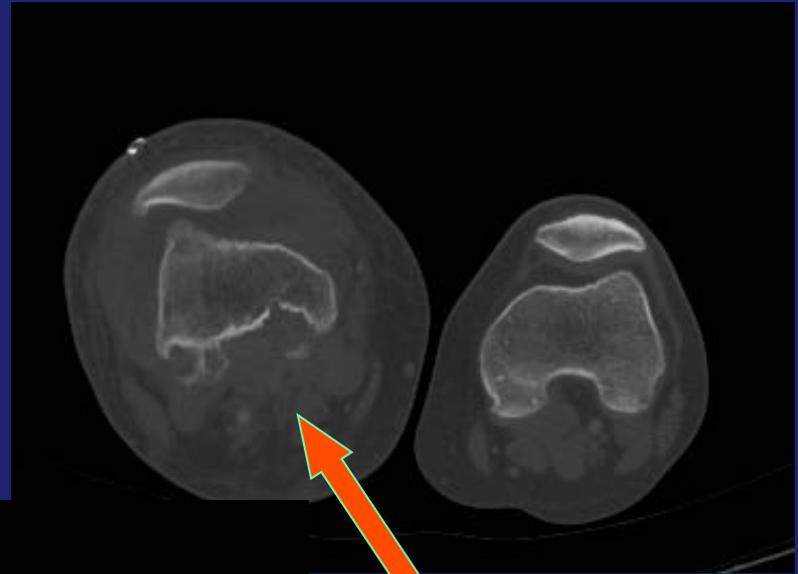
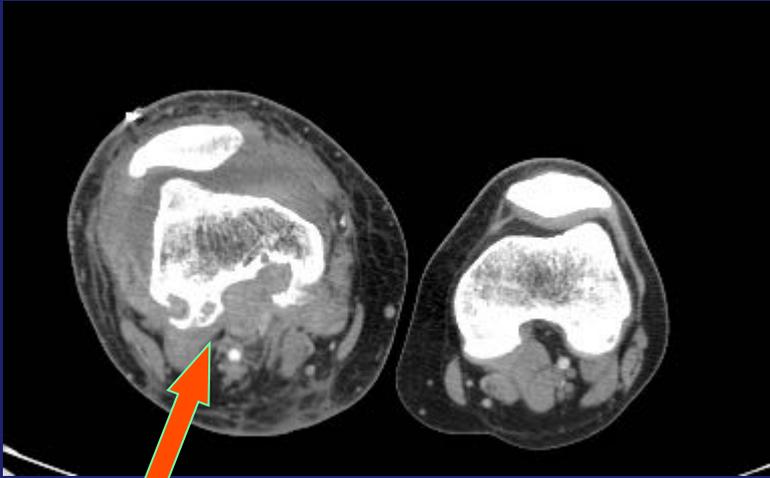
Liposarcoma lipoma-like

EXTRA.....in another muscular compartment



Liposarcoma

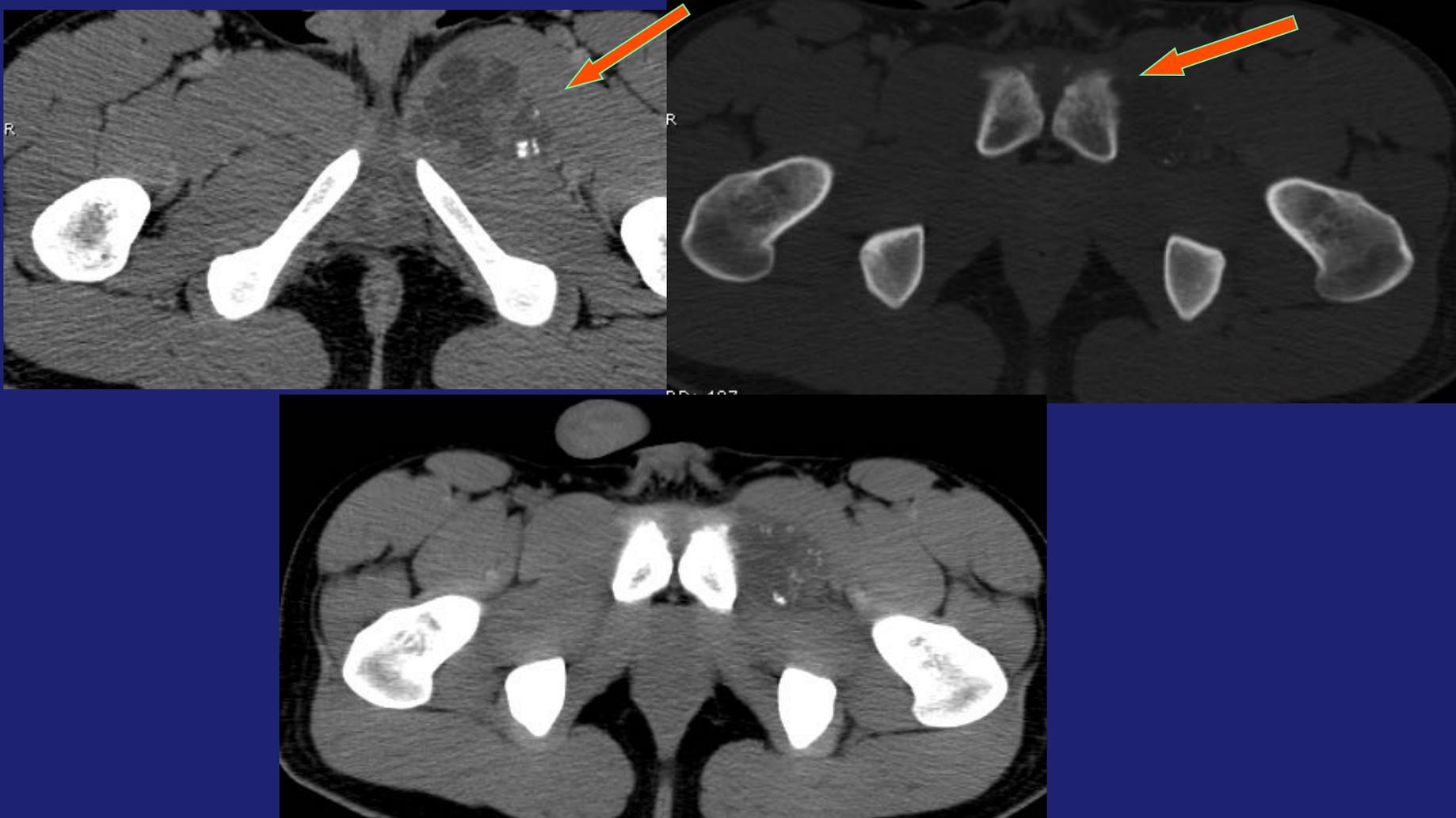
Extra...in the bone



Sarcoma



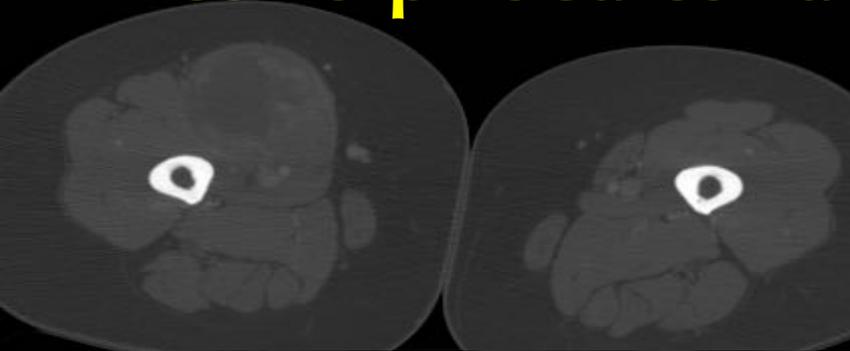
# Calcifications + Erosion



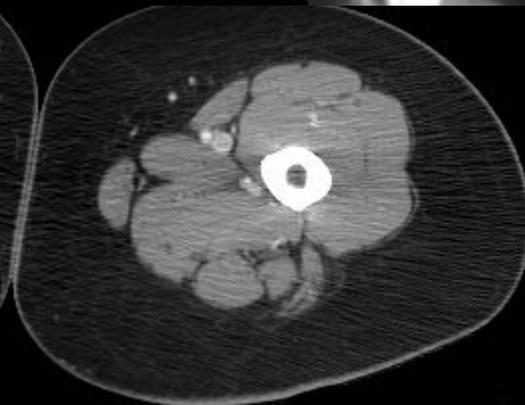
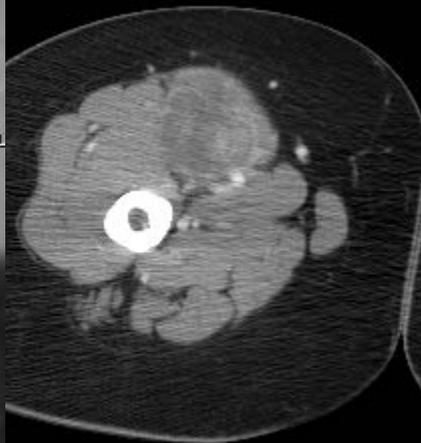
Pubic-bone excision!

# NEUROVASCULAR RELATIONSHIP ...

## Pleomorphic Sarcoma

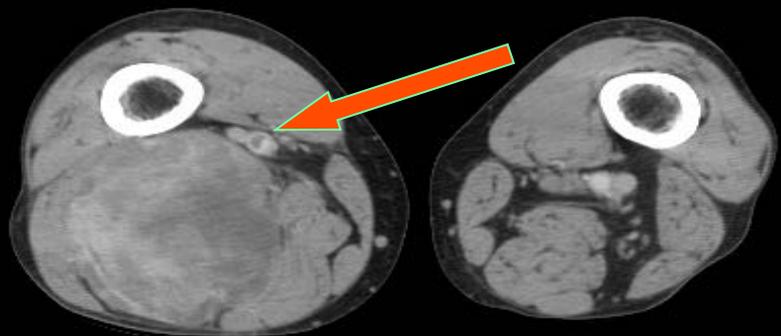
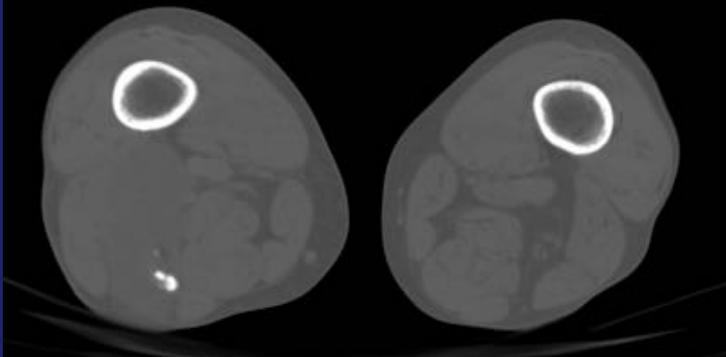
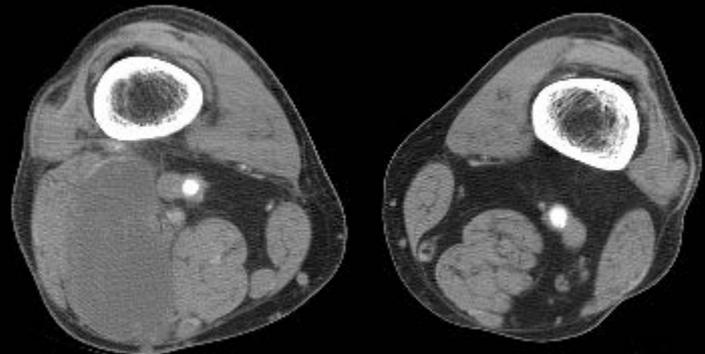
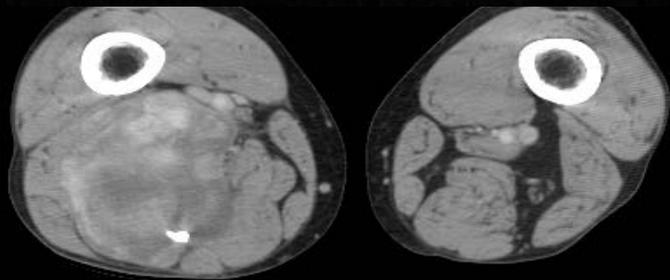
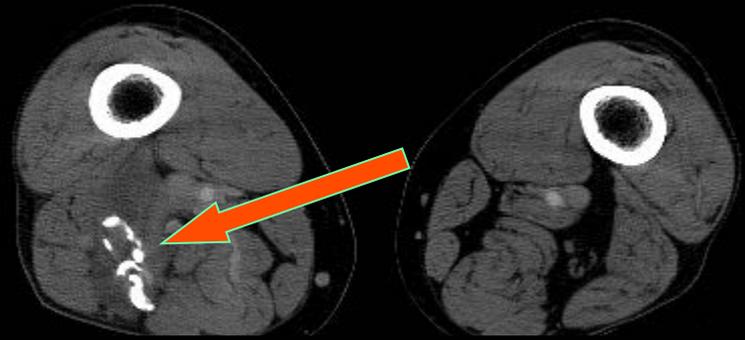
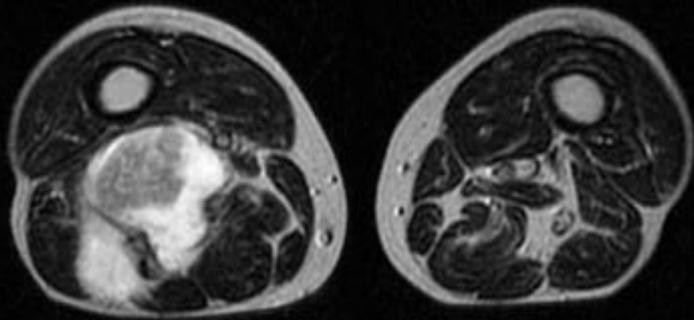


...englobed



... "en bloc" resection!!

# THROMBOSIS .....

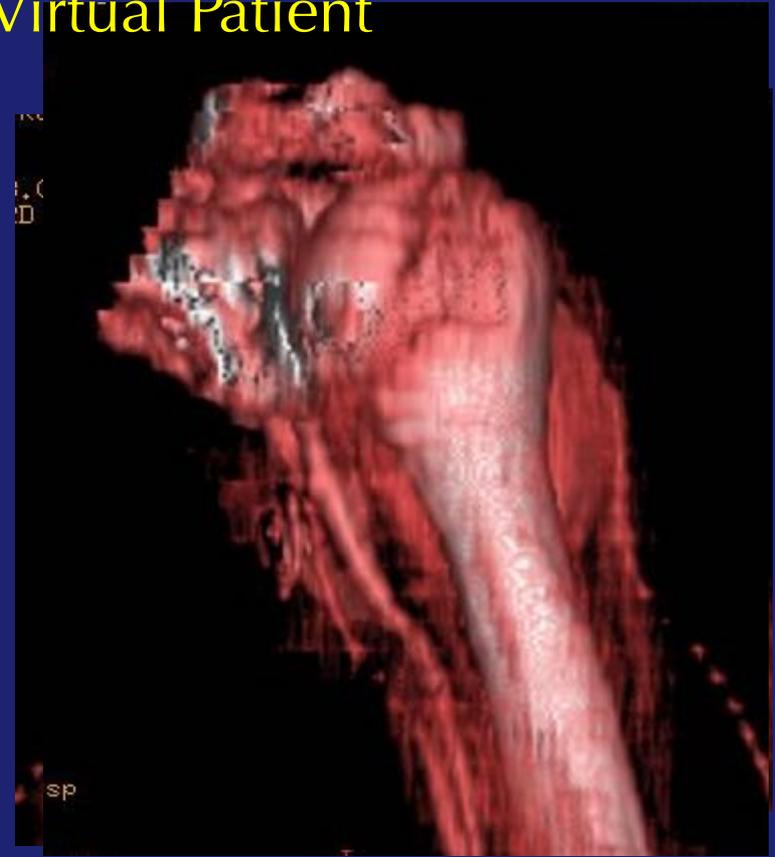
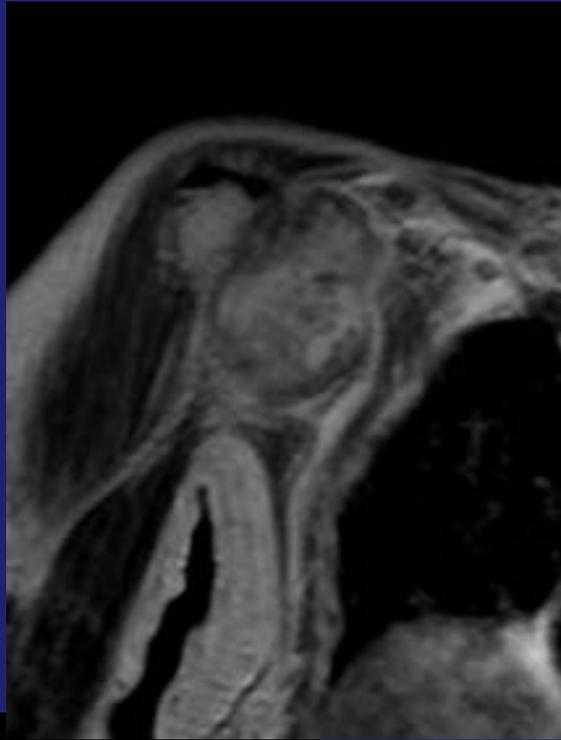
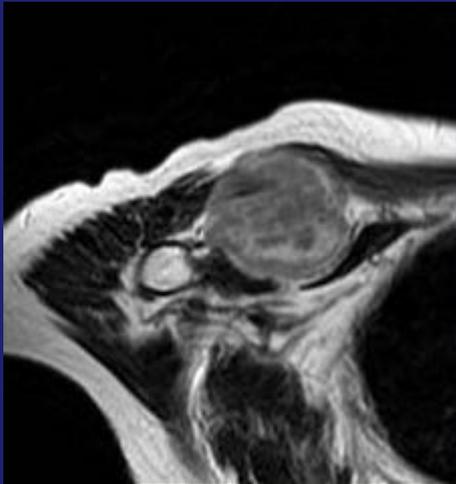


Mixoid Liposarcoma

3D CT



Virtual Patient

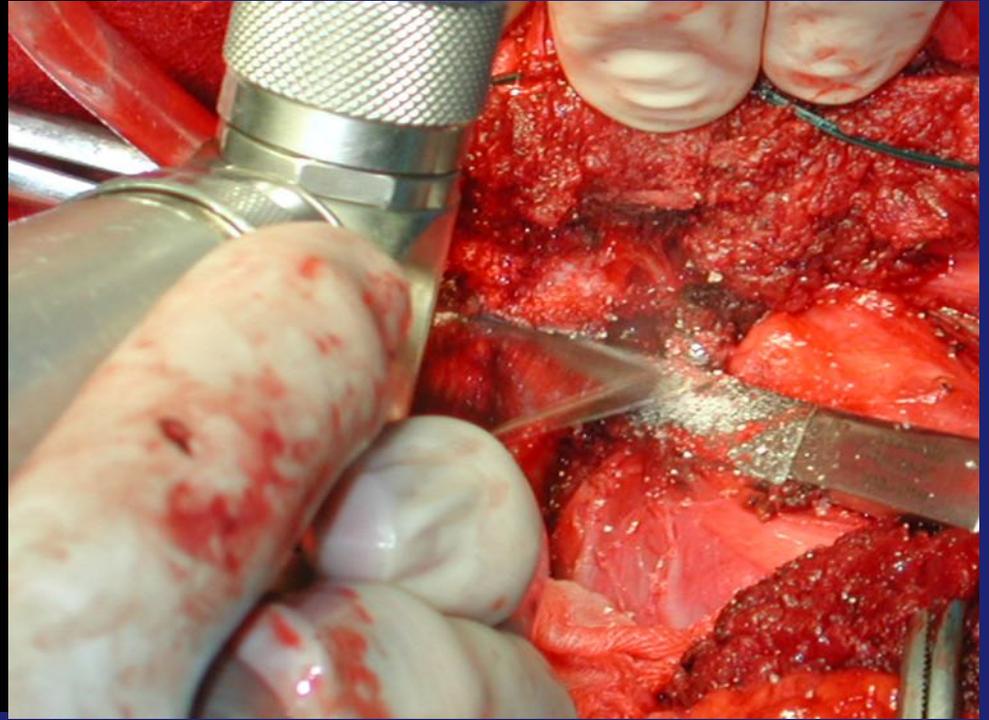


Pre-operative Planning

Surface 1 C.T.O. TORINO Radiologia D.E.A.  
Ex: 18055 S \*\*\*\*  
Se: 4 F 83 20232717  
Volume Rendering No cut: Feb 24 2004

DFOV 18.0 cm  
STANDARD  
570/2

No VOI  
kv 120  
mA N/A  
1.4  
5.0 mmHQ/5.0sp  
Tilt: 0.0  
09:52:34 AM  
W = 268 L = 103

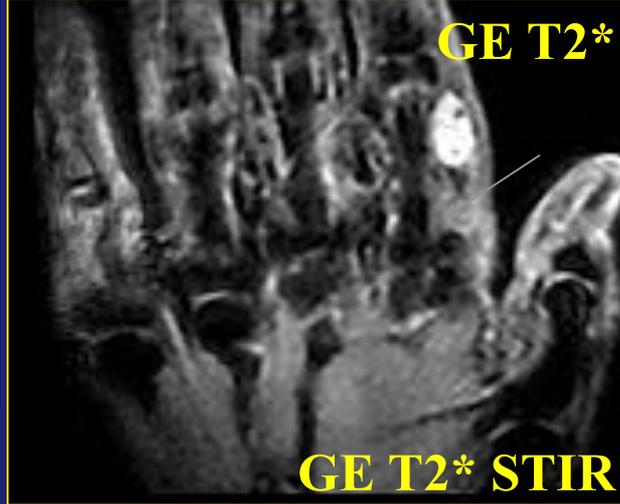
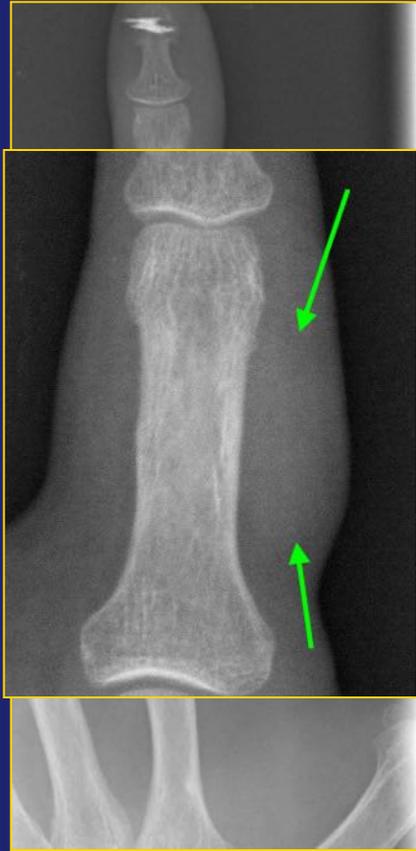


High Grade Sarcoma

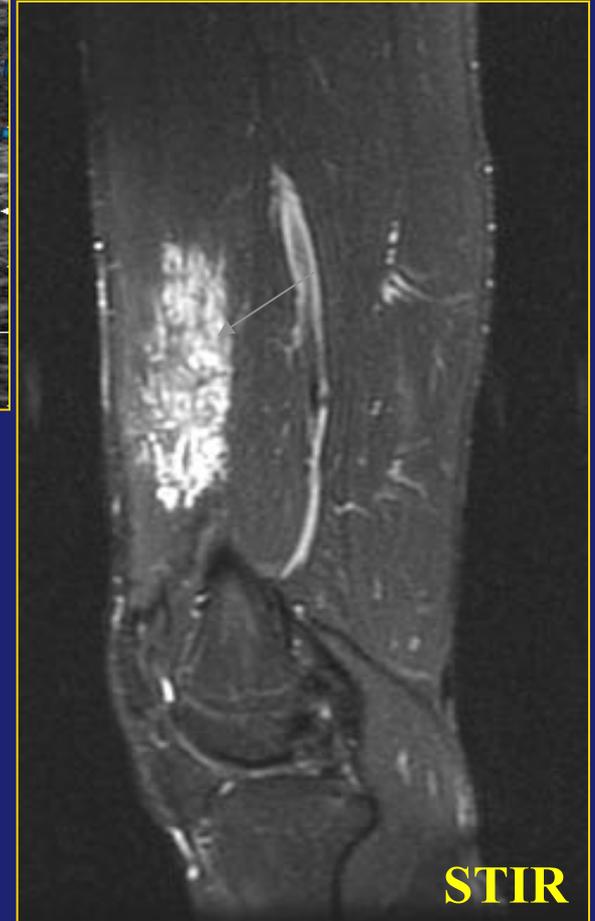
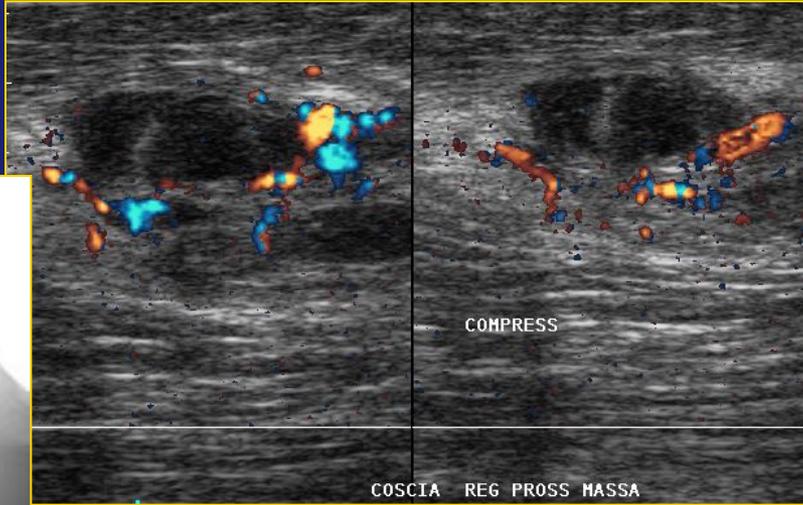
**MR**

“Gold standard” imaging modality for morphological and qualitative evaluation of tumors

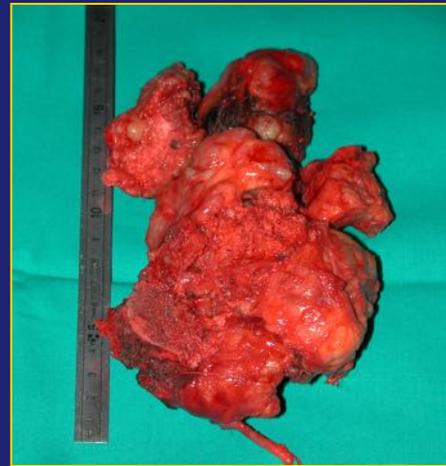
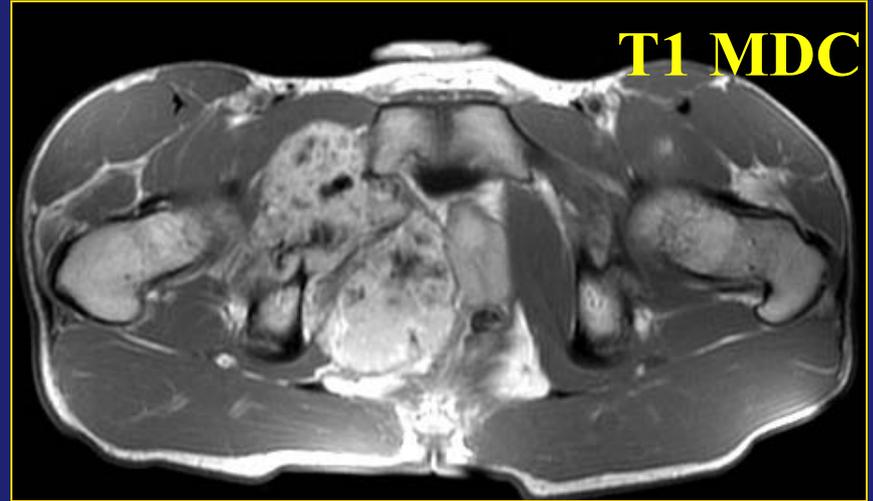
- multiplanarity
- high contrast resolution
- dynamic study



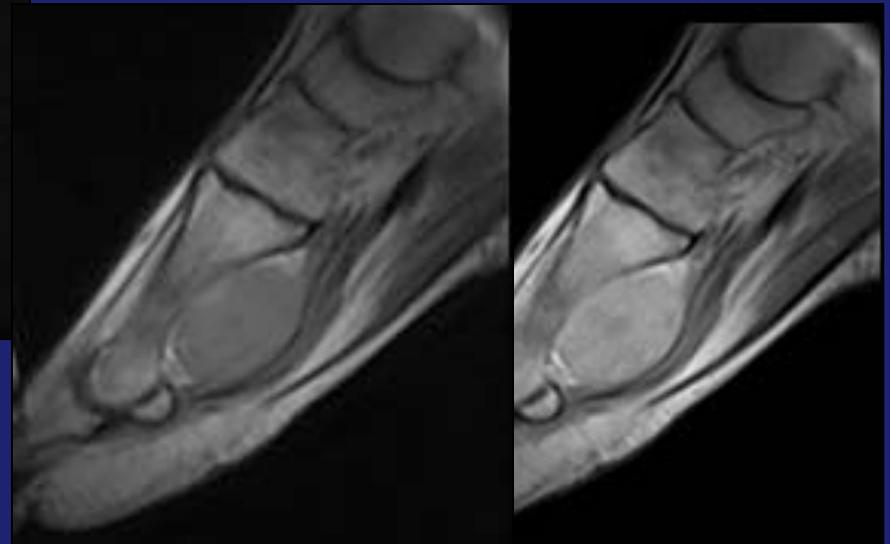
Emangioma



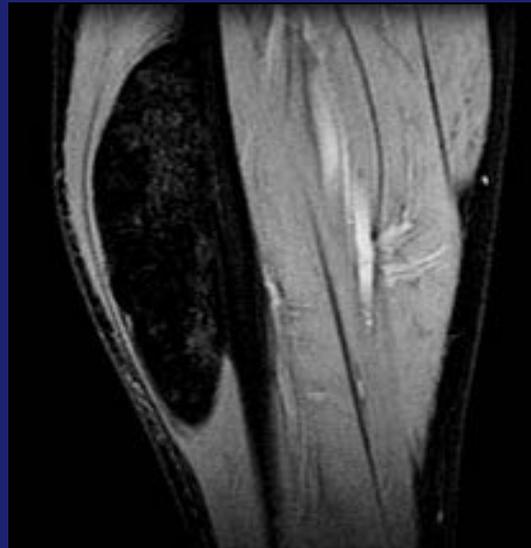
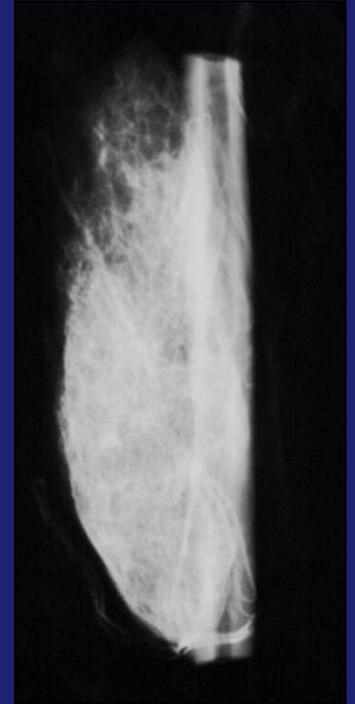
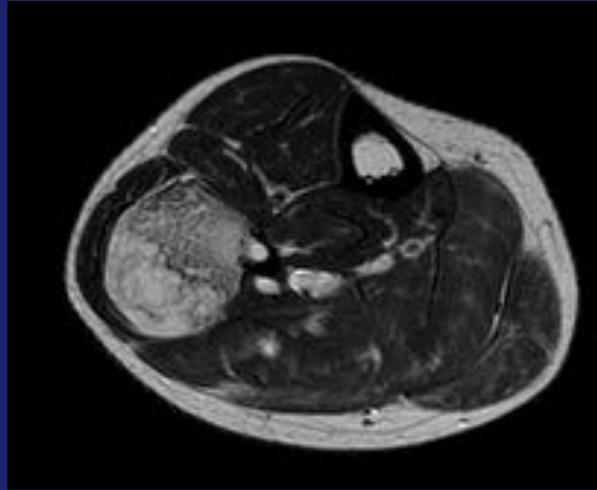
Angioma – MAV



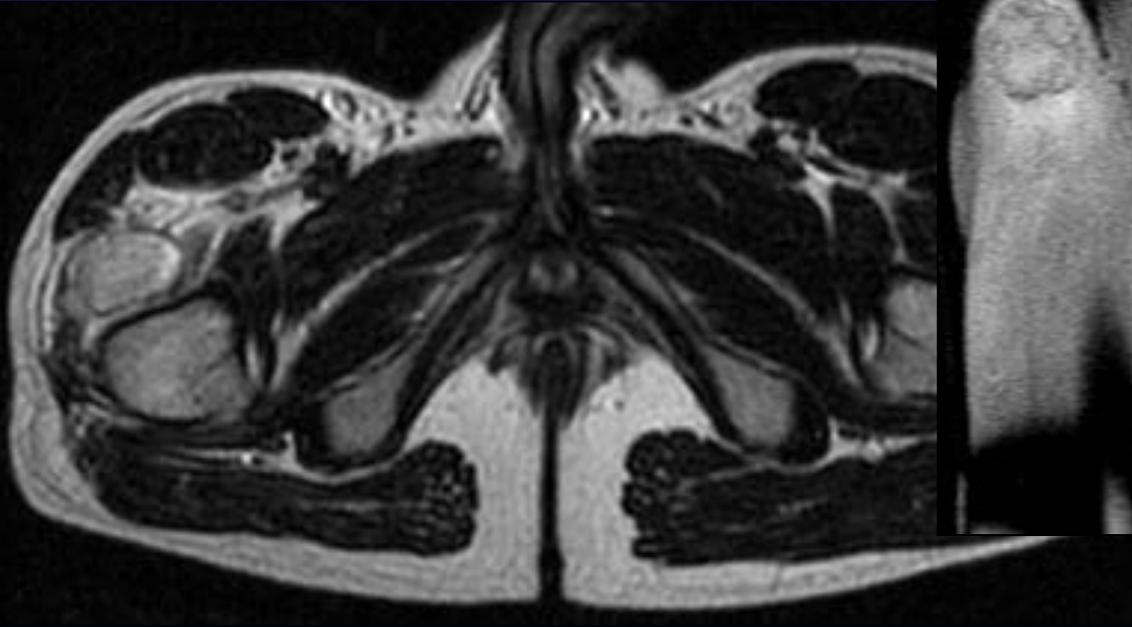
Calcific  
leiomyoma



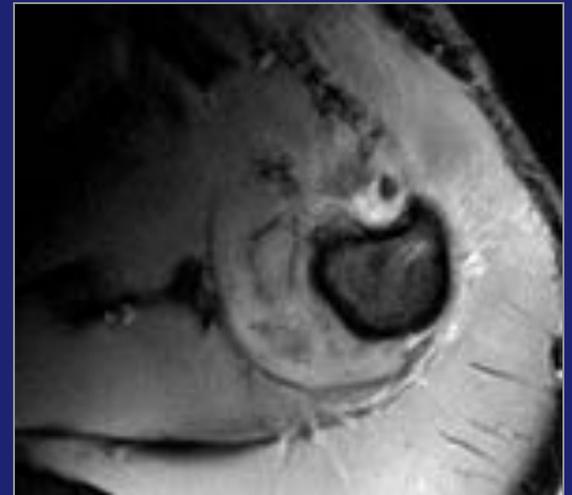
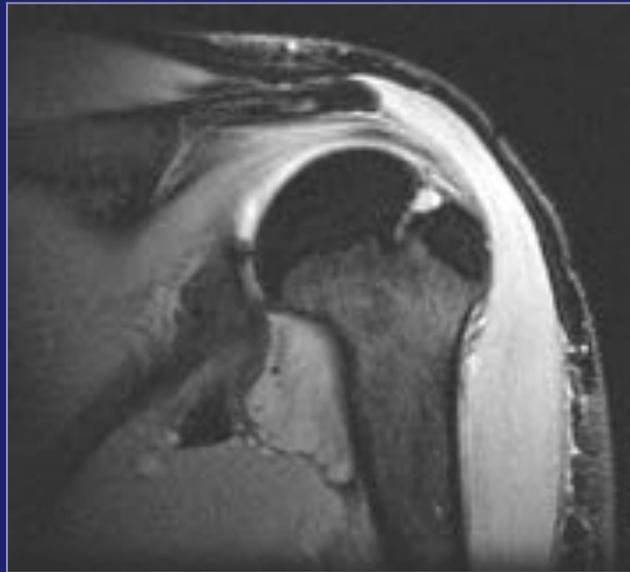
Synovial sarcoma



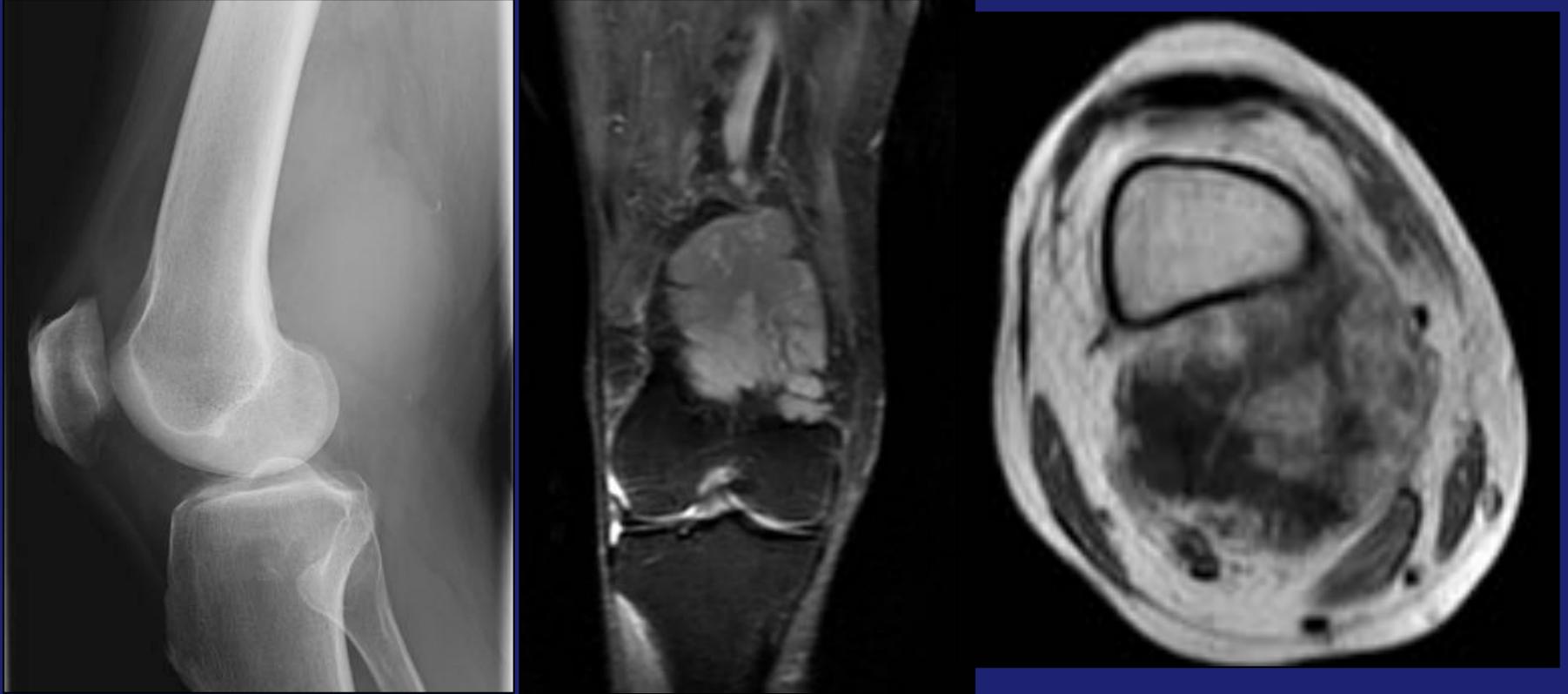
Ossified lipoma (lipoma periosteum?)



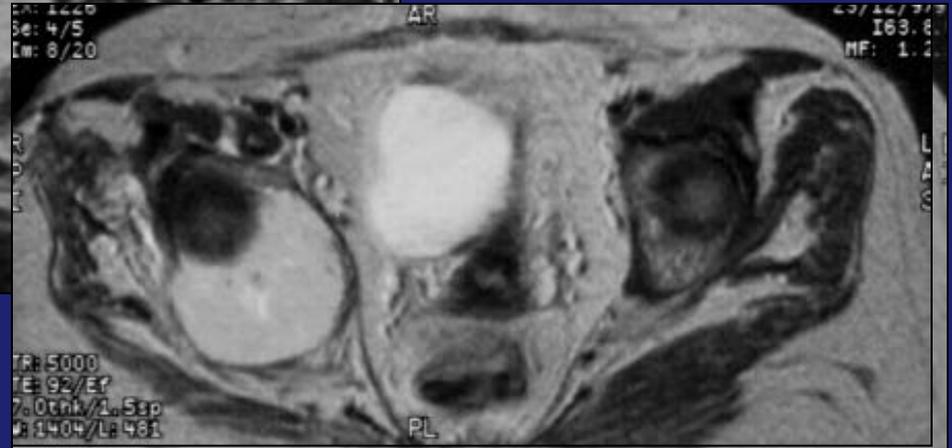
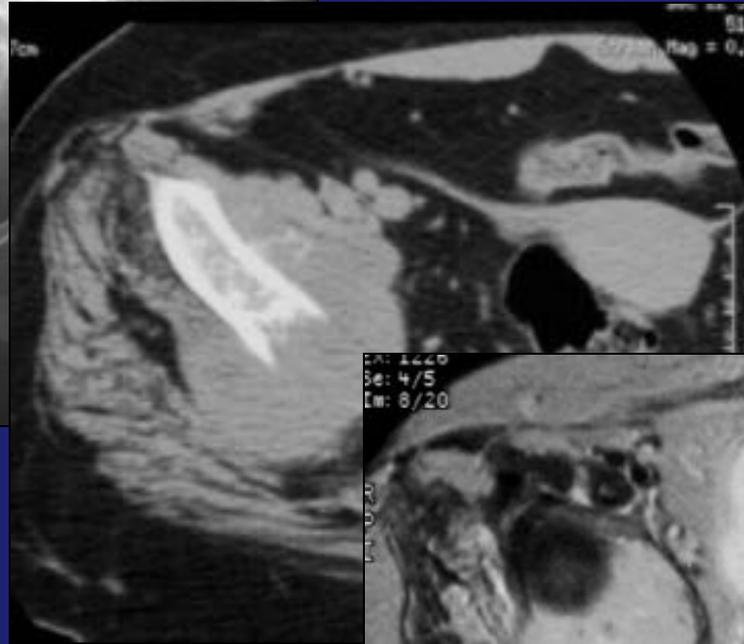
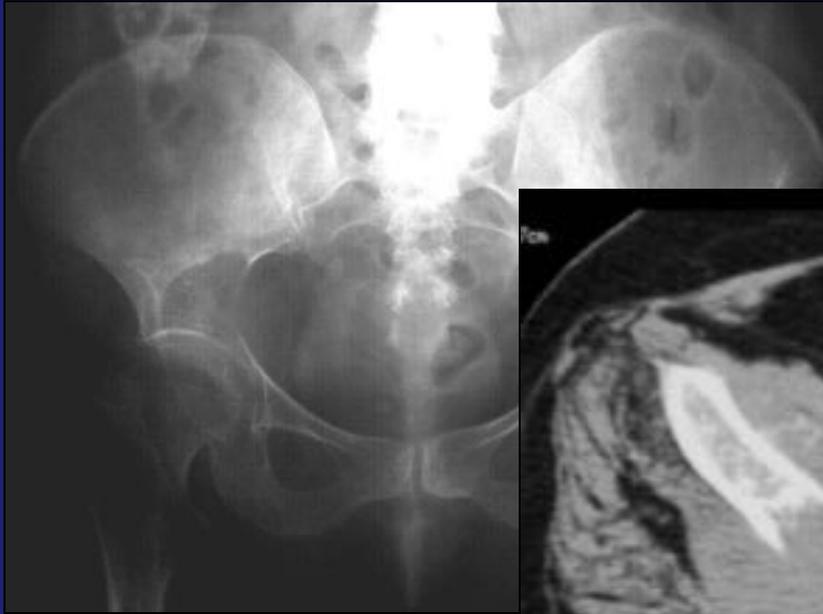
Ossificans myositis



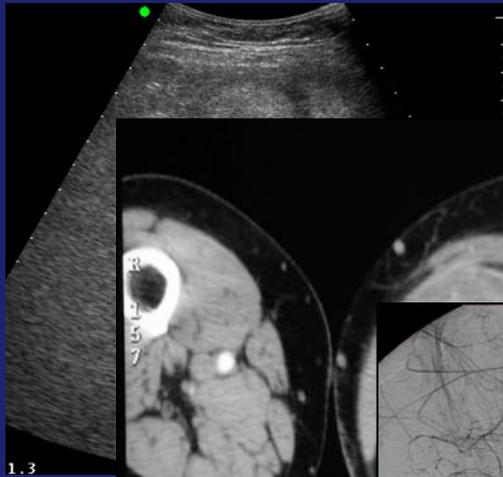
Pigmented villonodular synovitis



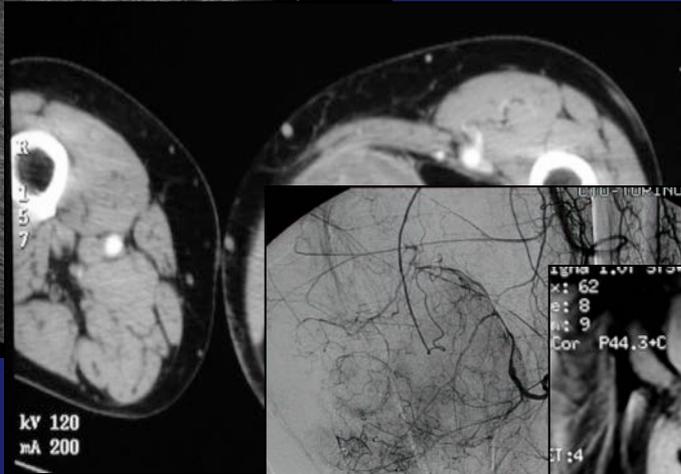
Extra-abdominal aggressive fibromatosis



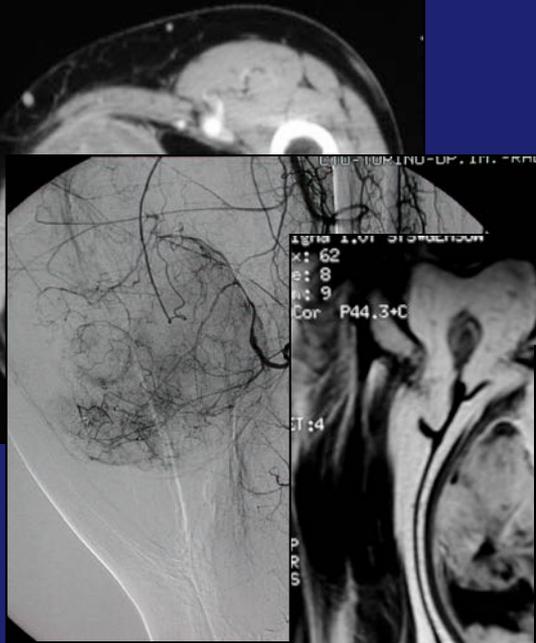
Neurofibrosarcoma



1.3



kV 120  
mA 200



T: 4  
FSE-TL/90  
TR: 7700  
TE: 18.3/EF  
EC: 1/1 31.2 Hz

1.01 31.2 Hz  
x: 62  
y: 8  
z: 9  
Cor P44.3+C



P40.6

90  
/EF  
15.6 kHz  
34

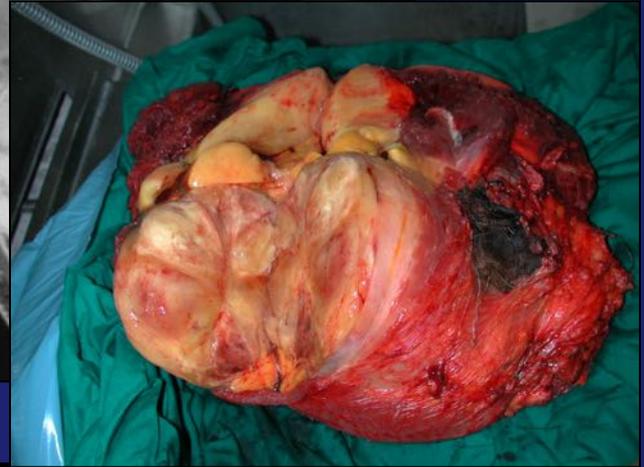


DOB: 02

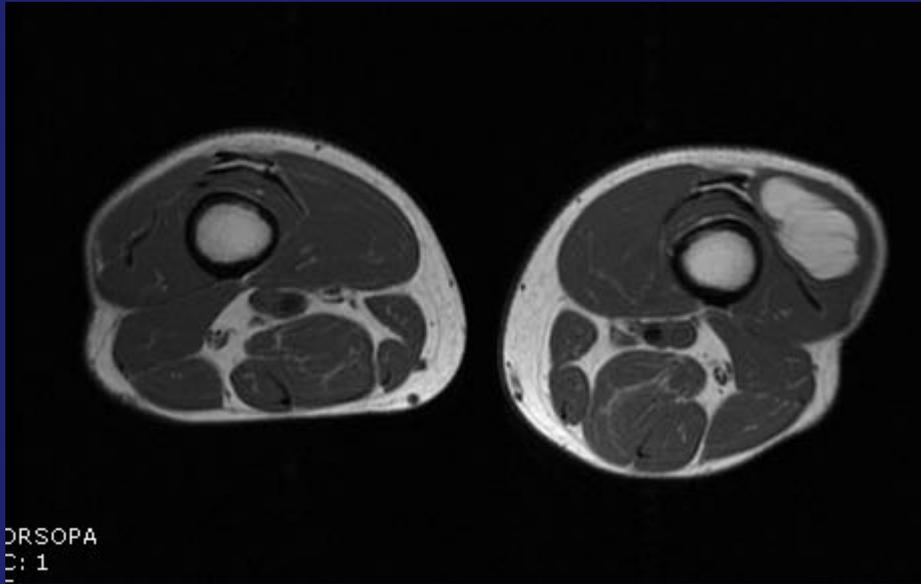
2

M

F



Liposarcoma



Lipoma → Lipoma like

**ANGIOCT**

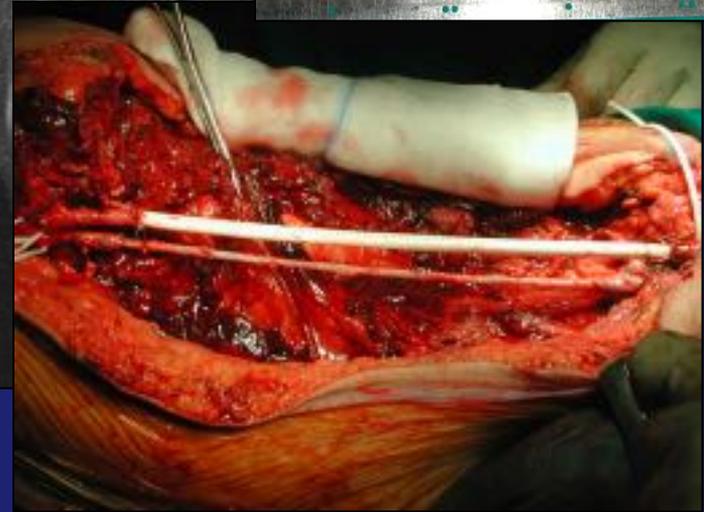


**ANGIO-MR:**

sequenze 3D ultraveloci:  
ricostruzioni multiplanari :  
valutazione morfologica



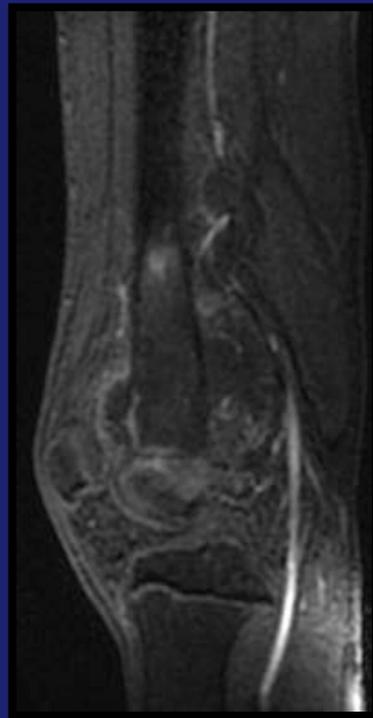
High-grade mixofibrosarcoma



High-grade pleomorphic sarcoma

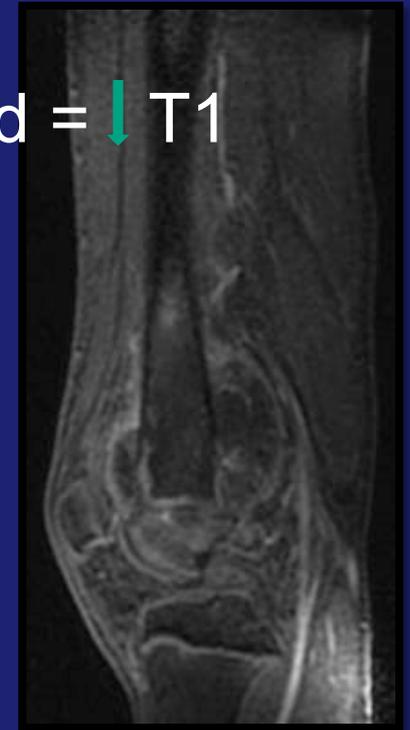
# RM dinamica (DCE MRI: dynamic contrast-enhanced MRI)

Iniezione e.v. mdc standardizzata



Acquisizione "dinamica" delle immagini post-mdc

Passaggio Gd = ↓ T1



Reddick WE et al. **Dynamic contrast-enhanced MR imaging evaluation of osteosarcoma response to neoadjuvant chemotherapy**. J Magn Reson Imaging 1995;5:689–694

Guo JY et al. **DCE-MRI pixel-by-pixel quantitative curve pattern analysis and its application to osteosarcoma**. Magn Reson Imaging 2009 Jul;30(1):177-84

## Iniezione e.v. mdc standardizzata

- utilizzo iniettore
- dose in fx del peso
- flusso 0,7- 3 ml/sec,  
seguiti da almeno 20 ml di fisiologica



MO Leach et al: *Assessment of antiangiogenetic and antivascular therapeutics using MRI: recommendations for appropriate methodology for clinical trials*. The British Journal of Radiology, special issue 76(2003),S87-91

## Acquisizione dinamica immagini

- **sequenze ultraveloci:**
  - ✓ eco di gradiente 3D,
  - ✓ fortemente T1 pesate (TR e TE minimi e gradienti spoiler che abbattano la magnetizzazione residua dopo ogni eco),
  - ✓ FA circa 40° per abbattere il segnale dei tessuti stazionari in cui non è presente la riduzione del T1 determinata dal Gd,
  - ✓ preimpulsi di inversione per presaturare il grasso senza allungare il TA,
  - ✓ tempo di acquisizione < 30 sec per separare la fase arteriosa da quella venosa

Es: SPGR 3D, LAVA, etc
- acquisizione immagini contemporaneamente o con tempo di ritardo rispetto all'inizio dell'iniezione, calcolandolo con la tecnica del *test-bolus* (sequenze centriche : ritardo=TA, sequenze sequenziali ritardo=TC-1/3TA)
- ripetute continuamente per i primi 150 sec e poi anche non continuamente per almeno 8 minuti

# Curve Intensità di Segnale – Tempo:

Quali applicazioni? a. Tipizzazione b. Ristadiazione

Reddick WE, Bhargava R, Taylor JS, Meyer WH, Fletcher BD. Dynamic contrast-enhanced MR imaging evaluation of osteosarcoma response to neoadjuvant chemotherapy. *J Magn Reson Imaging* 1995;5:689–694.

Woude HJ, Verstraete KL, Bloem JL: Muscoloskeletal tumors: does fast dynamic contrast-enhanced subtraction MR imaging contribute to the characterization? *Radiology* 1998, 208(3):821-8

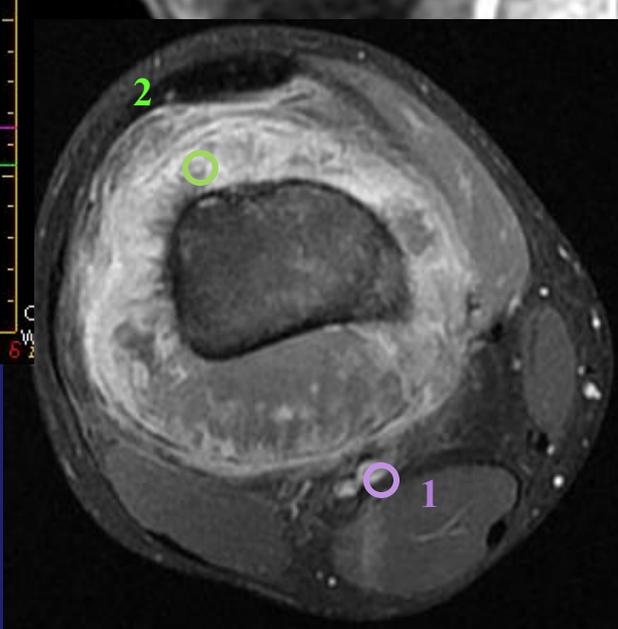
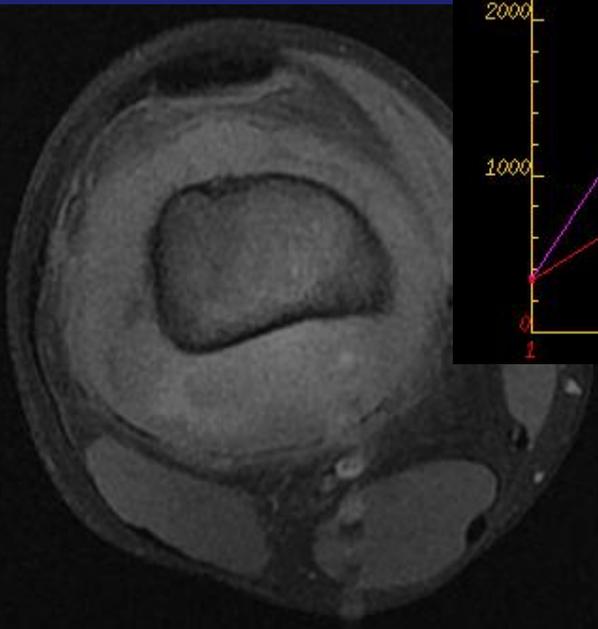
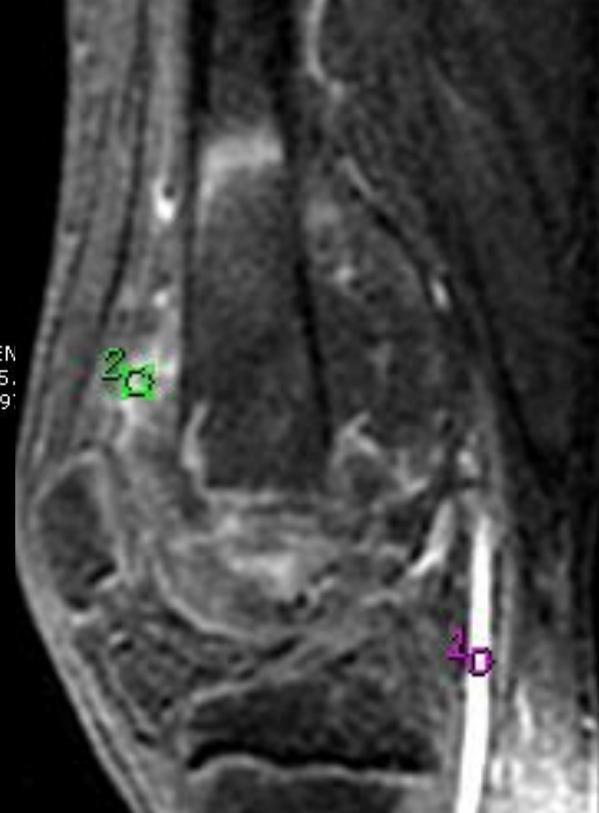
Woude HJ, Verstraete KL, Bloem JL: Dynamic contrast-enhanced MR imaging of muscoloskeletal tumors: basic principles and clinical applications. *J Magn Reson Imaging* 1996;6(2):311-21

Egmont-Petersen M, Hogendoorn PC, Van der Geest RJ, et al. Detection of areas with viable remnant tumor in postchemotherapy patients with Ewing's sarcoma by dynamic contrast-enhanced MRI using pharmacokinetic modeling. *Magn Reson Imaging* 2000;18:525–535.

Dyke JP, Panicek DM, Healey JH, et al. Osteogenic and Ewing sarcomas: estimation of necrotic fraction during induction chemotherapy with dynamic contrast-enhanced MR imaging. *Radiology* 2003;228:271–278.

Guo JY, Reddick WE. DCE-MRI pixel-by-pixel quantitative curve pattern analysis and its application to osteosarcoma. *Magn Reson Imaging* 2009 Jul;30(1):177-84.

# Approccio semiquantitativo: Curve Intensità di Segnale- Tempo



# Approccio quantitativo: **Perfusione**

## Parametri di farmacodinamica per il microcircolo

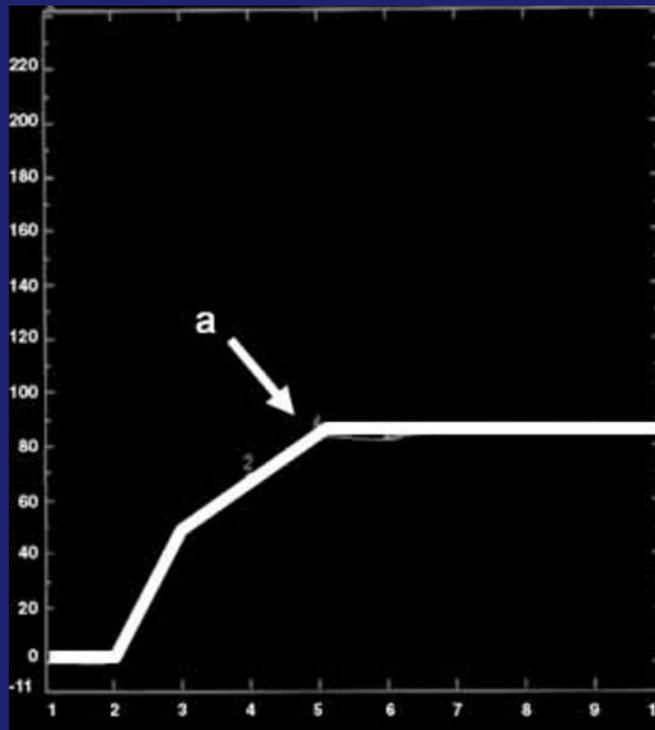
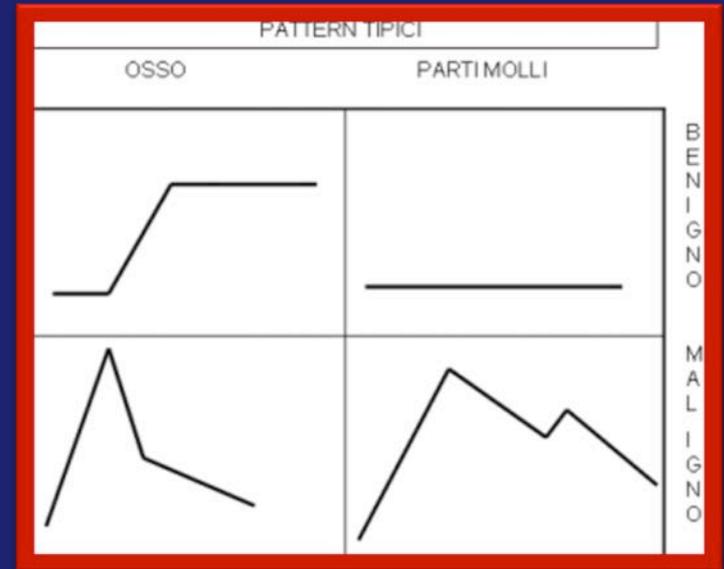
- Transfer constant (**K<sub>trans</sub>**, min<sup>-1</sup>): riflette il circolo (perfusione) e il passaggio del mdc attraverso l'endotelio vascolare (permeabilità)
- Initial area under the gadolinium concentration time curve (**IAUGC**, mM<sub>Gd</sub>.min; over first 1.5 min after contrast arrival)
- **V<sub>b</sub>**: volume ematico (mdc nei vasi)
- **V<sub>e</sub>**: volume extracellulare (mdc nello spazio interstiziale), dipende dalla cellularità
- **PS**: permeabilità dell'endotelio

# RM dinamica

## a. Tipizzazione

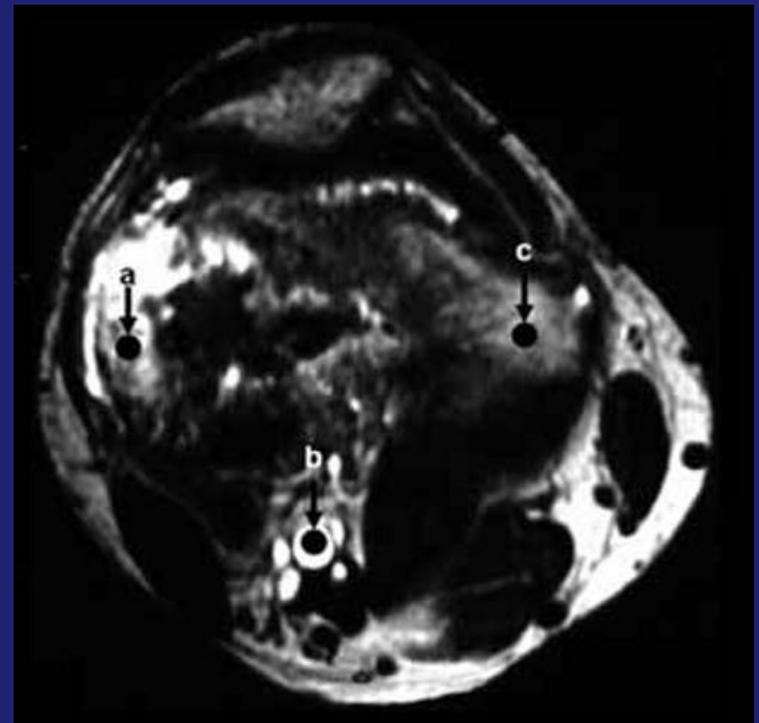
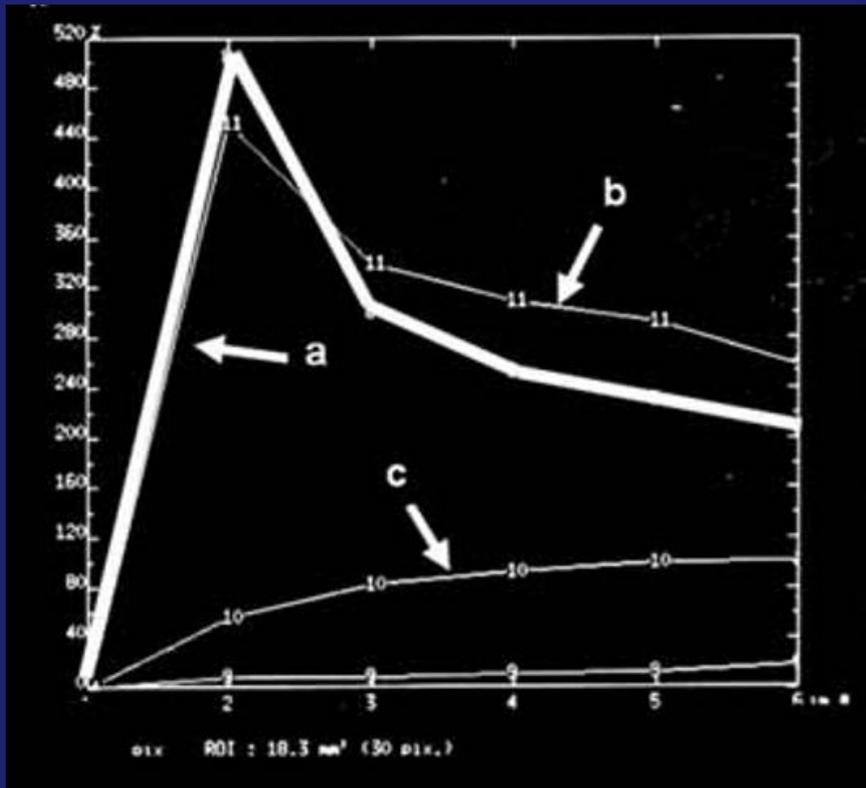
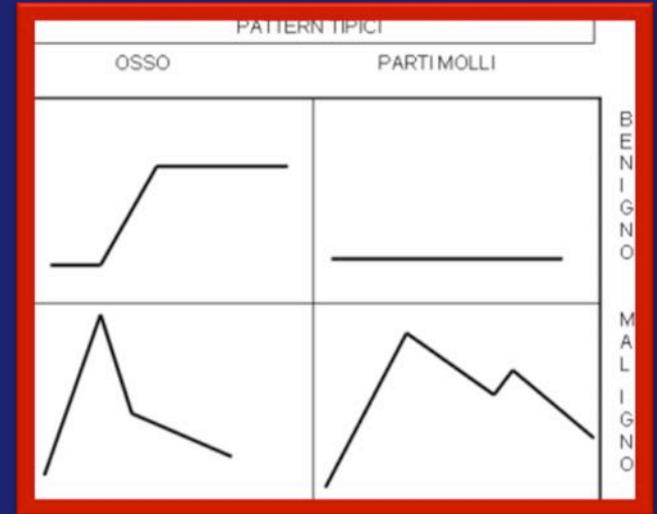
A.Barile, G. Regis, R. Masi, M.Maggiori, A. Gallo, C. Faletti, C. Masciocchi

Musculoskeletal tumours: preliminary experience with perfusion MRI. Radiol med (2007) 112:550–561



Lesione benigna:  
**encondroma**

# Lesione ossea maligna: Osteosarcoma osteoblastico



## RM dinamica:

### b. Valutazione della risposta tumorale

**Quale** valutazione della risposta tumorale nei sarcomi delle parti molli alla terapia neoadiuvante?



Il diametro massimo (RECIST) non può essere l'unico parametro valutato dal radiologo e dall'oncologo

#### **Purpose**

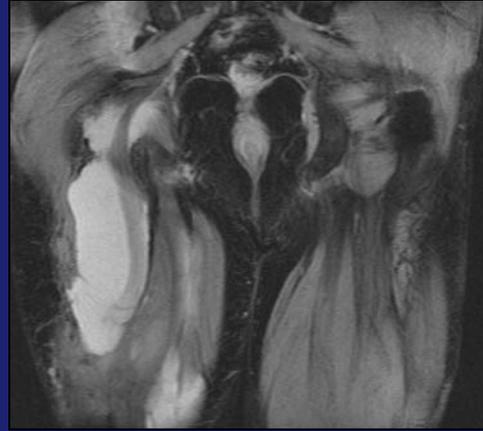
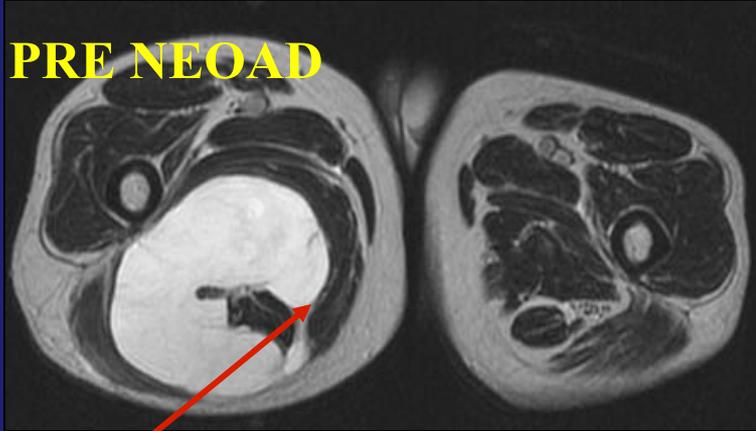
Response Evaluation Criteria in Solid Tumors (RECIST) are insensitive in evaluating imatinib-treated **gastrointestinal stromal tumors (GISTs)**. Response by Choi criteria, a 10% decrease in size or a 15% decrease in density on contrast-enhanced CT, correlated well in a small training set of patients who showed response as measured by positron emission tomography, and was more predictive of time to **tumor progression (TTP)** than response by RECIST. This study was designed to validate these observations in an independent data set.

# Molteplici strumenti: applicazione Choi criteria (valutazione quantitativa del contrast-enhancement in CT e MR)

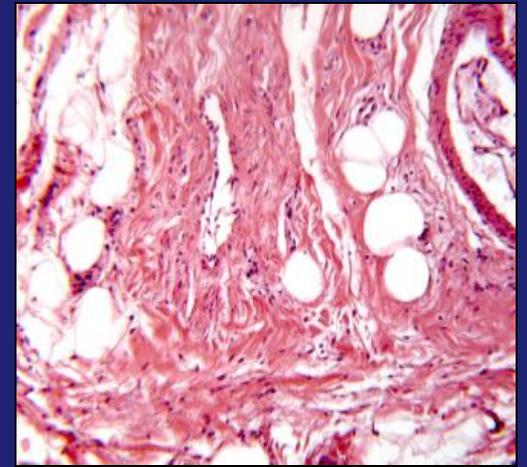
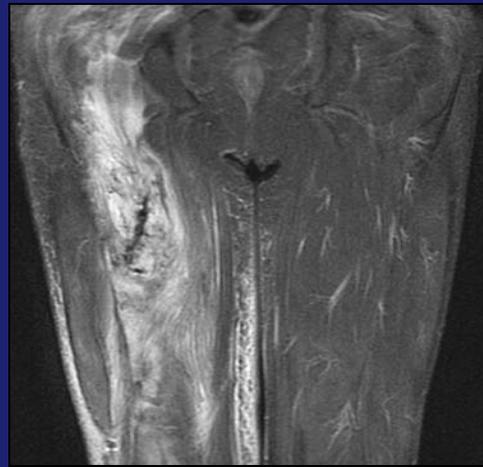
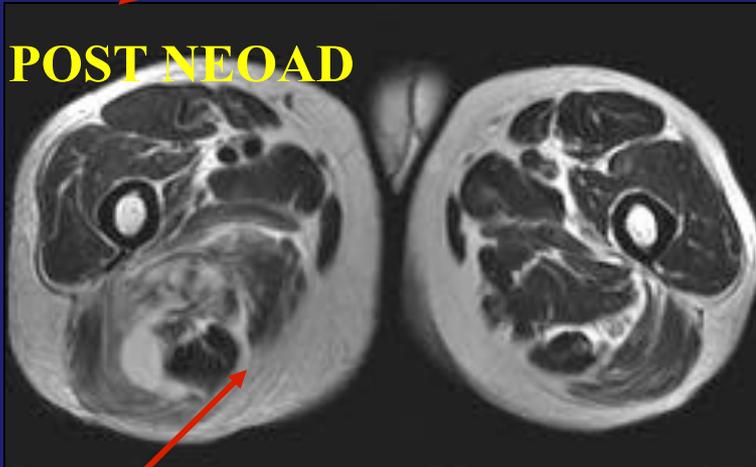
Tumor Response according to RECIST and Choi Criteria 		
Response	RECIST Criteria	Choi Criteria
Complete response	Disappearance of all lesions No new lesions	Disappearance of all lesions No new lesions
Partial response	$\geq 30\%$ decrease in the sum of greatest diameters No new lesions	$\geq 10\%$ decrease in the greatest maximal diameter or a $\geq 15\%$ decrease in tumor attenuation at CT or contrast enhancement at MR imaging No new lesions
Stable disease	Does not meet criteria for complete response, partial response, or progressive disease	Does not meet criteria for complete response, partial response, or progressive disease
Progressive disease	$\geq 20\%$ increase in the sum of greatest diameters	$\geq 10\%$ increase in the greatest maximal diameter and does not meet criteria for partial response by using tumor attenuation at CT or contrast enhancement at MR imaging or $\geq 15\%$ increase in tumor attenuation at CT or contrast enhancement at MR imaging and does not meet the criteria for partial response by using tumor size
	New lesion	New lesion New intratumoral nodule or increase in the size of existing intratumoral nodule

**Stacchiotti S:** High-Grade Soft-Tissue Sarcomas: Tumor Response Assessment-Pilot Study to Assess the Correlation between Radiologic and Pathologic Response by Using RECIST and Choi Criteria. **Radiology 251 (2): 447-456**

**PRE NEOAD**



**POST NEOAD**



High-grade mixofibrosarcoma

# Molteplici strumenti: imaging funzionale

in oncologia msk la maggior esperienza è sicuramente sull'osteosarcoma, ma vi sono alcuni lavori sui sarcomi delle parti molli

## 1. DCE MRI

### Phase I Trial of the Antivascular Agent Combretastatin A4 Phosphate on a 5-Day Schedule to Patients With Cancer: Magnetic Resonance Imaging Evidence for Altered Tumor Blood Flow

By James P. Stevenson, Mark Rosen, Weijing Sun, Maryam Gallagher, Daniel G. Heller, David Vaughn, Bruce Giordano, Ross Zimmer, William P. Petras, Michael Striford, David Chaplin, Scott L. Young, Mitchell Schindl, and Peter J. O'Dwyer

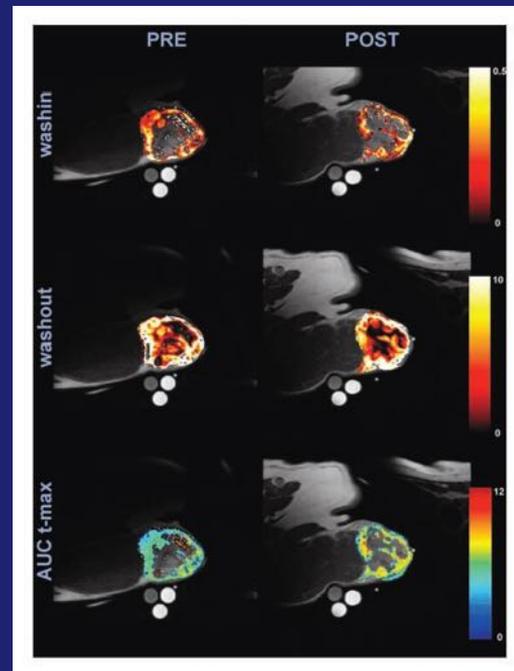
*Journal of Clinical Oncology*, Vol 21, No 23 (December 1), 2003; pp 4428-4438

### Dynamic Contrast-enhanced Magnetic Resonance Imaging as a Predictor of Clinical Outcome in Canine Spontaneous Soft Tissue Sarcomas Treated with Thermoradiotherapy

Benjamin L. Viglianti,<sup>1</sup> Michael Lora-Michiels,<sup>1</sup> Jeanie M. Poulson,<sup>1</sup> Lan Lan,<sup>2</sup> Dahio Yu,<sup>5</sup> Linda Sanders,<sup>2</sup> Oana Craciunescu,<sup>1</sup> Zeljko Vujaskovic,<sup>1</sup> Donald E. Thrall,<sup>4</sup> James MacFall,<sup>3</sup> Cecil H. Charles,<sup>3</sup> Terence Wong,<sup>3</sup> and Mark W. Dewhurst<sup>1</sup>

*Clin Cancer Res* 2009;15(15) August 1, 2009

9/37 pts sarcoma  
Valutazione  $K^{tr}$



Meeting Report

Diffusion-Weighted Magnetic Resonance Imaging as a Cancer Biomarker: Consensus and Recommendations

Anwar R. Pathani<sup>1</sup>, Guoying Liu<sup>2</sup>, Dow Mu-Koh<sup>3</sup>, Thomas L. Chenevert<sup>4</sup>, Harriet C. Thoeny<sup>5</sup>, Taro Takahara<sup>6</sup>, Andrew DeK-Jung<sup>7</sup>, Brian D. Ross<sup>8</sup>, Marc Van Cauteren<sup>9</sup>, David Collins<sup>10</sup>, Dima A. Hammoud<sup>11</sup>, Gordon J.S. Rustin<sup>12</sup>, Bachir Taouli<sup>13</sup> and Peter L. Choyke

## Molteplici strumenti: imaging funzionale

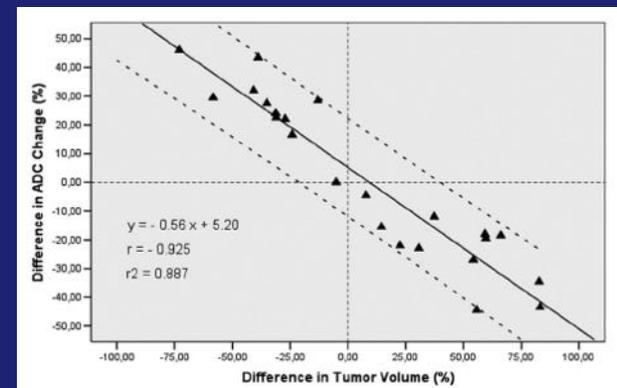
- 2) **DWI**: Diffusion-Weighted Imaging

### Diffusion-Weighted Magnetic Resonance Imaging Allows Monitoring of Anticancer Treatment Effects in Patients With Soft-Tissue Sarcomas

Oliver Dudeck, MD,<sup>1,2\*</sup> Martin Zeile, MD,<sup>3</sup> Daniel Pink, MD,<sup>4</sup> Maciej Pech, MD,<sup>2</sup> Per-Ulf Tunn, MD,<sup>5</sup> Peter Reichardt, MD,<sup>4</sup> Wolf-Dieter Ludwig, MD,<sup>4</sup> and Bernd Hamm, MD<sup>1</sup>

**Conclusion:** DWI can be used as a supplement to morphologic imaging for the evaluation of tumor response to anticancer therapy in patients with soft-tissue sarcomas. As cellular changes are expected to precede morphologic changes in treated tumors, DWI performed at an early stage of fractionated therapy may provide unique prognostic information of its effectiveness.

JOURNAL OF MAGNETIC RESONANCE IMAGING 27:1109-1113 (2008)



**Apparent Diffusion Coefficient:** rappresenta lo spazio percorso da una molecola di acqua nel suo moto stocastico nell'unità di tempo (mm<sup>2</sup>/sec) in funzione del mezzo biologico non omogeneo e spesso anisotropo in cui le molecole di acqua diffondono

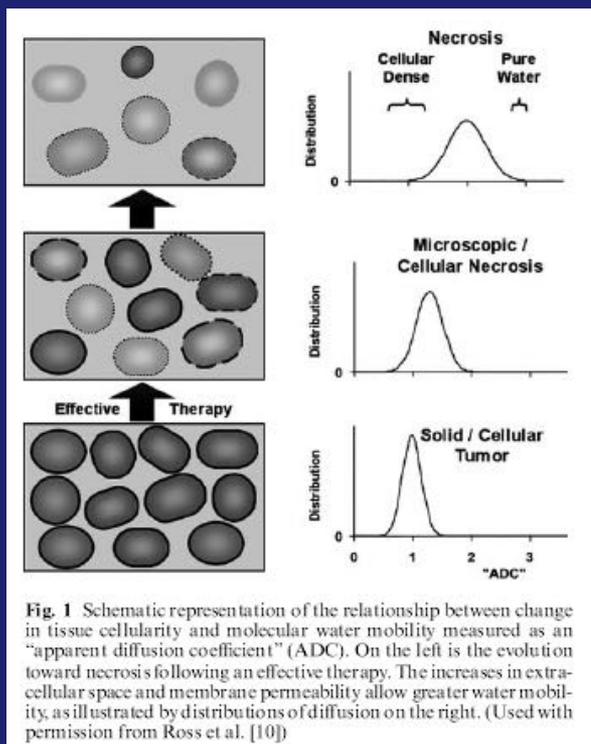
$$ADC = (lnS_0 - lnS) / b$$

Terapia → Necrosi Tumorale  
 ↑ ADC

↑ Permeabilità parete cellulare

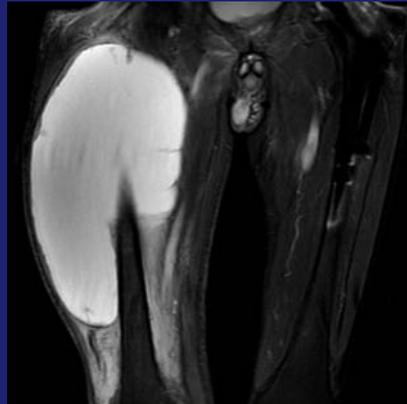
↓ Cellularità

↑ Spazi intercellulari  
 Acqua spazi intercellulari

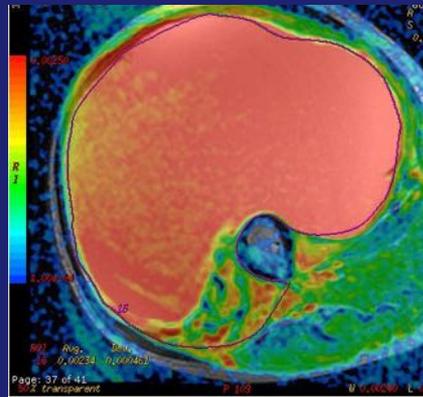
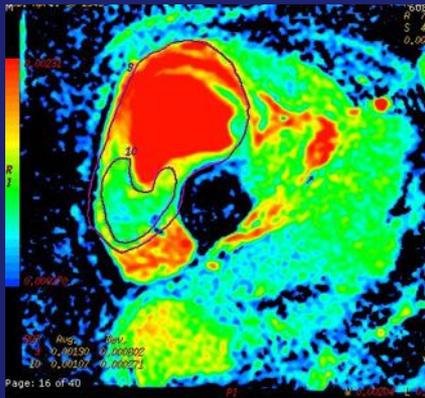


B. A. Moffat  
 D. E. Hall  
 J. Stojanovska  
 P. J. McConville  
 J. B. Moody  
 T. L. Chenevert  
 A. Rehemtulla  
 B. D. Ross

**Diffusion imaging for evaluation of tumor therapies in preclinical animal models**



**Leiomyosarcoma G3:**  
estesa necrosi  
colliquativa post-CTC  
(intervallo tra RM 161 gg)  
(da 175 mm a 295 mm diam. max)

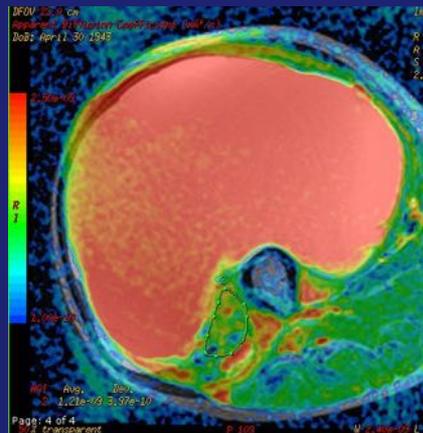


ADC medio "globale":

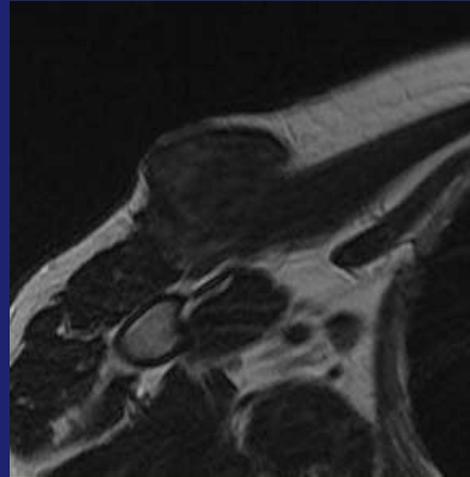
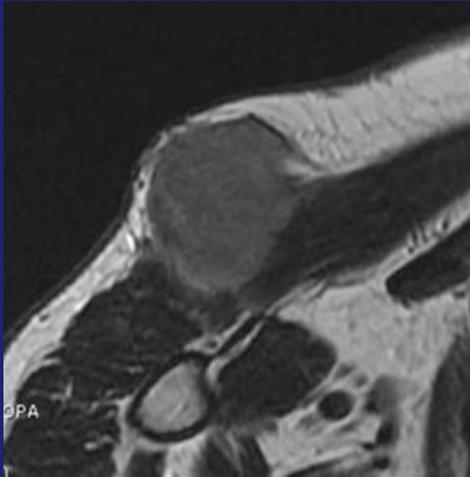
↪ 2,00E-03 mm/sec<sup>2</sup>  
↪ 2,36E-03 mm/sec<sup>2</sup>

ADC medio "a campione":

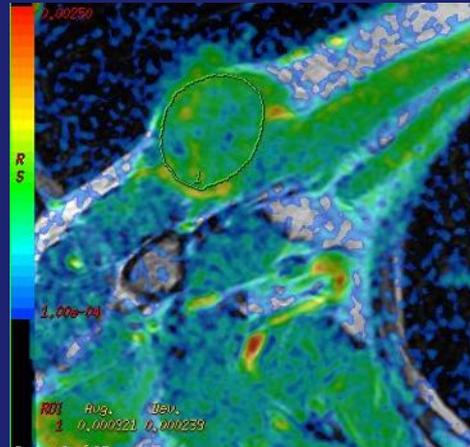
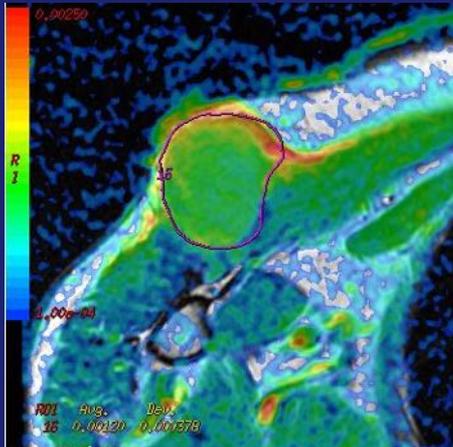
↪ 1,04E-03 mm/sec<sup>2</sup>  
↪ 1,21E-03 mm/sec<sup>2</sup>



Come interpretare l'aumento dimensionale?  
Causato dalla necrosi tumorale  
indice di buona risposta?



CASI DUBBI  
Sarcoma dedifferenziato (IFM)  
Diminuzione dimensionale  
Involuzione fibrotica,  
Scarsa necrosi colliquativa  
(intervallo 85 gg)  
(da 57 a 53)

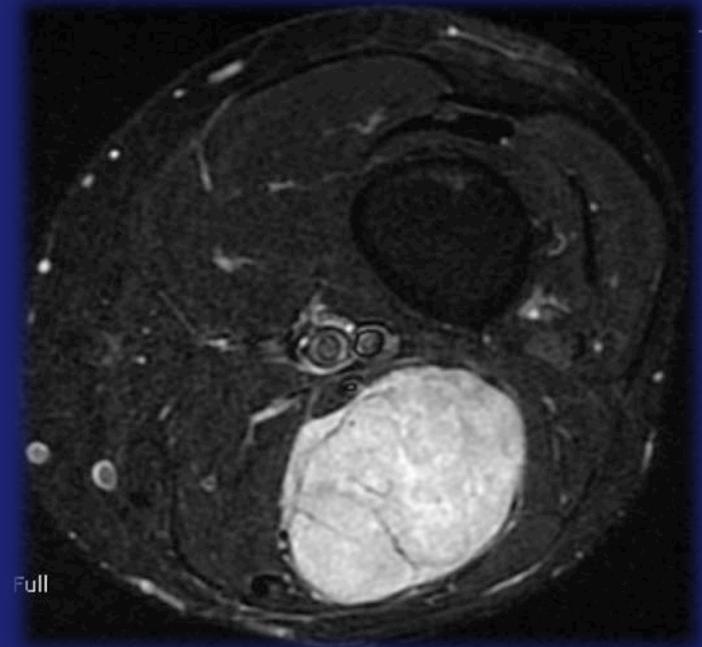
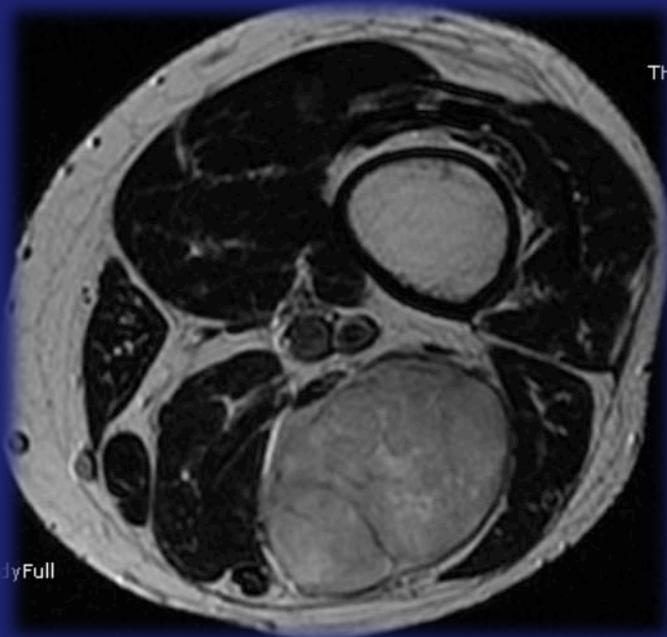


↓ ADC medio "globale"  
1,00E-03 → 9,21E-04

↓ ADC medio "a campione"  
1,41E-03 → 1,00E-03

- **Paziente M 38 anni,**
- **Tumefazione coscia sn**
- **Asintomatico**

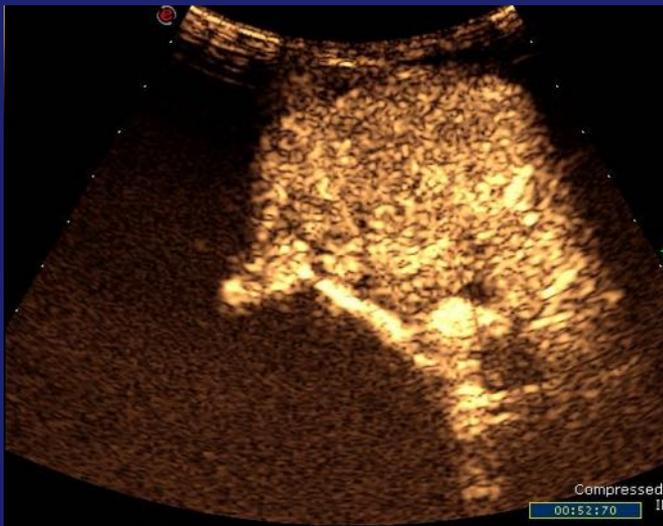
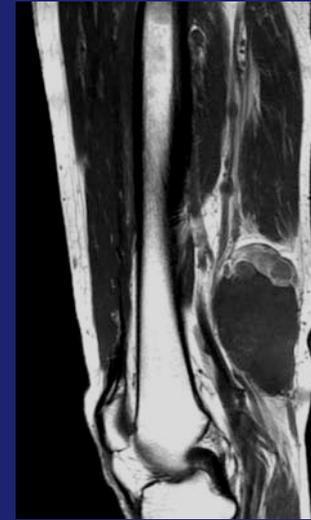
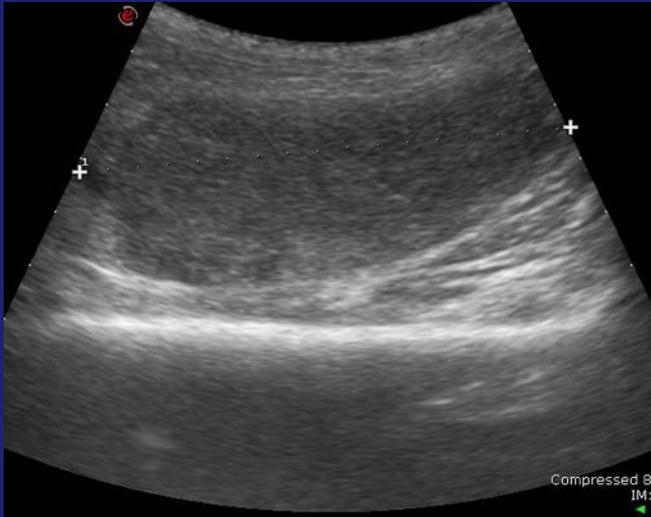
## **RM basale**



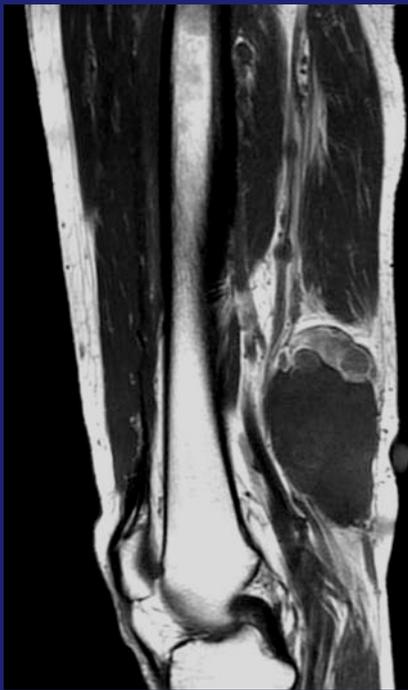
- **ax FR FSE T2:**  
**Iperintensa con**  
**aspetto “zolle”**

- **ax STIR**  
**Nettamente iperintensa**

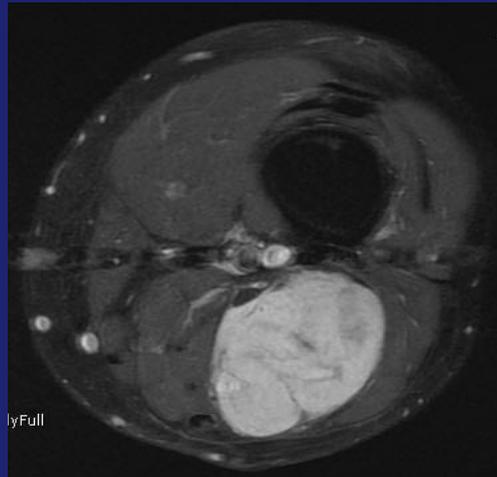
**In corrispondenza della  
biforcazione tra nervo tibiale e  
nervo peroneo comune**



CEUS: dopo mdc ev a 24 sec  
si evidenziano vasi che dalla  
periferia si portano verso il centro  
con distribuzione a carattere  
reticolare



- sag FR FSE T1 senza e con mdc:  
Netto contrast-enhancement
- ax FSE fat sat T1 senza e con mdc  
Netto contrast-enhancement

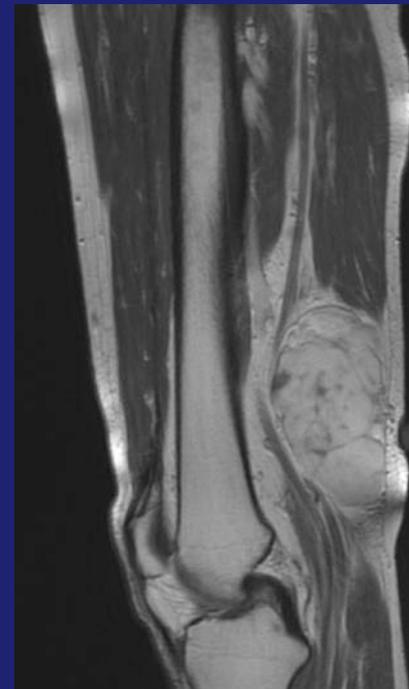
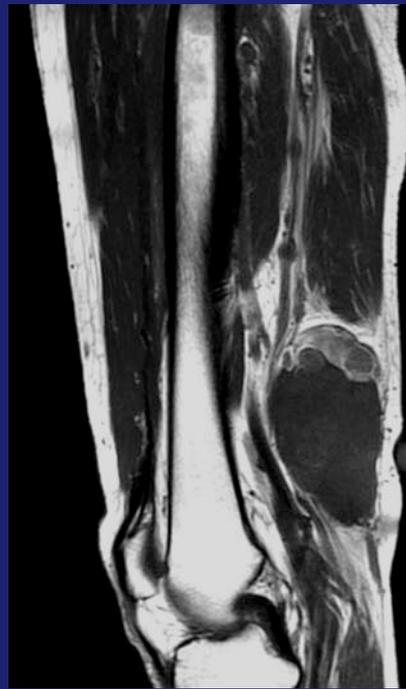


- E' possibile ipotizzare la linea cui appartiene la massa?

- a. sì, è un tumore a matrice mixoide

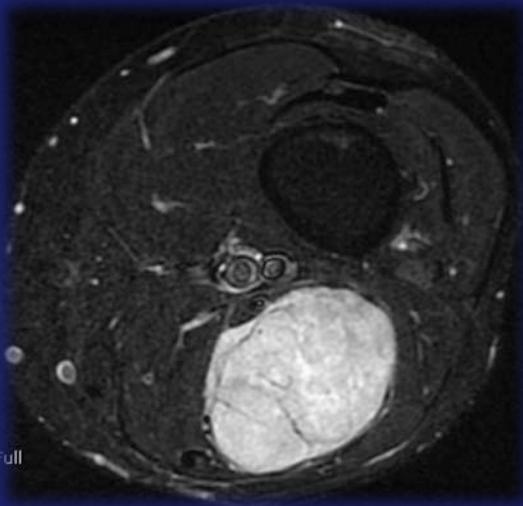
- b. sì, è tumore delle guaine nervose

- c. no



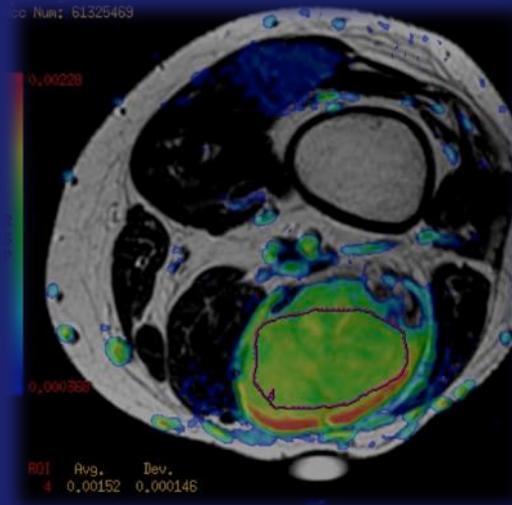
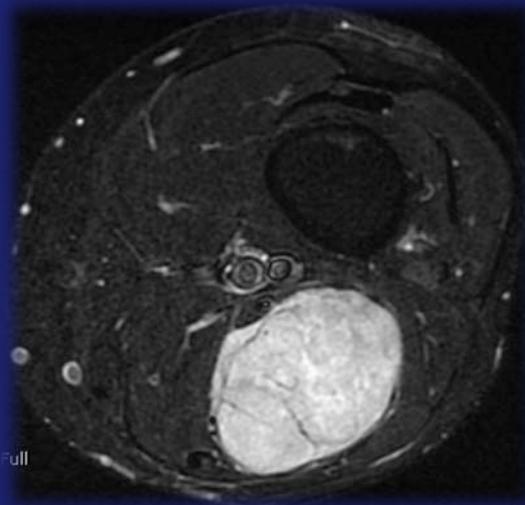
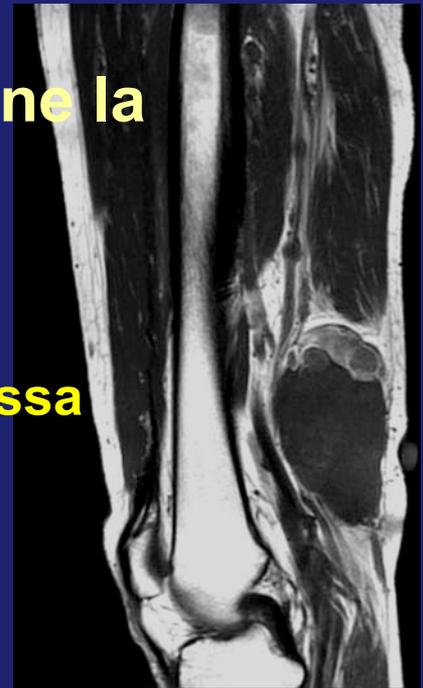
- **E' possibile ipotizzare la linea cui appartiene la massa?**

- a. sì, può essere un tumore a matrice mixoide, il nervo tibiale non fuoriesce dai poli della massa**



- E' possibile ipotizzare la linea cui appartiene la massa?

- sì, può essere un tumore a matrice mixoide, il nervo tibiale non fuoriesce dai poli della massa



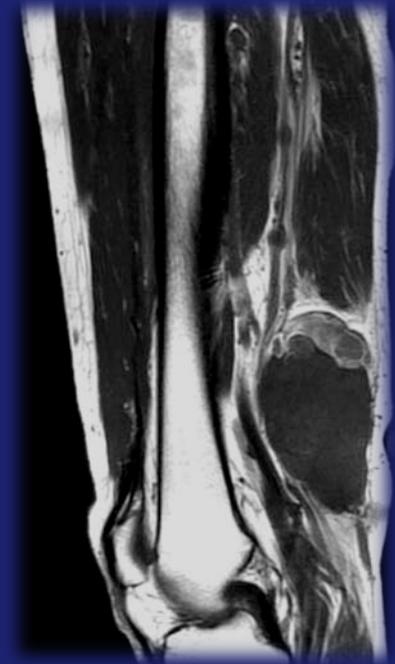
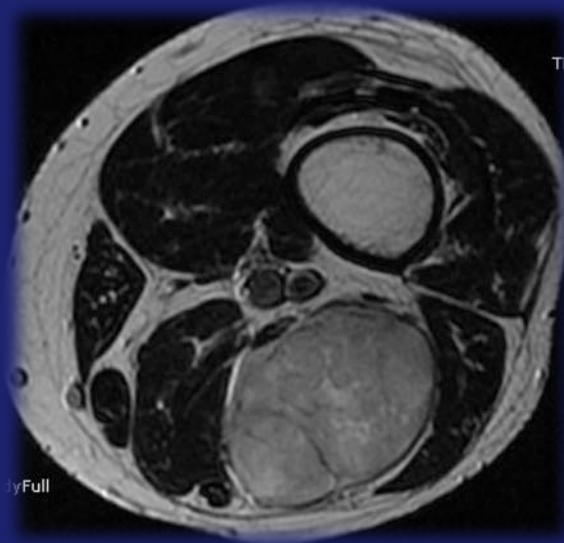
- DWI b 900 mm<sup>2</sup>/sec  
lesione disomogenea  
ADC intermedio  
(1,52 sec/mm<sup>2</sup>)

# Esame istologico core-needle

## Liposarcoma mixoide G3.

G3. maggior cellularità, matrice mixoide meno estesa: valori di ADC inferiori

- Intracompartmentale?
- Extracompartmentale?

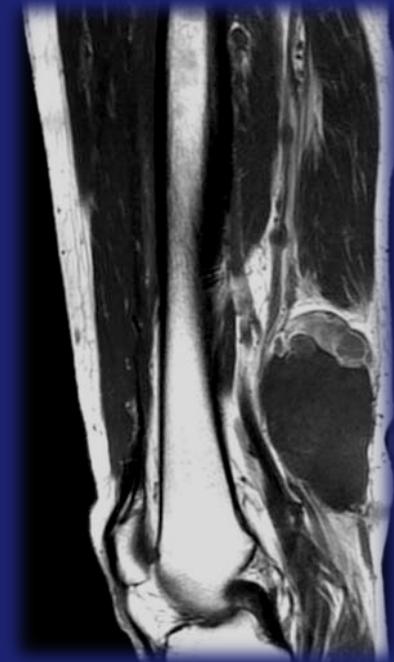
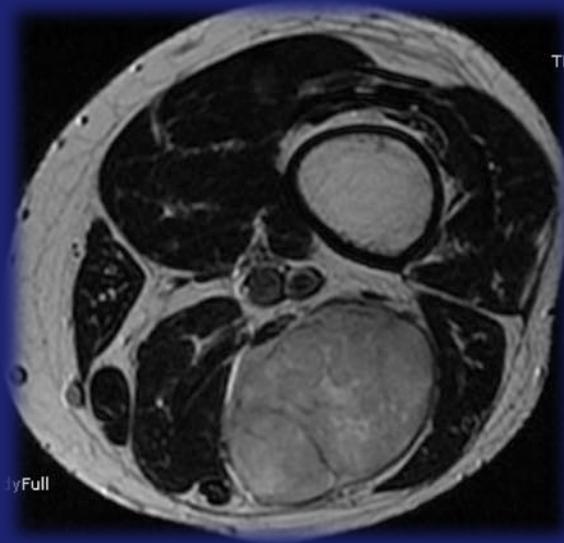


# Esame istologico core-needle

## Liposarcoma mixoide G3.

**G3. maggior cellularità, matrice mixoide meno estesa: valori di ADC inferiori**

- **Cavo popliteo, intimo contatto con i rami nervosi: Extracompartmentale**



- CTC neo adiuvante: netta riduzione dimensionale



Resezione con  
epinevrectomia

**“THE DIAGNOSTIC OF SOFT TISSUE  
SARCOMAS IMPLIES PRIMARILY A  
MULTISCIPLINARY APPROACH FOR A  
STAGE ASSOCIATED THERAPY”.....**

**Tunn PU, Gebauer B., Fritzmann J., e al.**

**Chir.2004**

## **“TAKE-HOME” MESSAGE**

- X-ray is always necessary
- US is often the first step, but with color-power Doppler ( c.m.)
- MR is the gold standard ( c.m.,dynamic,diffusion.....)
- CT with c.m. for the presurgical planning

**THANK YOU**

