

Unità Operativa di Neurologia
Ospedale "Madonna del Soccorso"
San Benedetto del Tronto (AP)
Direttore M.Ragno

**NEUROSONOLOGIA ED
ECOGRAFIA CEREBRALE:
Brain Parenchyma Sonography
e malattie extrapiramidali**

**Perché il TCCD è
diventato un esame
insostituibile nella
gestione del paz con
patologia cerebrale?**

Dr. Sandro Sanguigni

***Neurosonologia
Stroke-Unit***

**San Benedetto del Tronto
6-8 Novembre 2017**



**Società Italiana
Interdisciplinare Neurovascolare**





STUDI ARTERIOSI

STUDI VENOSI

STUDI PARENCHIMALI



NEUROSONOLOGIA

STUDIO ECOGRAFICO PARENCHIMALE

Indicazioni cliniche

BRAIN PARENCHYMA SONOGRAPHY

Anatomia e metodologia

IN TEMP DX

2.0MHz

TCI

PWR

11

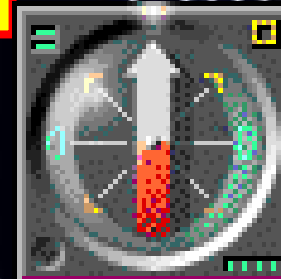
0.70

11

11

Why ultrasound parenchymal study ?

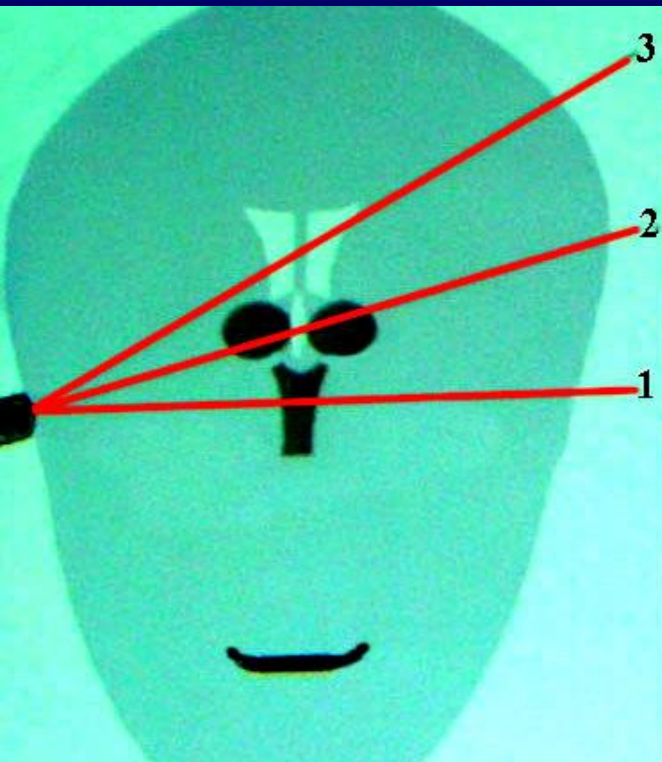
➤ Mastery of anatomy: Where are we? What are we seeing ??



DOVE SIAMO?? COSA STIAMO INSONANDO??

axial Plane:

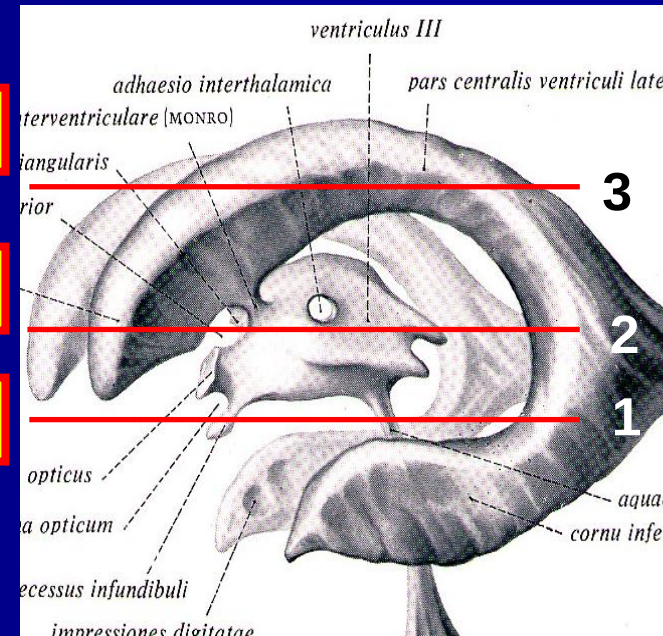
(orbito-meatal line)



Sovradiencephalic (3)

Diencephalic plane(2)

Mesencephalic (1)

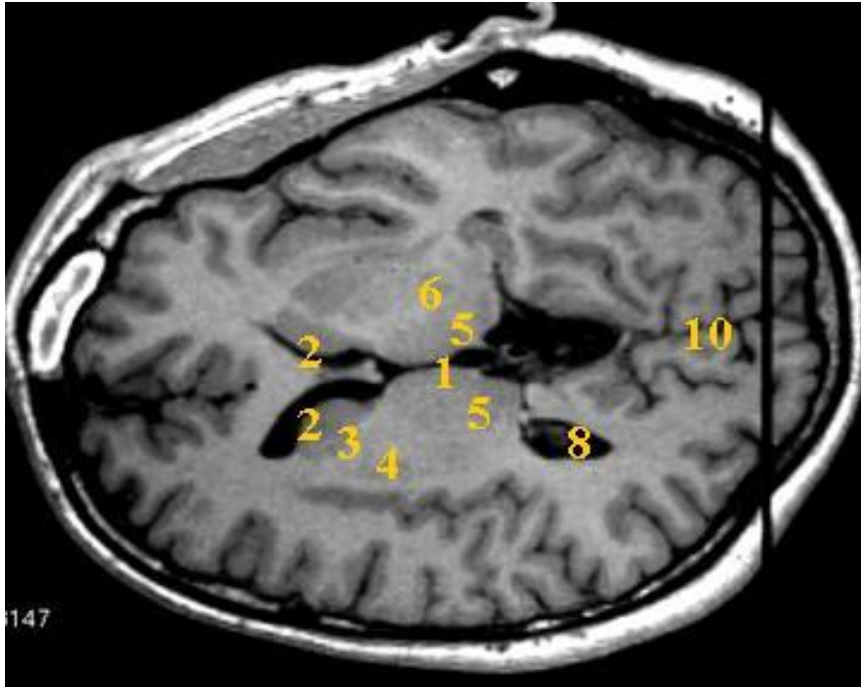


IL CONCETTO DI «FUSION» STA CAMBIANDO TALI PIANI !!

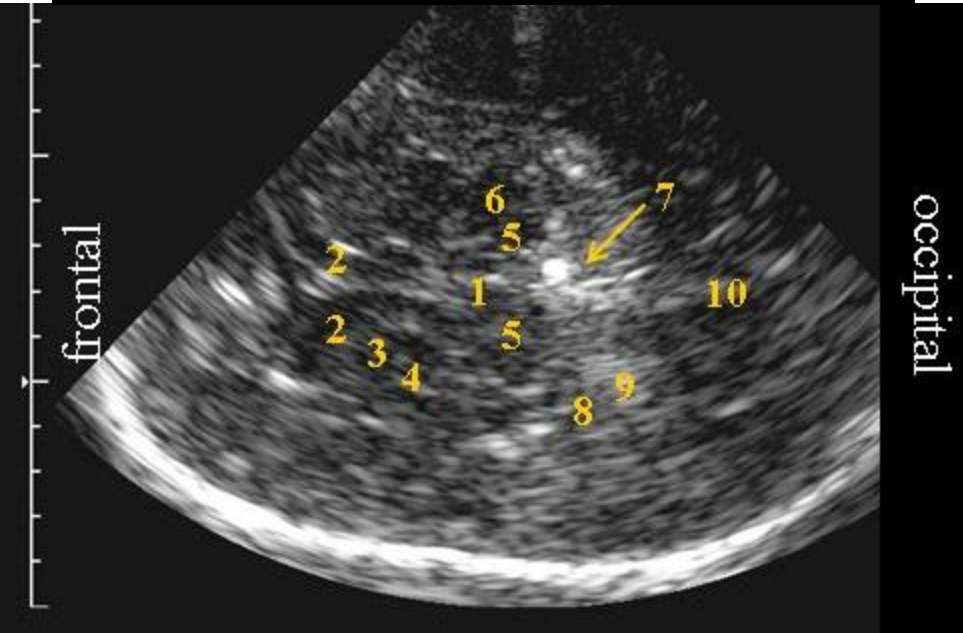
Il piano lo stabilisco io ma lo valido con TAC/RMN !

Non ho più bisogno di punti di repere ecografici !

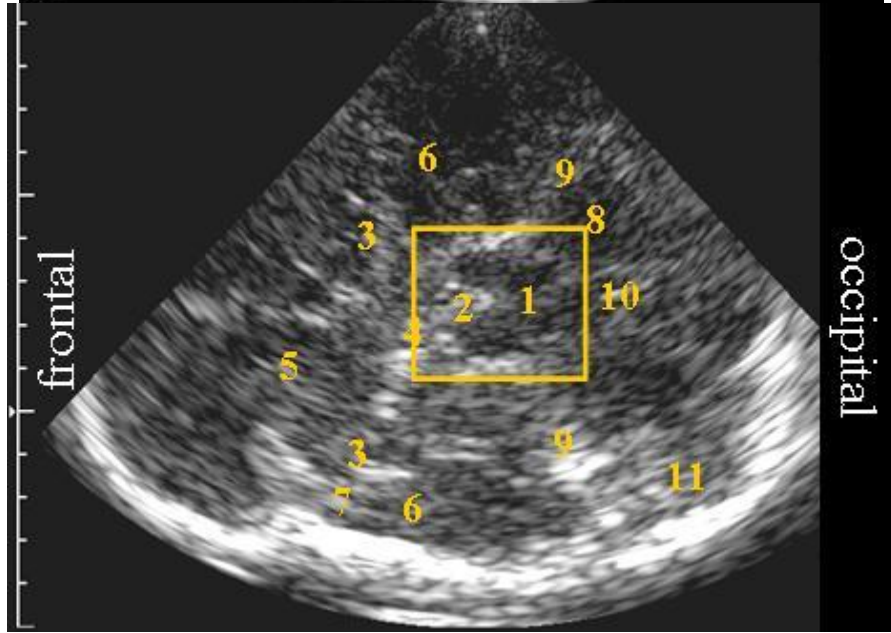
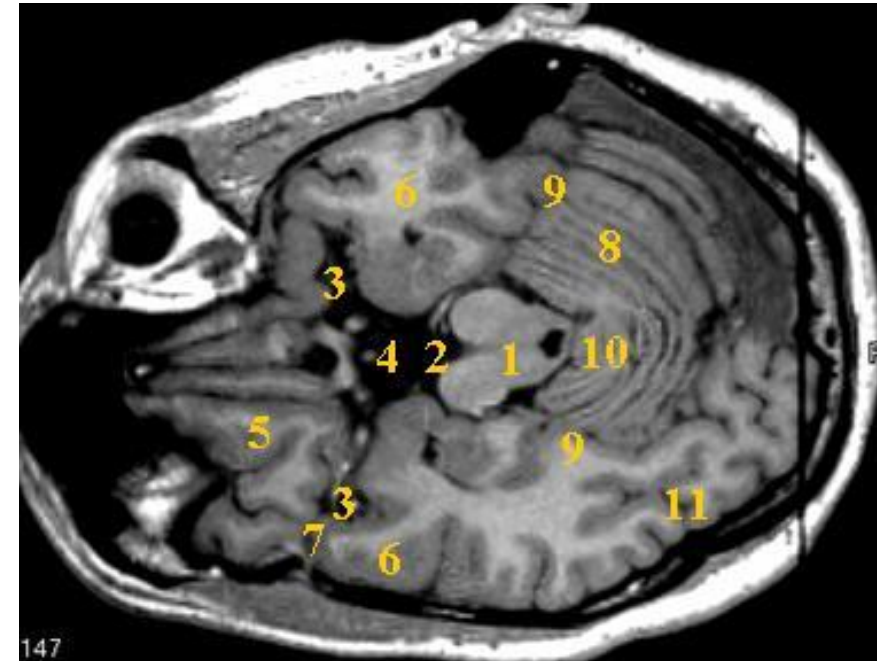
Anatomia e imaging ultrasonoro



1. 3.ventricle
2. anterior horn
3. nucleus caudatus
4. nucleus lentiformis
5. thalamus
6. capsula interna
7. glandula pinealis
8. posterior horn
9. plexus choroideus
10. falx cerebri

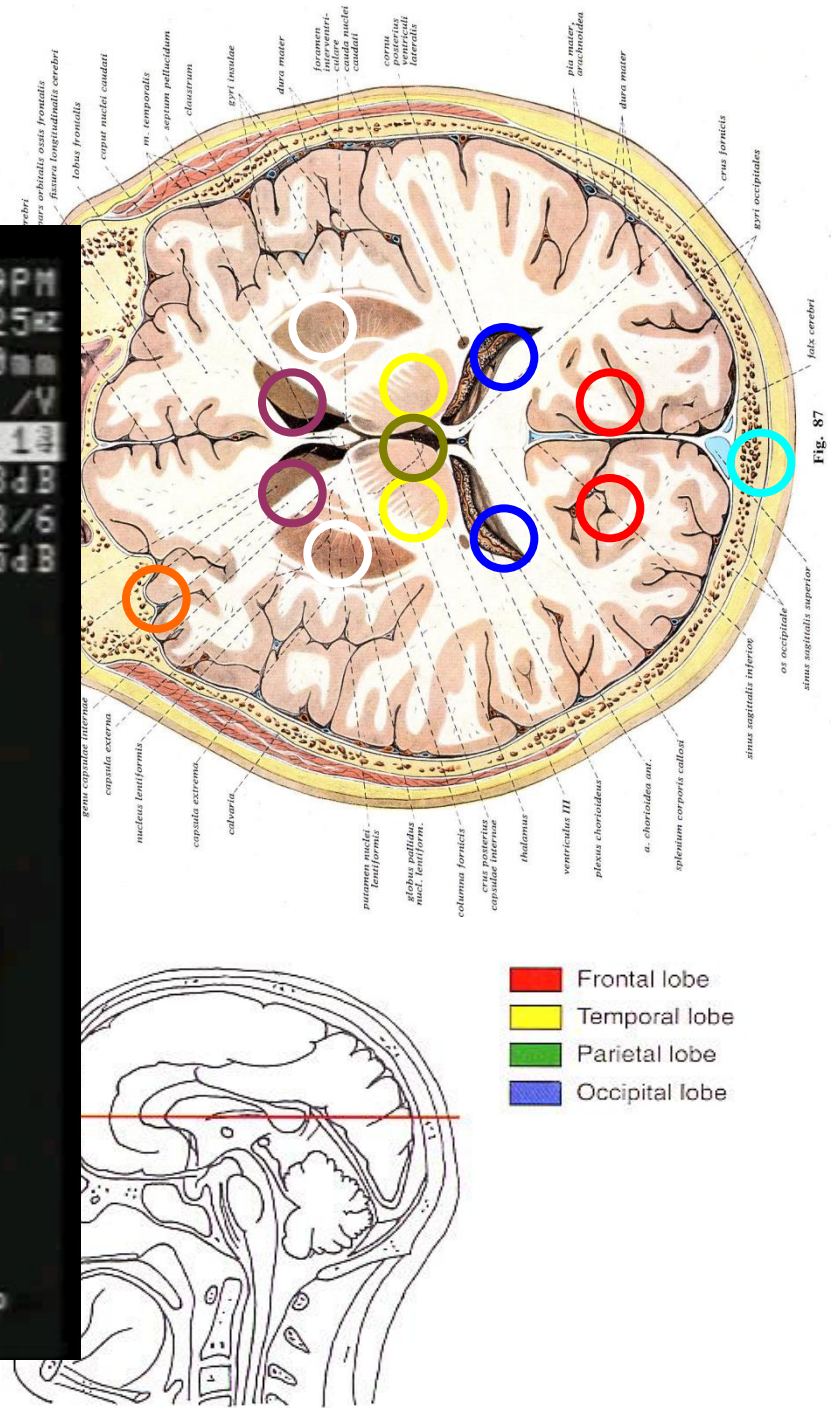
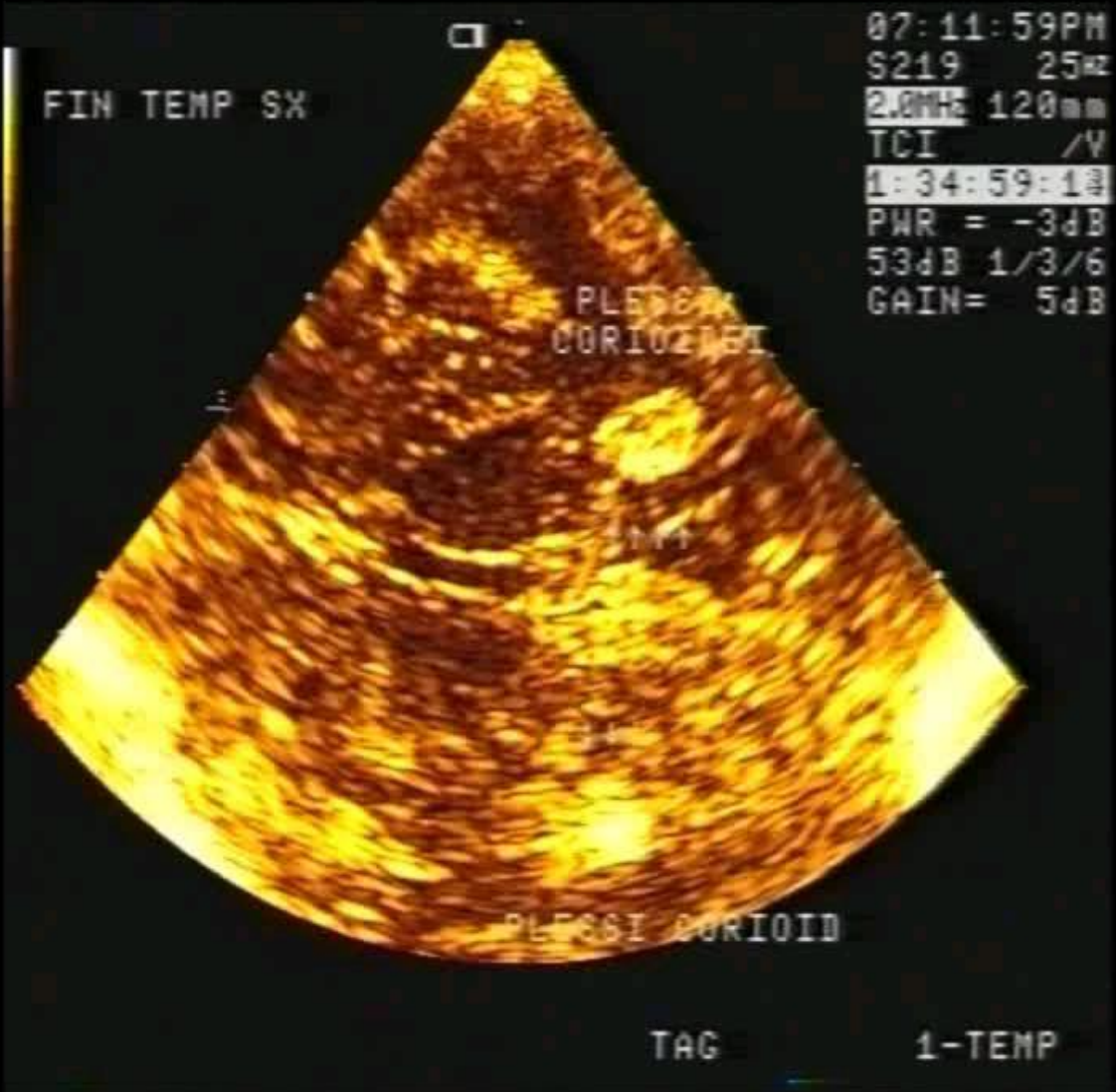


Anatomia e imaging ultrasonoro



1. mesencephalic brainstem
2. basal cistern
3. mca
4. chiasma opticum
5. basal frontal lobe
6. temporal lobe
7. sylvic fissure
8. cerebellum
9. tentorium cerebelli
10. superior cerebellar vermis
11. occipital lobe

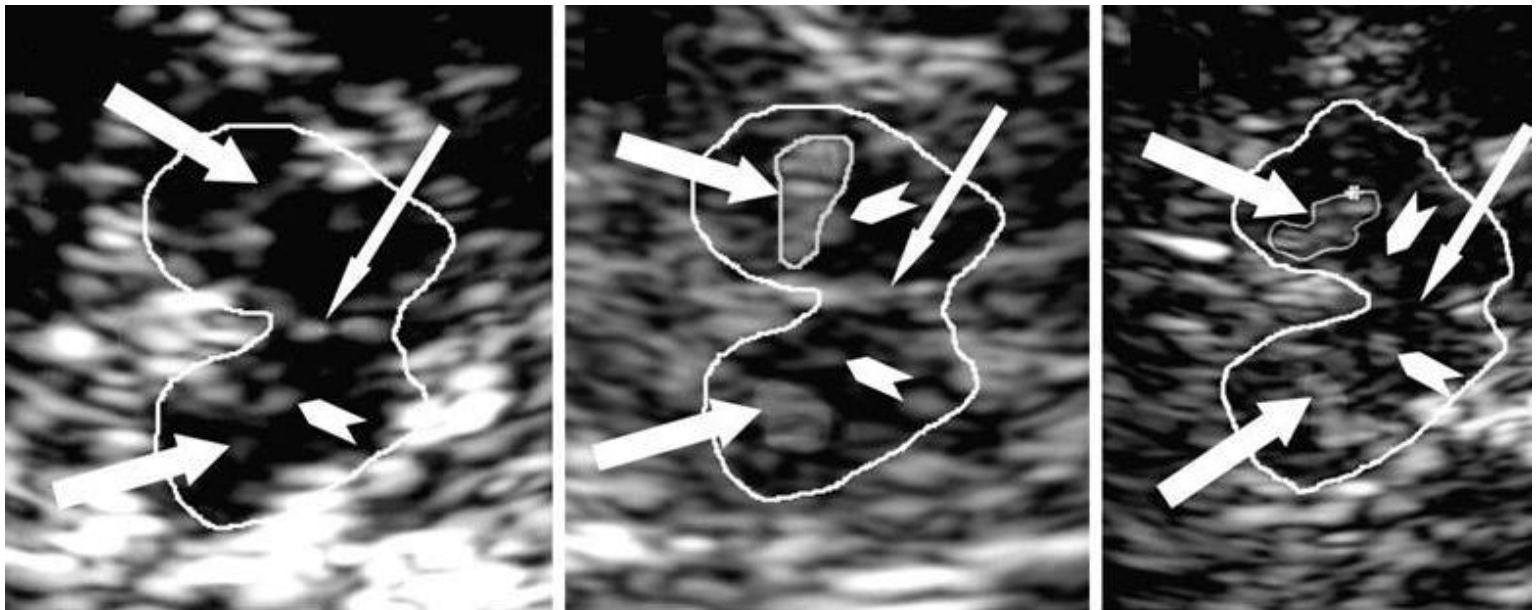
BRAIN PARENCHYMA



Alterata ecogenicità del rafe mediano mesencefalico

Presente nel:

- 10 % della popolazione normale,
- 50 – 70 % dei paz con disturbi depressivi.



Role of Transcranial Ultrasound in the Diagnosis of Movement Disorders

Neuroimaging Clin N Am. (2010)

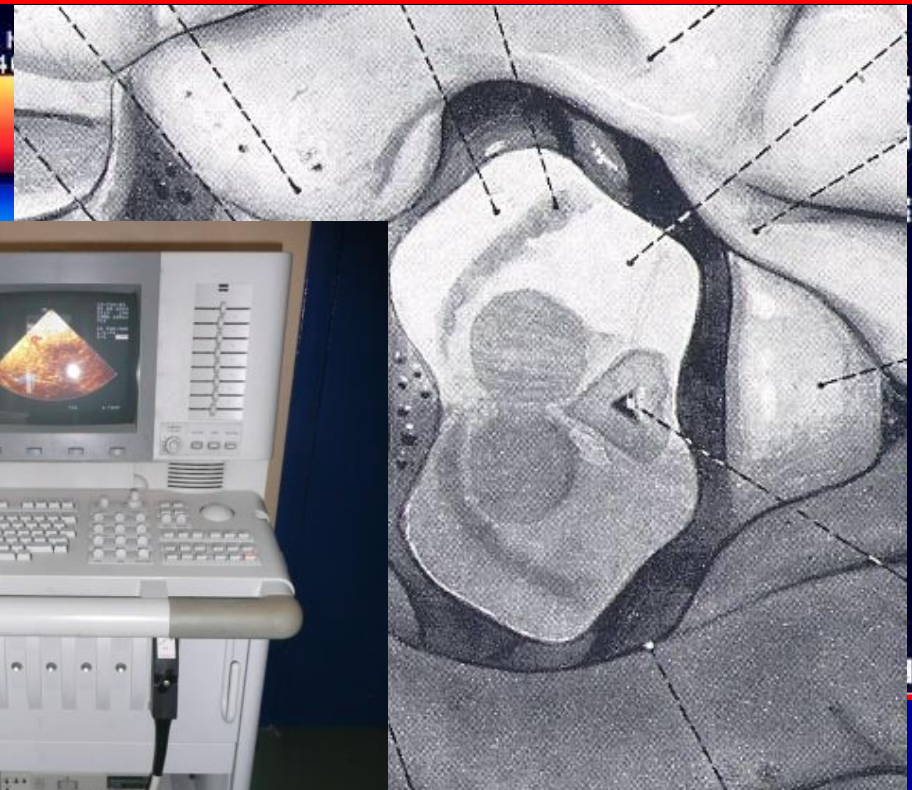
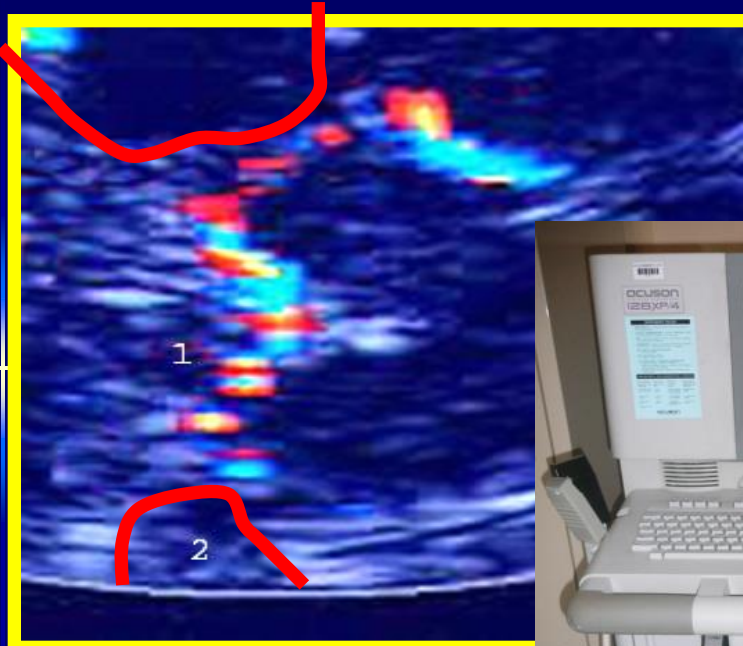
Jana Godau, MD*, Daniela Berg, MD

Table 2
System settings

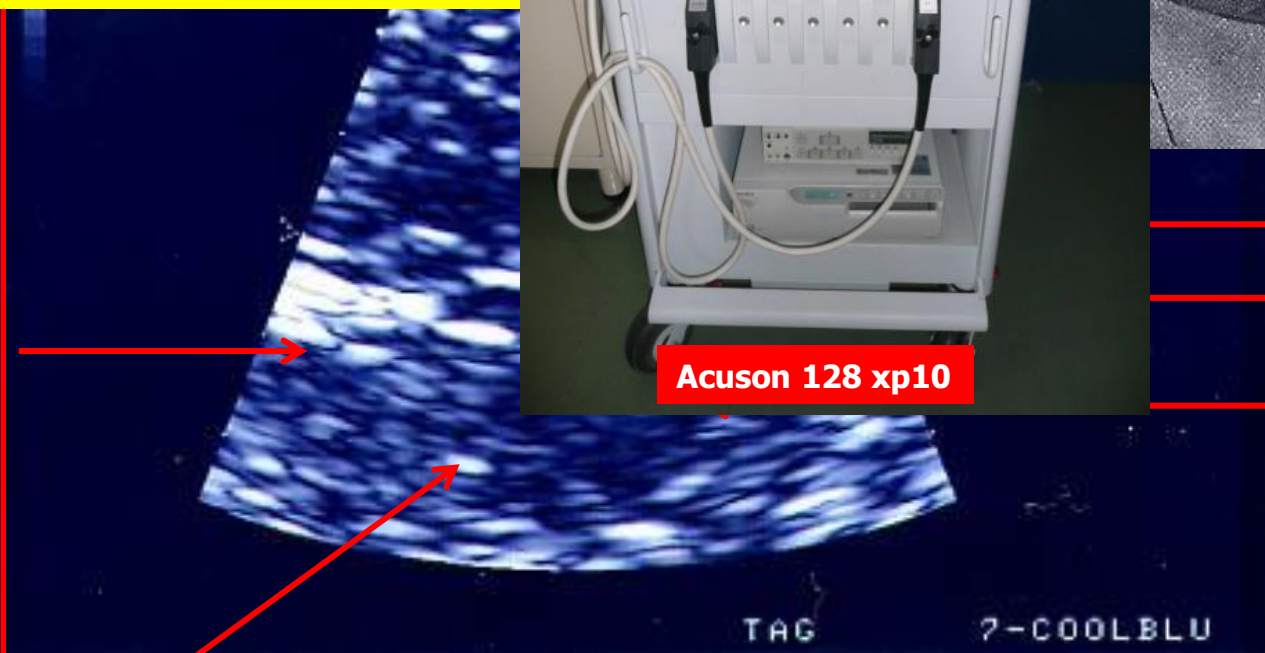
Ultrasound transducer	1.8–3.5 MHz
Penetration depth	14–16 cm
Dynamic range	45–55 dB
Contour amplification	Medium–high
Postprocessing parameters	Moderate suppression of low echo signals
Time gain compensation	Adjust as needed
Image brightness	Adjust as needed

Axial resolution : 0.3-0.7 mm

Lateral resolution :1.1 mm



Caudate



Lenticular Nucleus

Thalamus omolat
3° Ventr

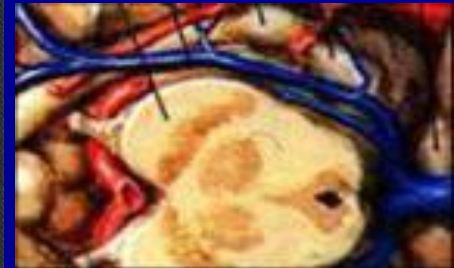
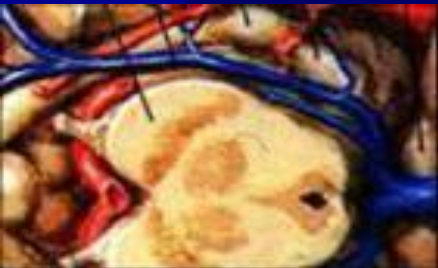
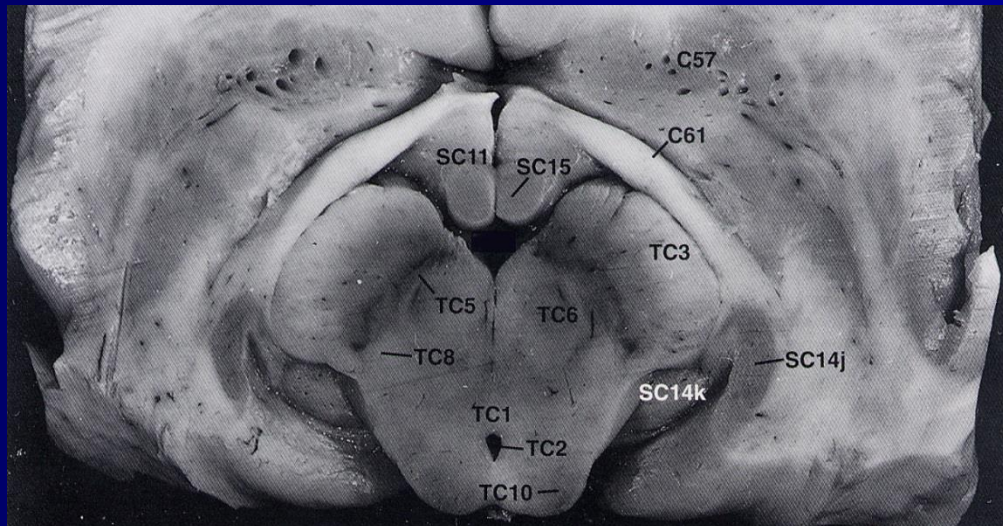
Thalamus control

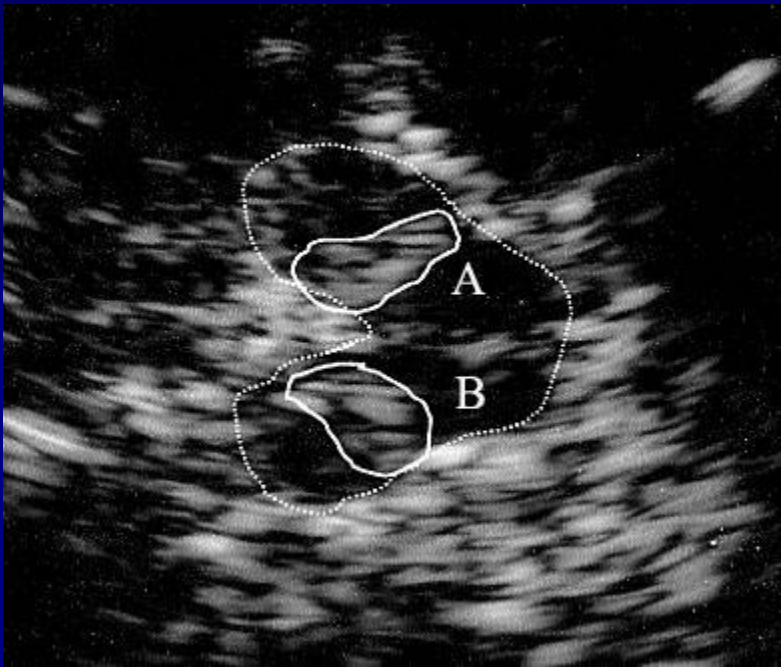
TAG

2-COOLBLU

TCCD e Morbo di Parkinson

**STUDI :
ECOGRAFICI
FLUSSIMETRICI
MED. NUCLEARE**





A=0,25 cm²

B=0,27 cm²

**Changes in cerebral blood flow
detected in the middle cerebral artery,
in patients affected by Idiopathic Parkinson Disease,
during ON-OFF phenomena.**

A transcranial color doppler study.

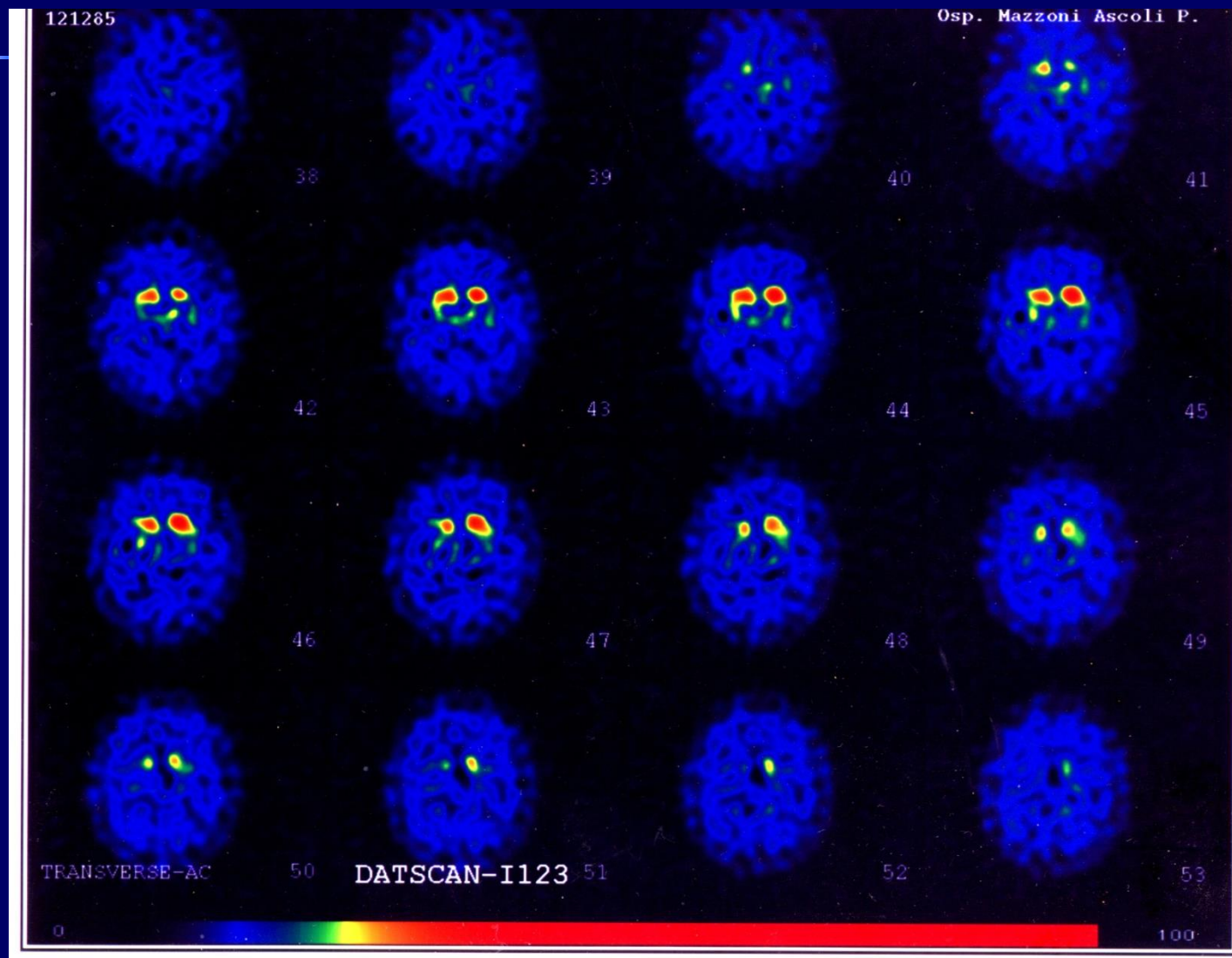
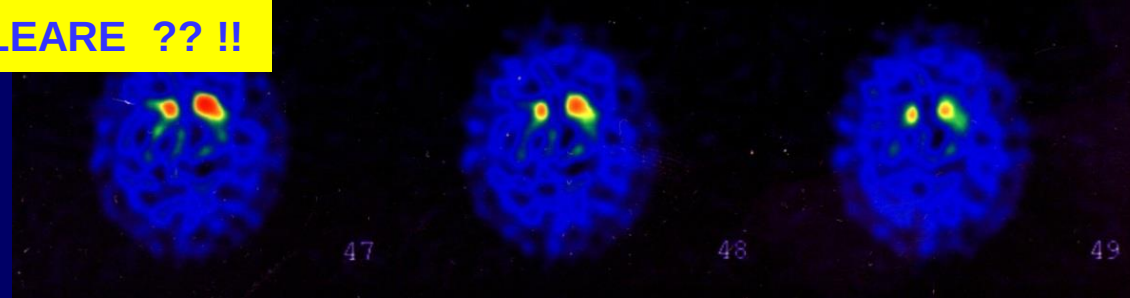
Curatola.Sanguigni.Malferrari,Carboni,Gobbato,Paci.
Journal of Neurology 2002;249-251

Esame eseguito:

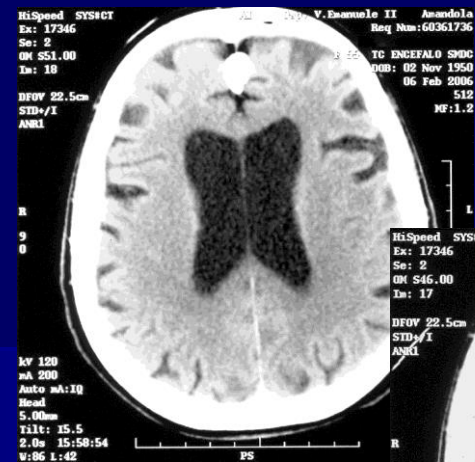
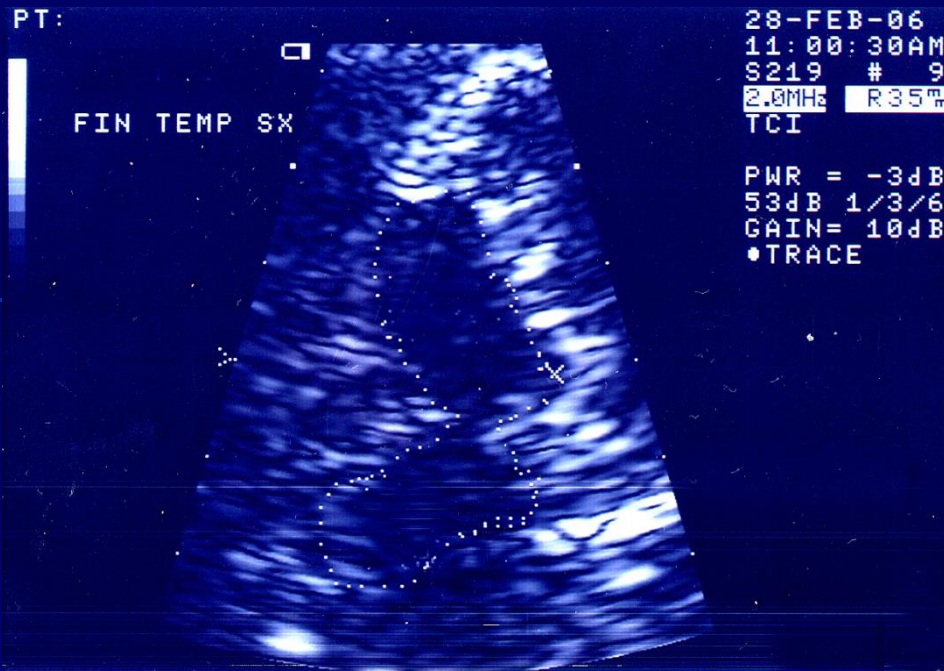
- **Precedentemente esclusi disturbi autonomici (Shy-Drager,Ipot.Posturale ecc.)**
- **Riposo psicosensoriale-occhi chiusi**
- **Stanza buia**
- **Controllo dei valori di P.Arteriosa**
- **Controllo dei valori di saturazione della pO_2**

Pazienti	Arteria Cerebrale Media Destra				Arteria Cerebrale Media Sinistra			
	PSV m/sec		PEDV m/sec		PSV m/sec		PEDV m/sec	
	ON	OFF	ON	OFF	ON	OFF	ON	OFF
Pz 1	1.23	0.87	0.60	0.46	1.46	0.96	0.69	0.49
Pz 2	0.87	0.70	0.43	0.34	0.94	0.67	0.47	0.31
Pz 3	1.33	0.96	0.64	0.42	1.32	0.96	0.67	0.42
Pz 4	1.30	1.03	0.47	0.33	1.30	1.03	0.50	0.33
Pz 5	1.10	0.75	0.37	0.24	1.10	0.79	0.38	0.26
Pz 6	1.01	0.80	0.48	0.35	1.08	0.90	0.50	0.36
Pz 7	1.07	0.78	0.55	0.40	1.15	0.75	0.53	0.38
Pz 8	1.05	0.75	0.48	0.35	1.03	0.68	0.47	0.31
Pz 9	1.15	0.83	0.38	0.25	1.20	0.87	0.41	0.29
Pz 10	1.06	0.80	0.40	0.28	1.01	0.77	0.37	0.26
Pz 11	1.17	0.90	0.39	0.25	1.25	0.96	0.40	0.25
Pz 12	1.13	0.79	0.50	0.29	1.10	0.80	0.52	0.30
Media	1.12± 0.13	0.83± 0.09	0.47± 0.09	0.33± 0.07	1.16± 0.15	0.84± 0.12	0.49± 0.10	0.33± 0.07
P	P<0.001		P<0.001		P<0.001		P<0.001	

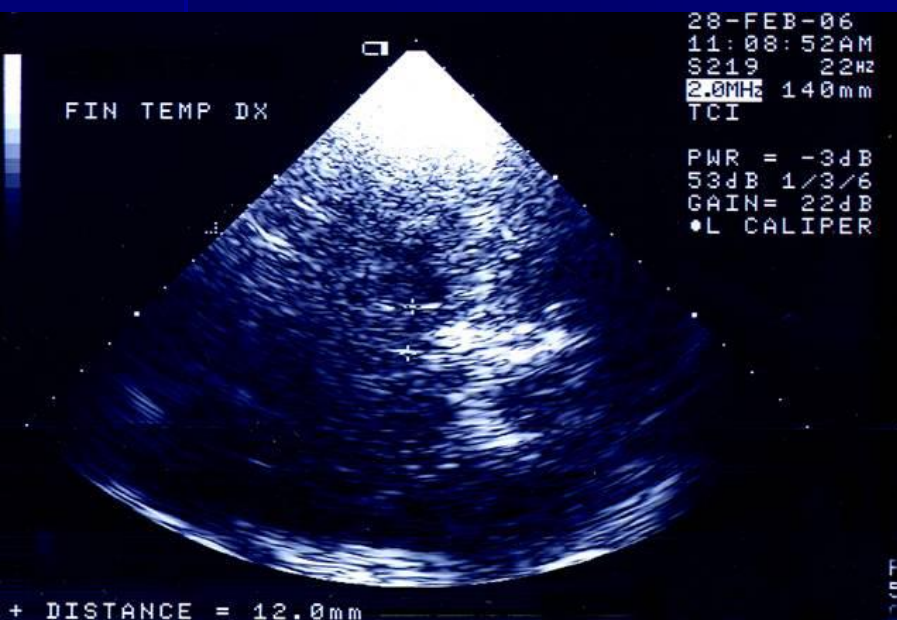
STUDI DI MED. NUCLEARE ?? !!



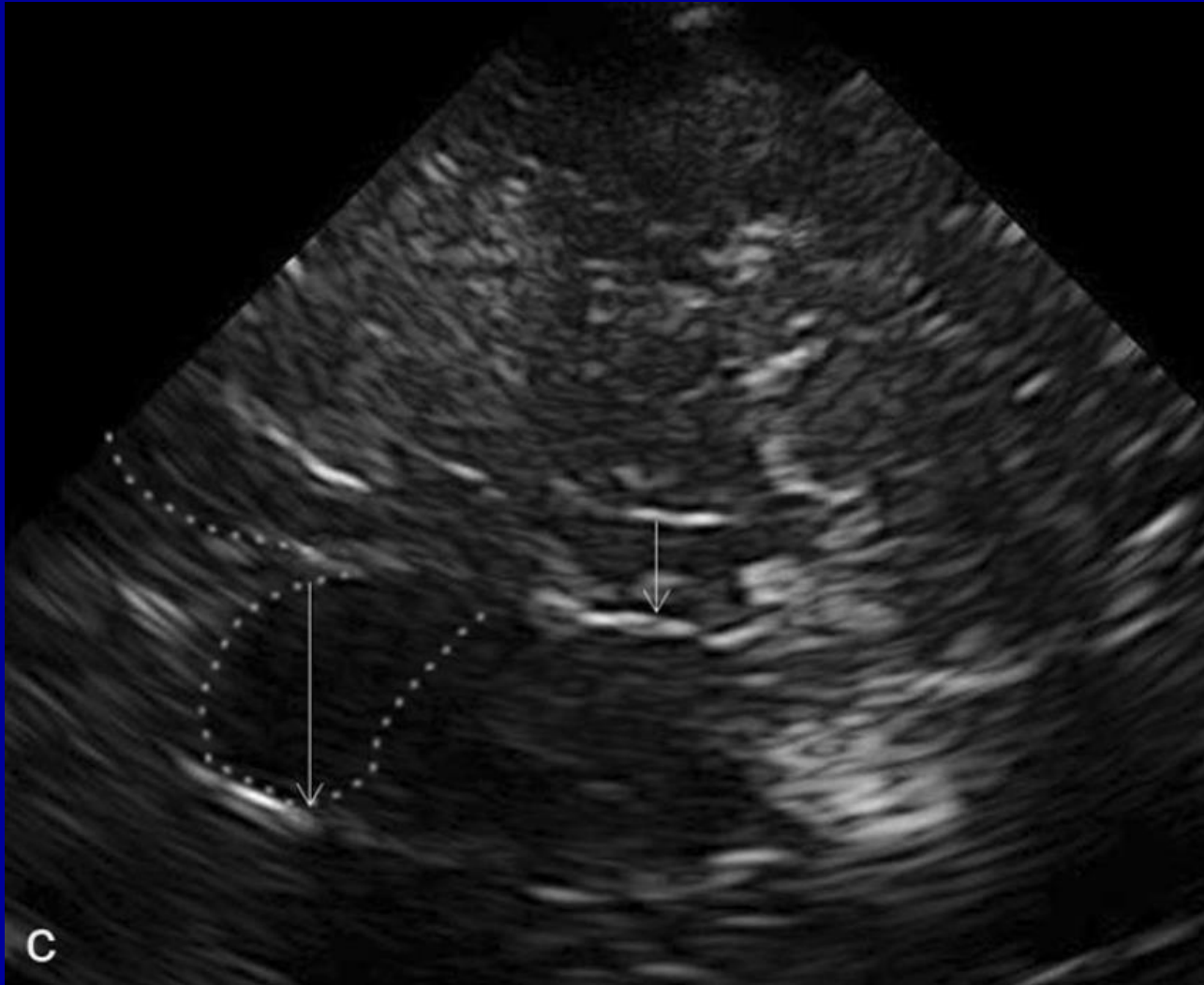
STUDI ECOGRAFICI 2006



PSP



Atypical parkinsonism



PSP

EFNS/MDS-ES recommendations for the diagnosis of Parkinson's disease

A. Berardelli, G. K. Wenning, A. Antonini, D. Berg, B. R. Bloem, V. Bonifati, D. Brooks, D. J. Burn, C. Colosimo, A. Fanciulli, J. Ferreira, T. Gasser, F. Grandas, P. Kanovsky, V. Kostic, J. Kulisevsky, W. Oertel, W. Poewe, J.-P. Reese, M. Relja, E. Ruzicka, A. Schrag, K. Seppi, P. Tabat, M. Vidailhet

European Journal of Neurology 2012 EFNS

The current evidence suggests that TCS is useful in the diagnosis of parkinsonian syndromes, especially with regard to:

- **Differentiation of atypical parkinsonian syndromes** (APS; class I evidence, Level A)
- **Differentiation of secondary parkinsonian syndromes** (class I and II evidence, Level A – for sensitivity, specificity and predictive value of parameters)
- **Early diagnosis of PD, in clinically unclear cases** (class II evidence)
- **Detection of subjects at risk for PD** (class I) including asymptomatic mutation carriers for monogenic forms of PD.

Enlarged substantia nigra hyperechogenicity and risk for Parkinson disease: a 37-month 3-center study of 1847 older persons.

Berg D., Seppi K., Behnke S., Liepelt I., et al

Arch Neurol. 2011 Jul;68(7):932-7.

OBJECTIVE:

To evaluate whether enlarged substantia nigra hyperechogenicity (SN+) is associated with an increased risk for Parkinson disease (PD) in a healthy elderly population.

CONCLUSIONS:

In this prospective study, we demonstrate for the first time a highly increased risk (17.4-fold) for PD in elderly individuals with SN+.

Enlarged hyperechogenic substantia nigra as a risk marker for Parkinson's disease.

Berg D., Behnke S., Seppi K., Godau J., et al

Mov Disord.2013 Feb;28(2):216-9

Recently, we reported a **17.4**-fold increased risk for PD in individuals with SN+ older than 50 years within 3 years.

Of the initial 1,847 at baseline PD-free participants 50 years or older, 1,271 underwent the 5-year reassessment.

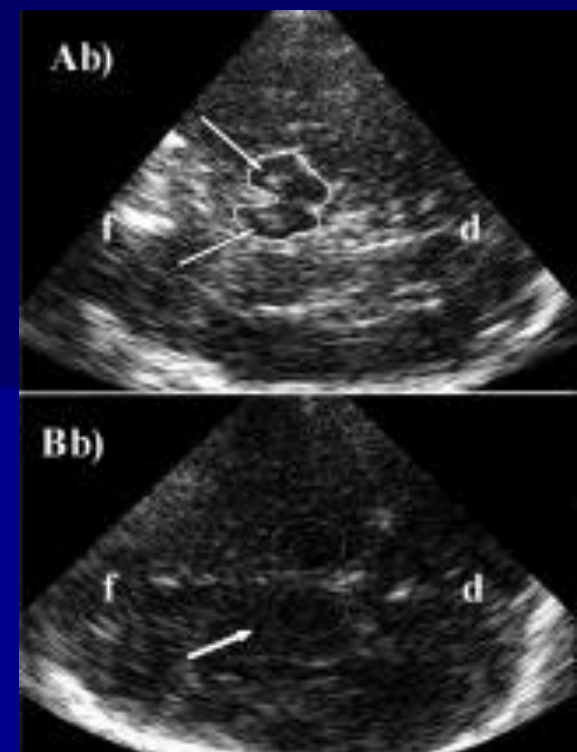
Participants with SN+ at baseline had a more than **20.6** times increased risk to develop PD in this time span than those without this echo feature.

Conclusions: We suggest SN+ as an important risk marker for PD.

Transcranial Sonography Findings in Welding-Related Parkinsonism(WRP) in Comparison to Parkinson's Disease

Uwe Walter, Dirk Dressler, Christian Lindemann,
Andrea Slachevsky, and Marcelo Miranda,
Movement disorders 2008

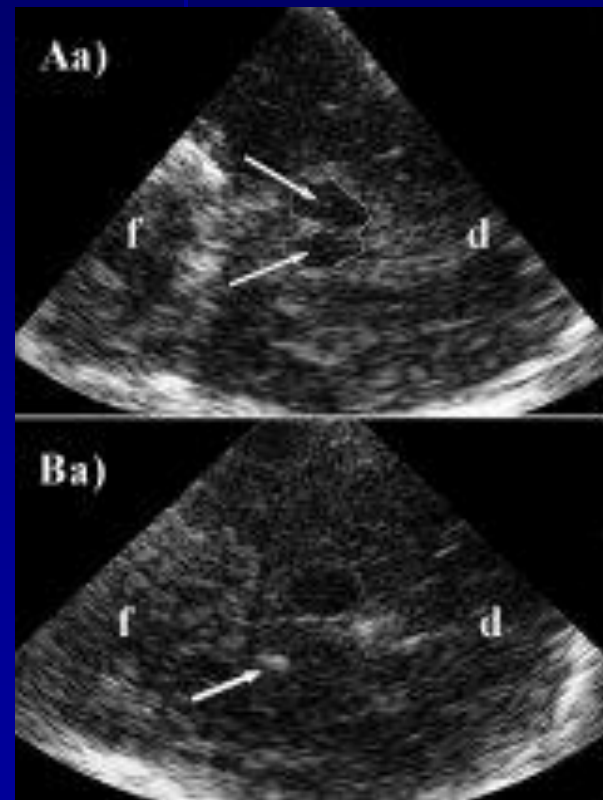
Idiopathic Parkinson's disease :
-marked hyperechogenicity of SN
lenticular nucleus with normal
echogenicity.



Welding-related Parkinsonism:



-Normal SN echogenicity
-lenticular nucleus with marked hyperechogenicity



Paz with WR-P developed a L-dopa-resistant akinetic-rigid parkinsonian syndrome at age <50 years

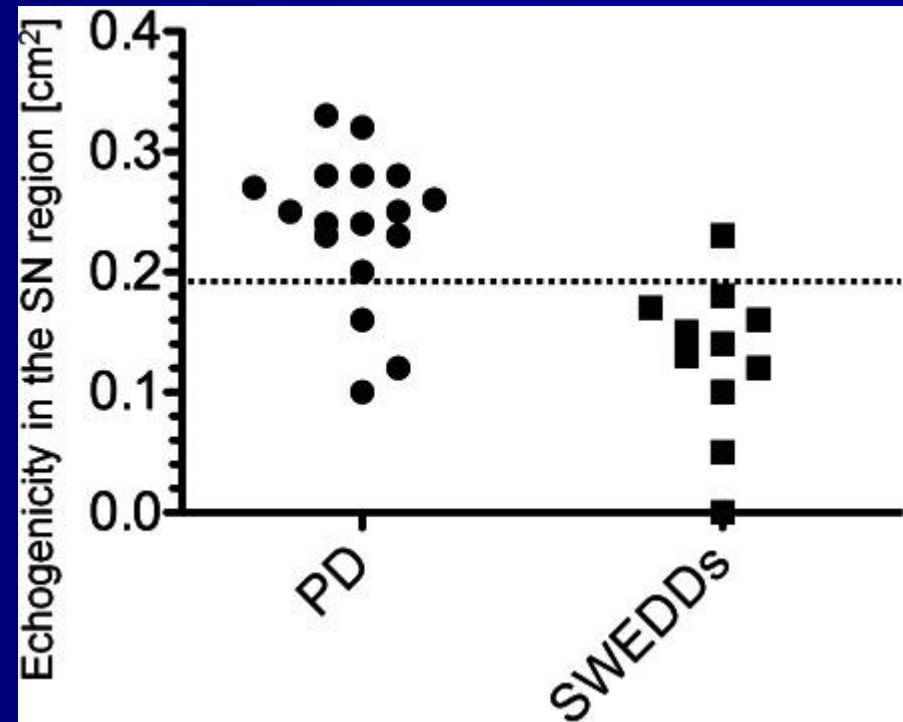
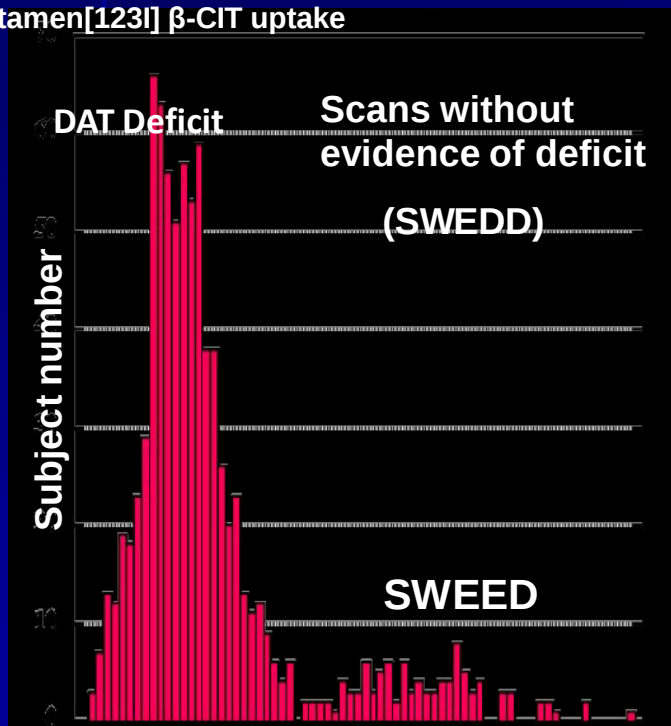
Is Transcranial Sonography Useful to Distinguish Scans Without Evidence of Dopaminergic Deficit Patients From Parkinson's Disease?

Heike Stockner, Petra Schwingenschuh, Atbin Djamshidian, Laura Silveira-Moriyama, Petra Katschnig, Klaus Seppi, John Dickson, Mark J. Edwards, Andrew J. Lees, Werner Poewe, Kailash P. Bhatia,

Movement Disorders 2012

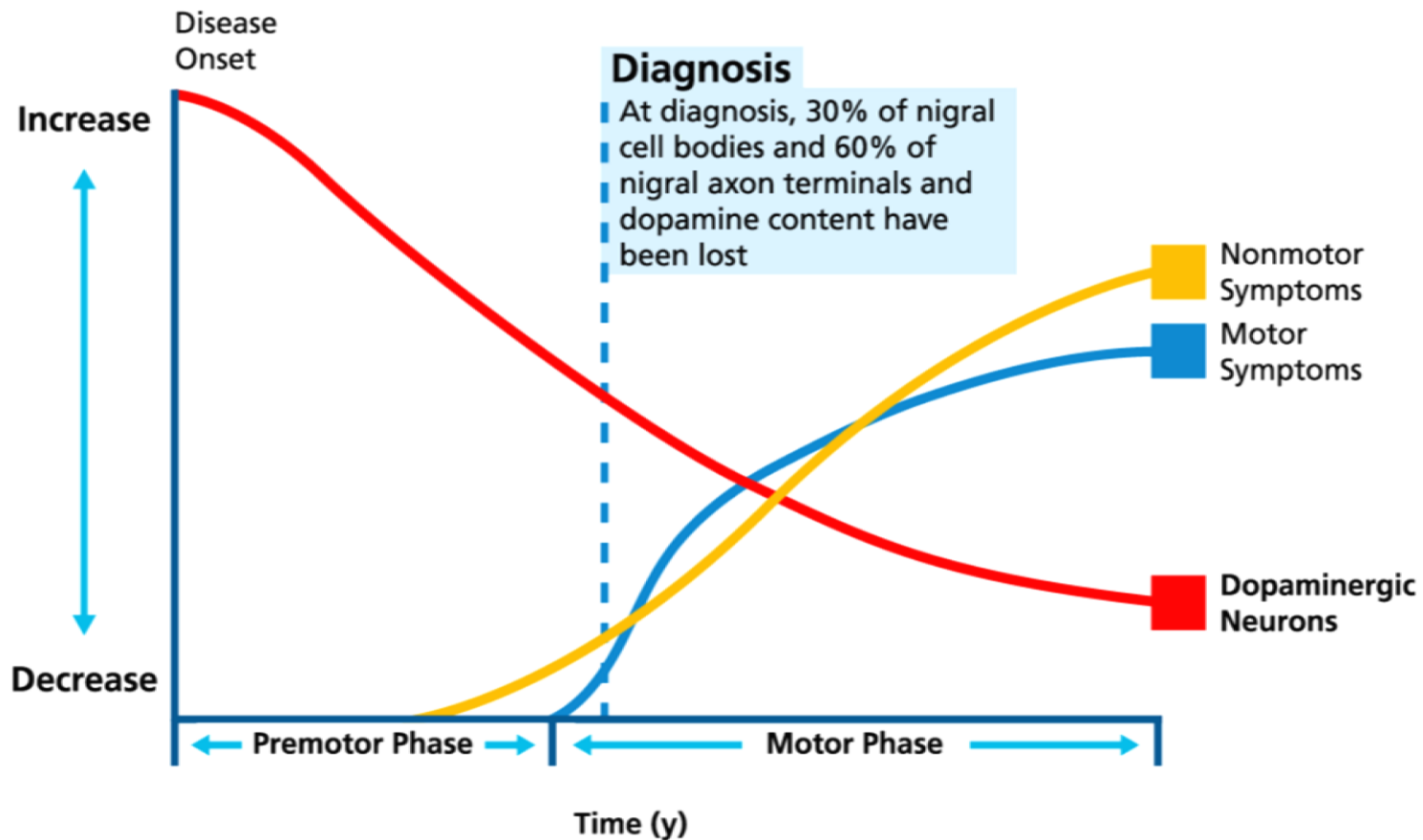
Baseline PRECEPT- % Age expected

Putamen[123I] β -CIT uptake

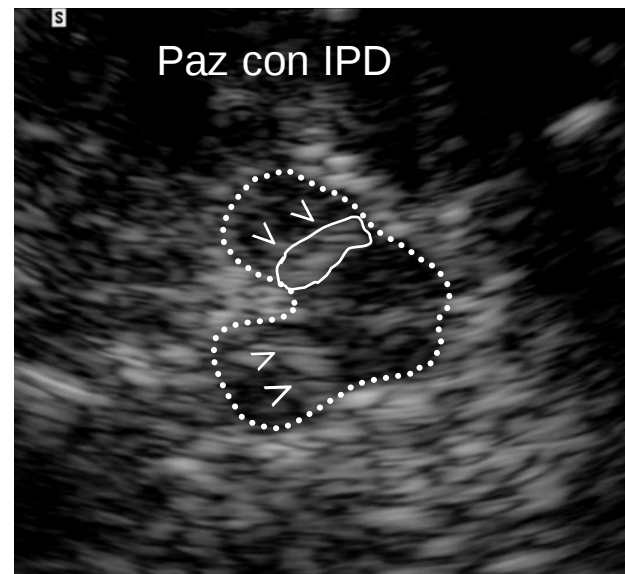
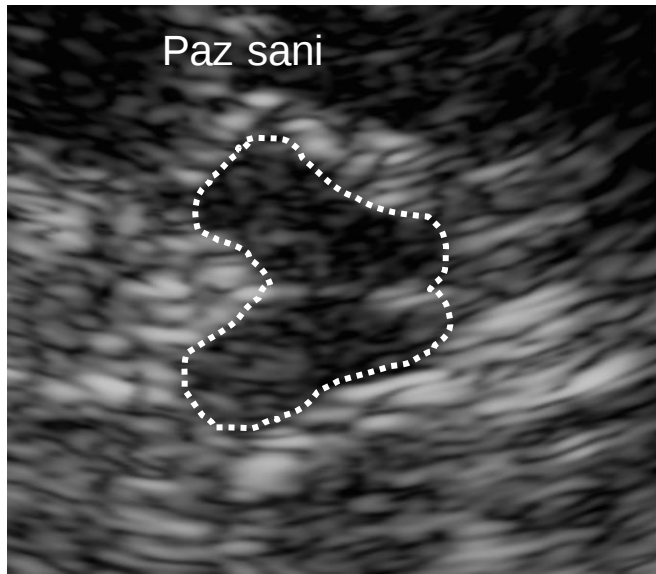


Significantly increased area of midbrain echogenicity in the IPD group compared with the SWEDD group ($P < 0.001$)

Storia naturale del Morbo di Parkinson



Echogenicity of the Substantia nigra



La iperecogenicità della SN è presente in oltre il 90% dei paz con Idiopathic Parkinson Disease(IPD):

(Berg et al., 2001; Walter et al., 2002)

Iron in the basal ganglia in
the Parkinson disease
Griffiths P.D. et al
Brain 1999

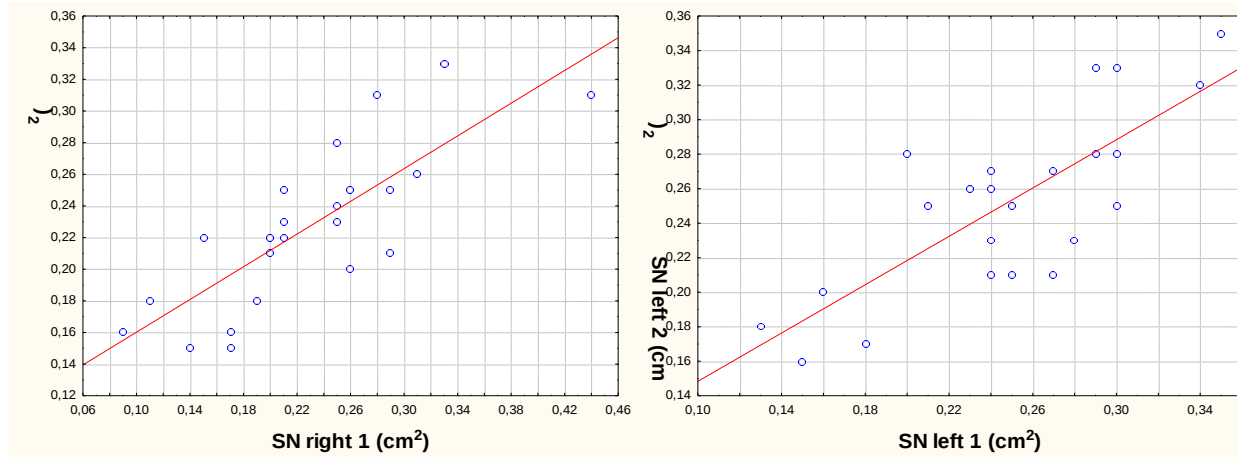
The relevance of iron in the pathogenesis of PD
Gotz ML et
ann NY Acad. Sci. 2004

Sequence variations in genes involved in iron metabolism in PD
Hochstrasser H. et al
Medgen 2003

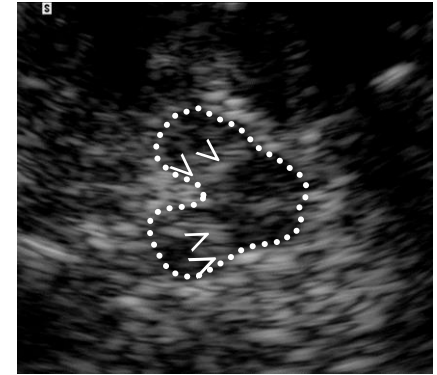
In vivo detection of iron and neuromelanin by transcranial sonography--a
new approach for early detection of substantia nigra damage
Berg D.
J Neural Transm.2006

«However, it is not clear yet, whether iron accumulation is a primary cause
or a secondary phenomenon in the disease process»

L'area della SN varia con l'età e il decorso del MDP?

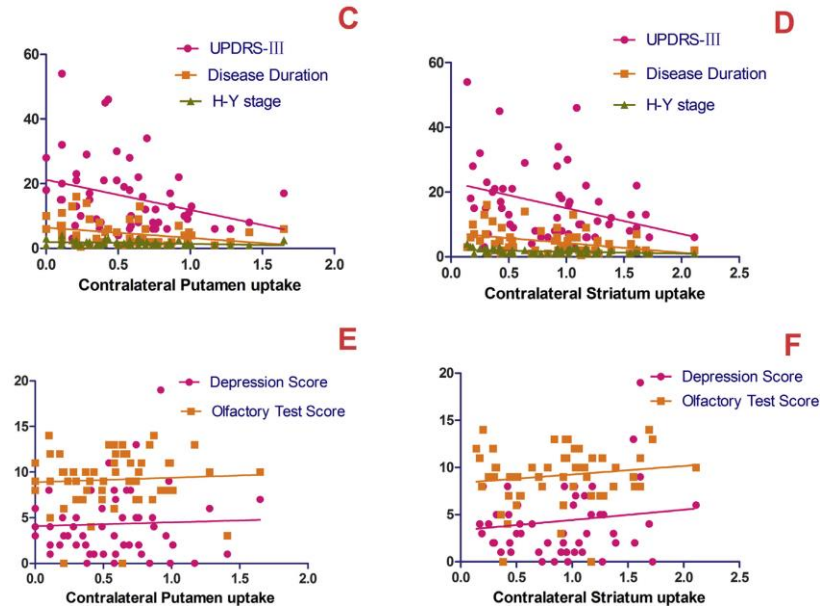


Berg et al. Mov Disord 2005



L'estensione dell'area della SN non è correlata con lo stadio della malattia

Li Park and related disorders 2015



Summary:neurosonologic patterns for differential diagnosis

Malattia	Substanza Nigra	Raphe Mesenc	Ventricoli	Gangli Basali
MDP Sporadico	↑↑	↓	-	-
Parkinsonismi Atipici	↑	-	↑↑ 3. Ventricolo	↑↑
Metabolico		-	-	↑↑
Idrocefalo	-	-	↑↑	-
Tremore Essenziale	(↑)	-	-	-
Depressione	(↑)	↓↓	-	-

Wilson Disease

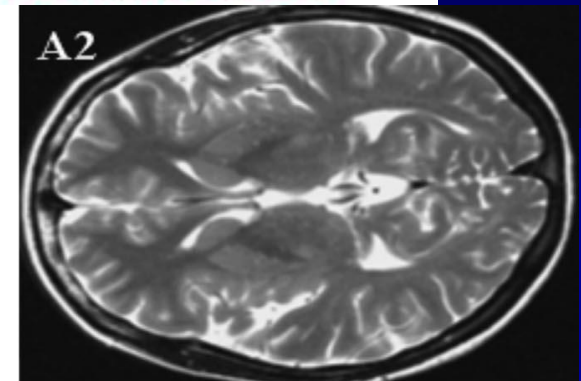
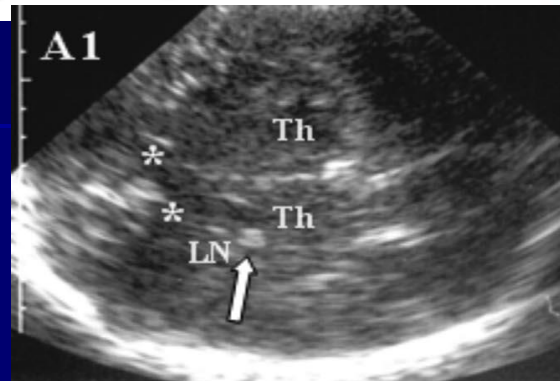


Sonographic detection of basal ganglia lesions in asymptomatic and symptomatic Wilson disease

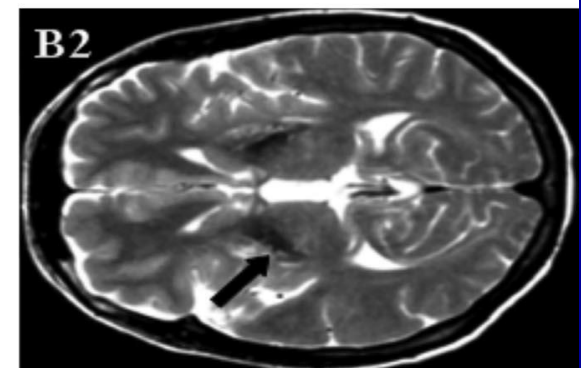
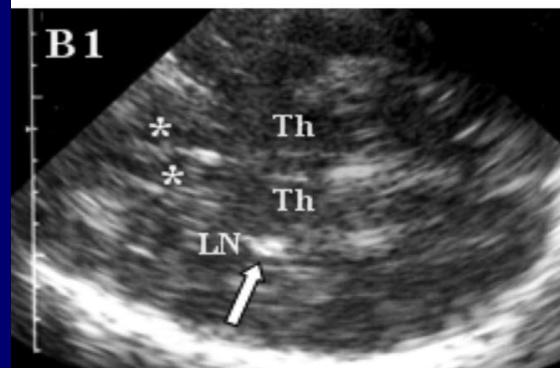
U. WALTER et al.

NEUROLOGY 2005

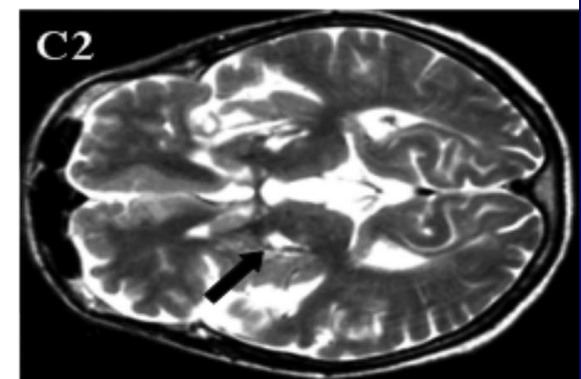
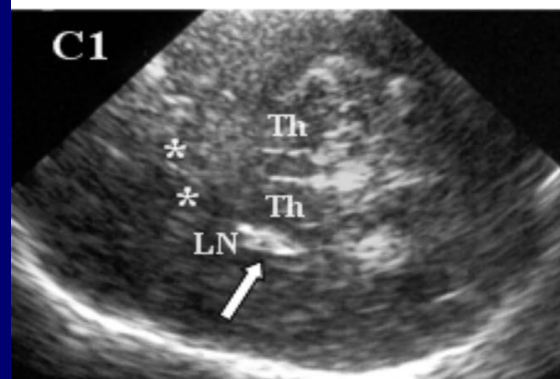
(A) Neurologically asymptomatic patient with WD. Whereas MRI (A2) shows normal LN, TCS (A1) depicts abnormal, dot-like hyperechogenicity of the lenticular nucleus (arrow).



(B) Neurologically moderately affected patient with WD. TCS (B1) shows abnormal hyperechogenicity (arrow) of the LN, corresponding to hypointense MRI lesion (B2, arrow).



(C) Neurologically severely affected patient with WD. TCS (C1) displays abnormal LN hyperechogenicity (arrow), corresponding to hyperintense and hypointense lesions on MRI (C2, arrow).

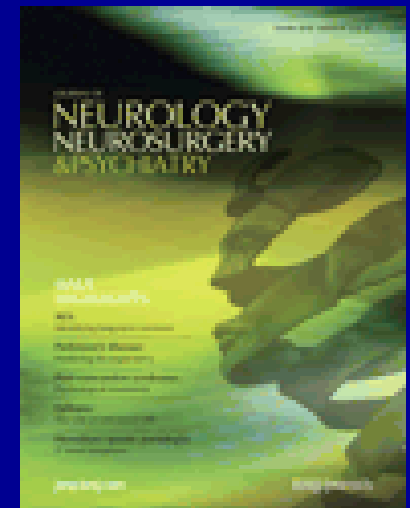


Reproducibility and diagnostic accuracy of substantia nigra sonography for the diagnosis of Parkinson's disease

van de Loo S., Walter U., Behnke S., Hagenah J.,
Lorenz M., Sitzer M., Hilker R., Berg D.

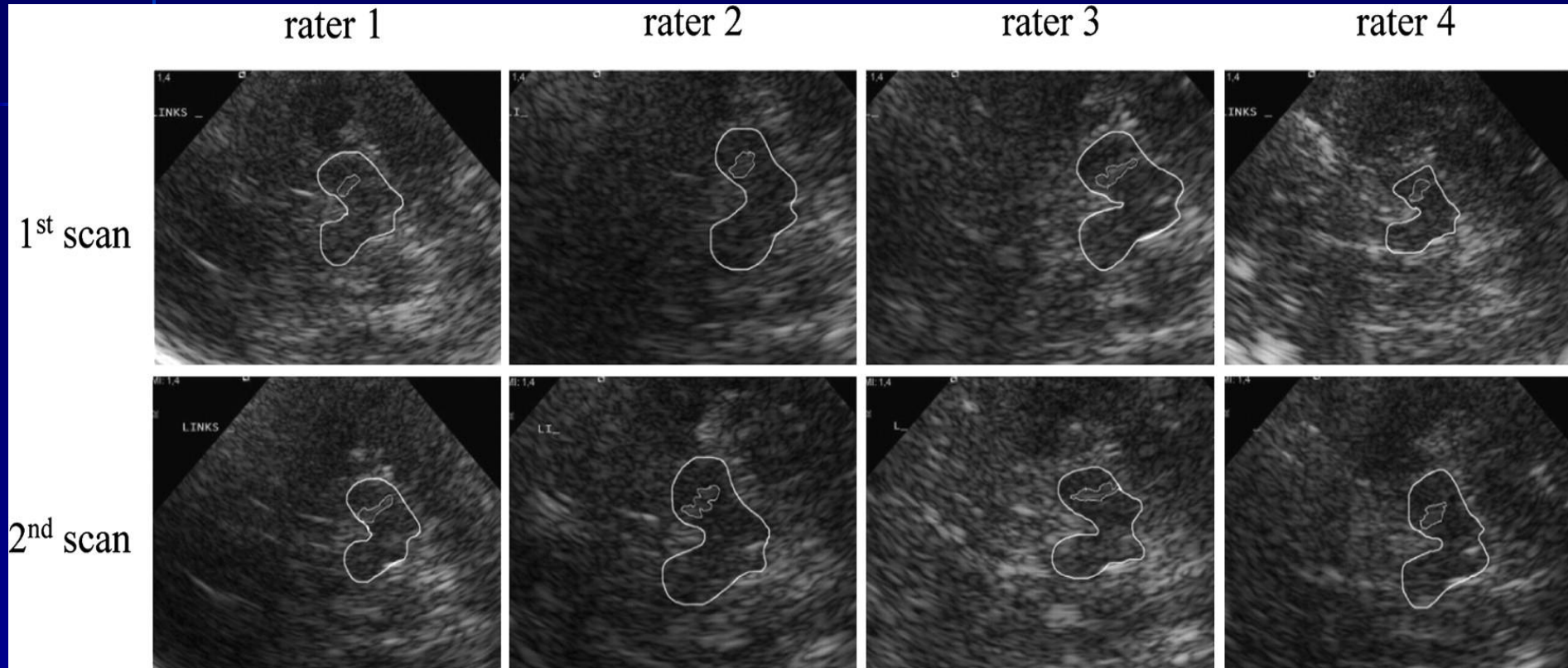
J Neurol Neurosurg Psychiatry.2010
Oct;81(10):1087-92

a hyperechogenic area of 0.24 cm^2 as the optimum cut-off value for the differentiation between PD and controls (sensitivity 79%, specificity 81%).



Transaxial midbrain B-mode sonography of the substantia nigra (SN) measured **twice by **four** raters in the **same** Parkinson's disease patient with good transtemporal bone window quality (first scan: upper row; second scan: lower row).**

van de Loo S et al. J Neurol Neurosurg Psychiatry
2010;81:1087-1092



Reliable SN TCS data on PD can be achieved in clinical routine and multicentre trials when standardised analysis protocols and certain quality criteria of brain parenchyma sonography are met.

JNNP

Conclusions: “The data demonstrate that the observer variability of SN planimetry is low only in the hands of experienced investigators”.

Cerebral Autosomal Dominant Arteriopathy With Subcortical Infarcts and Leukoencephalopathy (**CADASIL**)

Stroke

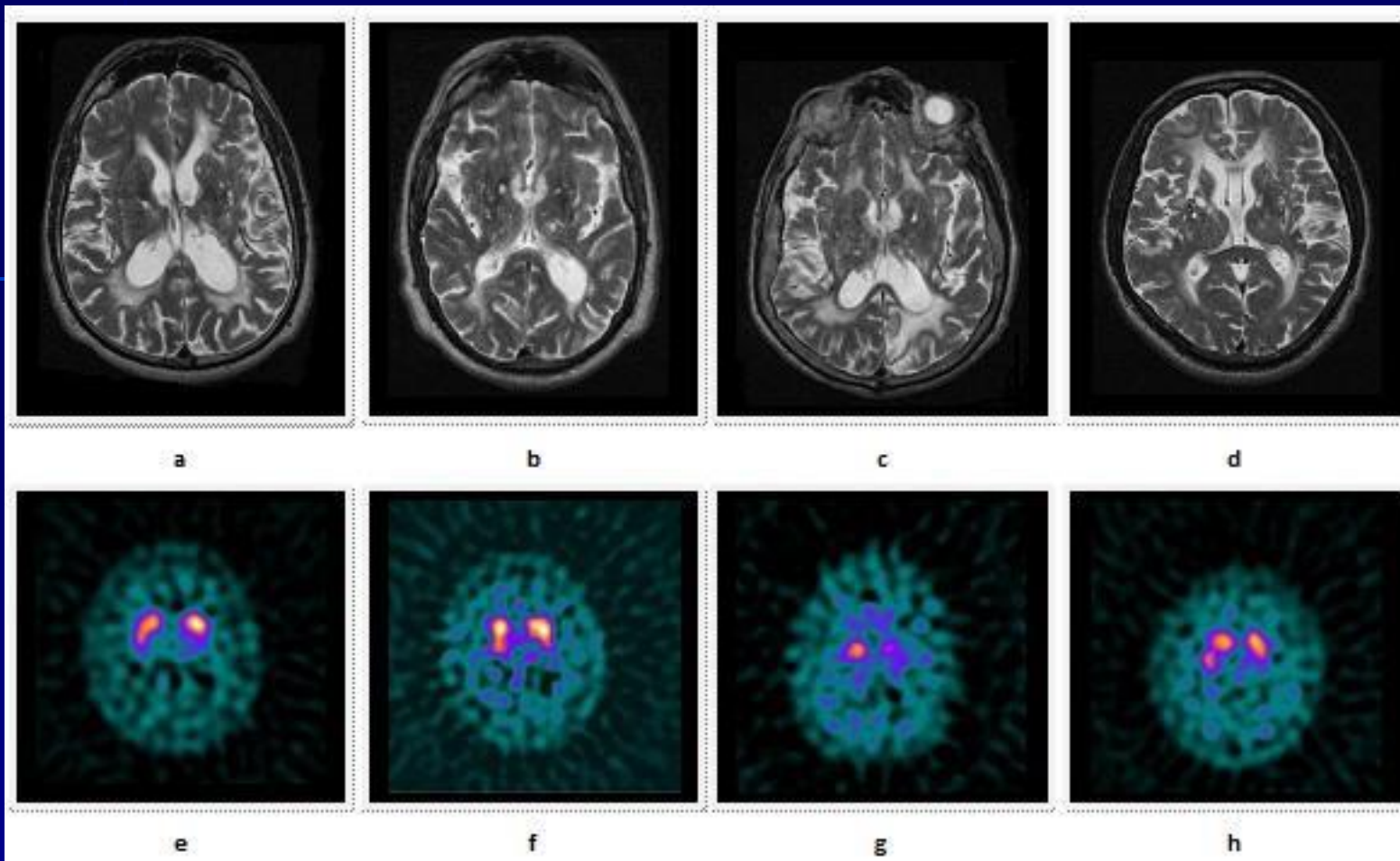
JOURNAL OF THE AMERICAN HEART ASSOCIATION



Parkinsonism is a late, not rare, feature of
CADASIL: a study on Italian patients carrying the
R 1006 C mutation.

Ragno M. et al

Stroke Apr;44(4):1147-9. 2013



Axial brain MRI showing: (a) multiple lacunae of the basal ganglia, more evident on the right pallidus and left caudate head on T2-weighted images ; (b) small basal ganglia lacunae and ischemic lesions of the left ventral thalamus and right globus pallidus on T2-weighted spin echo image ; (c) cortical infarct in the left temporo-occipital region and lacunae in the left pallidus and right thalamus on T2-weighted images ; (d) ischemic infarction of the right lentiform nucleus on T2-weighted spin echo images .

[123I]-FP-CIT showing: (e) slight reduction of left putamen uptake; (f) bilateral low putaminal uptake; (g) bilaterally reduced uptake in the caudate and putamen ; (h) moderate low uptake at the right caudate and bilaterally reduced putaminal uptake.

Brain Parenchyma Sonography(BPS) in pat. with Cadasil

Our ongoing experience

PHILIPS 17:06:29 TIC3.3 MI 1.4
08011720131010 NEUROLOGIA - S.B.T. S5-1/TCCD

FR 15Hz
R1

2D
55%
C 58
P Med.
AGen

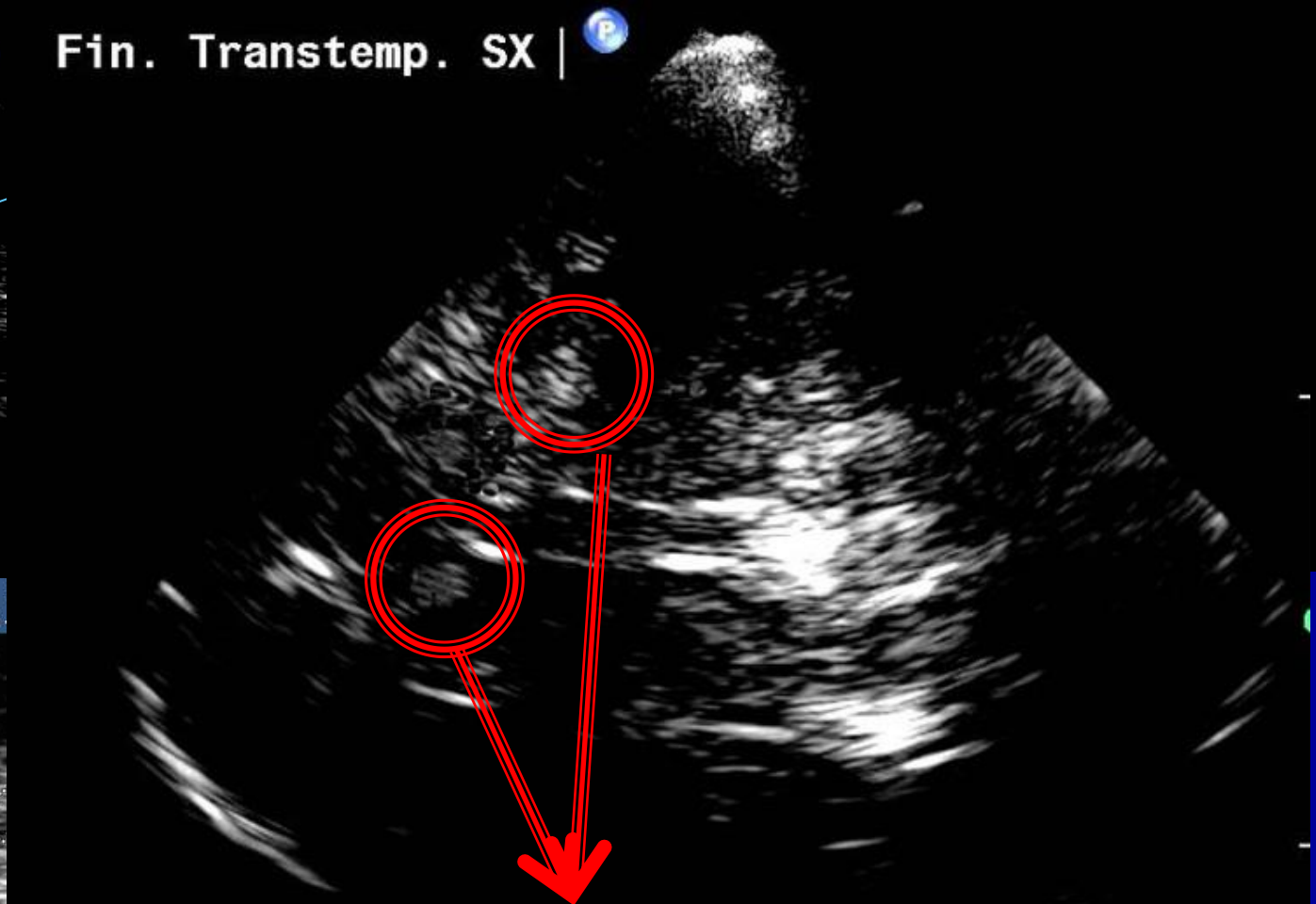
Fin. Transtemp. SX |

Fin. Transtemp. SX |

PHILIPS 17:07:00 TIC3.3 MI 1.4
08011720131010 NEUROLOGIA - S.B.T. S5-1/TCCD

FR 15Hz

M3



-Bilateral ipercogenicity of LN

PHILIPS 04092020130627 NEUROLOGIA

FR 15Hz
R1
Z 2.4
2D
58%
C 58
P Med.
AGen

DX

✧ Area 5.87 cm²
✧ Area 0.237 cm²

task force sottolinea che questa tecnica non è universalmente utilizzata e richiede una particolare esperienza. Inoltre a causa della bassa specificità dell'ecografia transcranica nella diagnosi di malattia di Parkinson, questa tecnica dovrebbe essere utilizzata insieme ad altri test.

Considerando le finalità delle raccomandazioni redatte in questa linea guida, orientate verso la pratica clinica corrente e la sanità pubblica, e le difficoltà nell'esecuzione del test (con la necessità di un operatore esperto e la limitazione imposta all'esecuzione dell'esame subordinata alla qualità della finestra trans temporale) si ritiene di non dover raccomandare l'uso dell'ecografia transcranica nella malattia di Parkinson.

Raccomandazioni

C

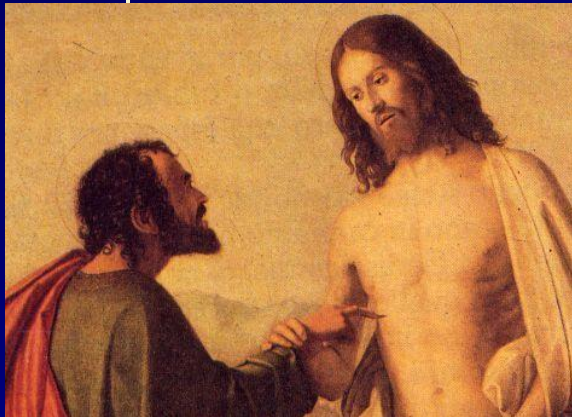
L'ecografia transcranica parenchimale non dovrebbe essere utilizzata per la diagnosi differenziale tra malattia di Parkinson idiopatica e altre condizioni associate, se non all'interno di specifici protocolli di ricerca.

Raccomandazione per la ricerca

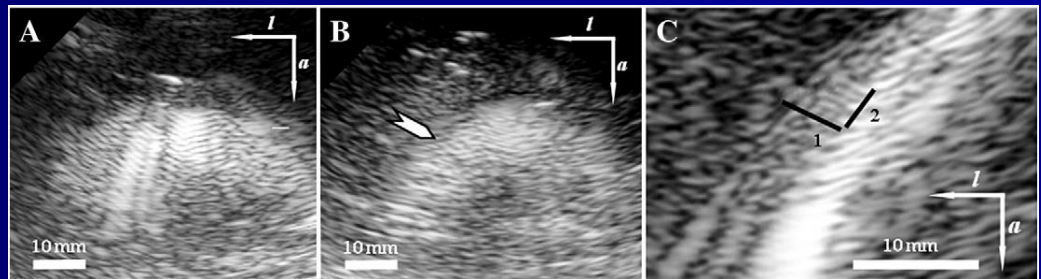
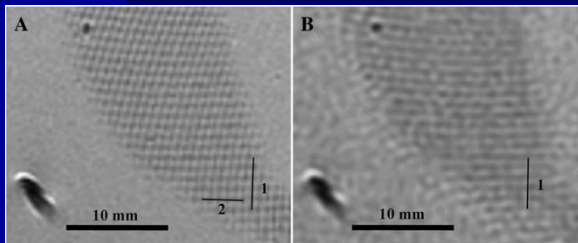
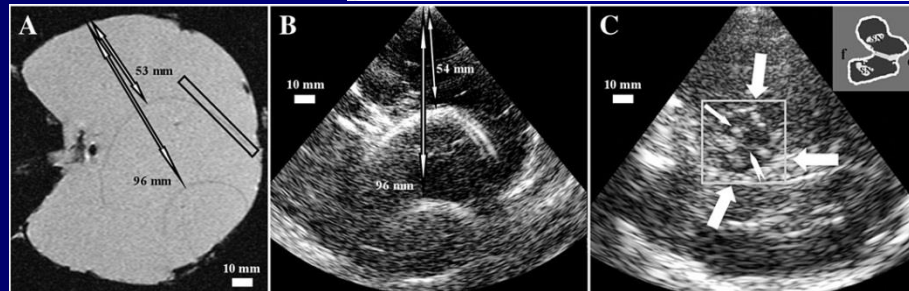
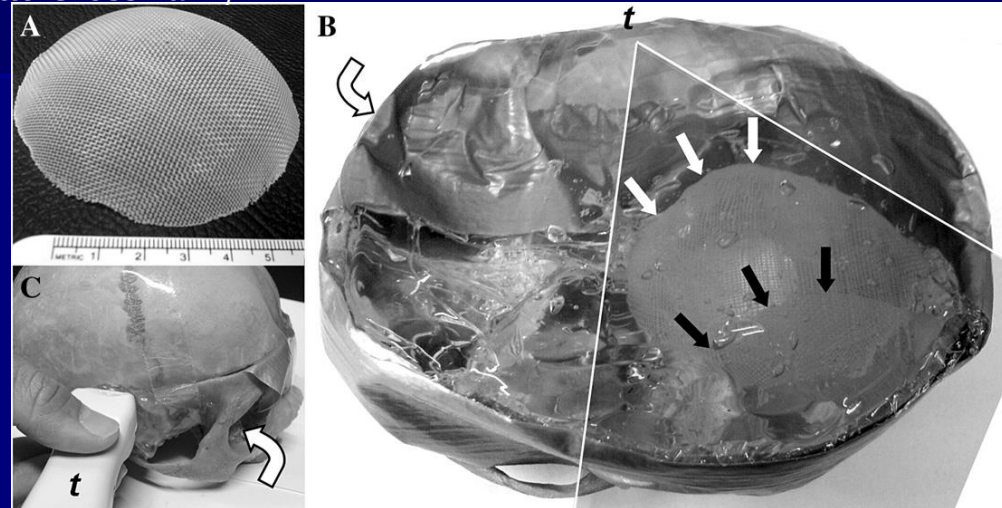
In considerazione dei risultati raggiunti sono sollecitati ulteriori studi relativi all'impiego dell'*imaging* strutturale cerebrale (TC, RM, ecografia transcranica parenchimale) per la diagnosi differenziale tra malattia di Parkinson e altri parkinsonismi degenerativi.

Contemporary ultrasound systems allow high-resolution transcranial imaging of small echogenic deep intracranial structures similarly as MRI: A phantom study

Uwe Walter, Martin Kanowski, Jörn Kaufmann, Annette Grossmann, Reiner Benecke, and Ludwig Niehaus



L-558



Future applicazioni

Se:8
Im:18

[A]

[R]

[P]

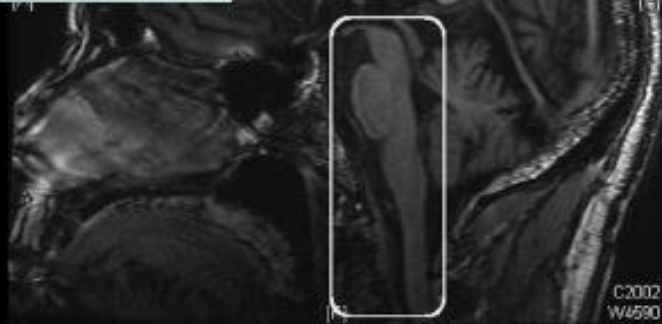


Hot-cross bun

Se:401
Im:26

[H]

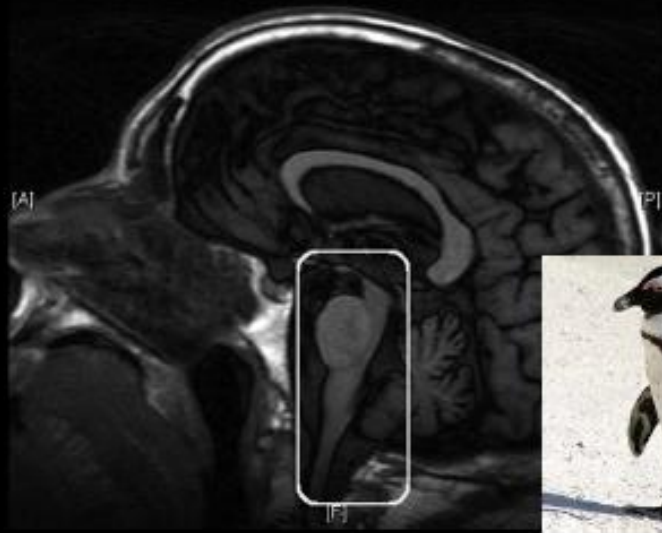
A



Se:2
Im:12

[H]

B

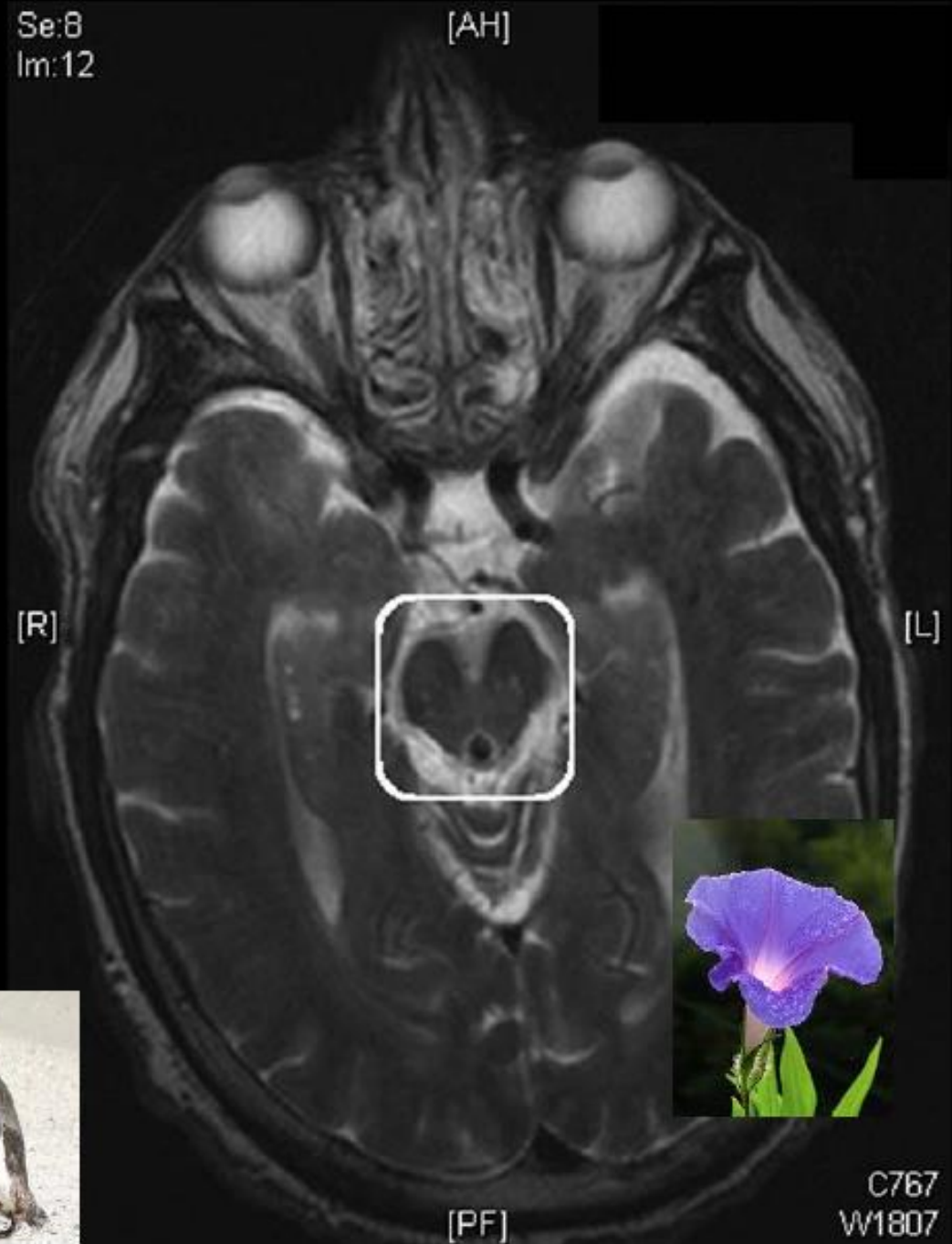


Se:8
Im:12

[AH]

[R]

[L]

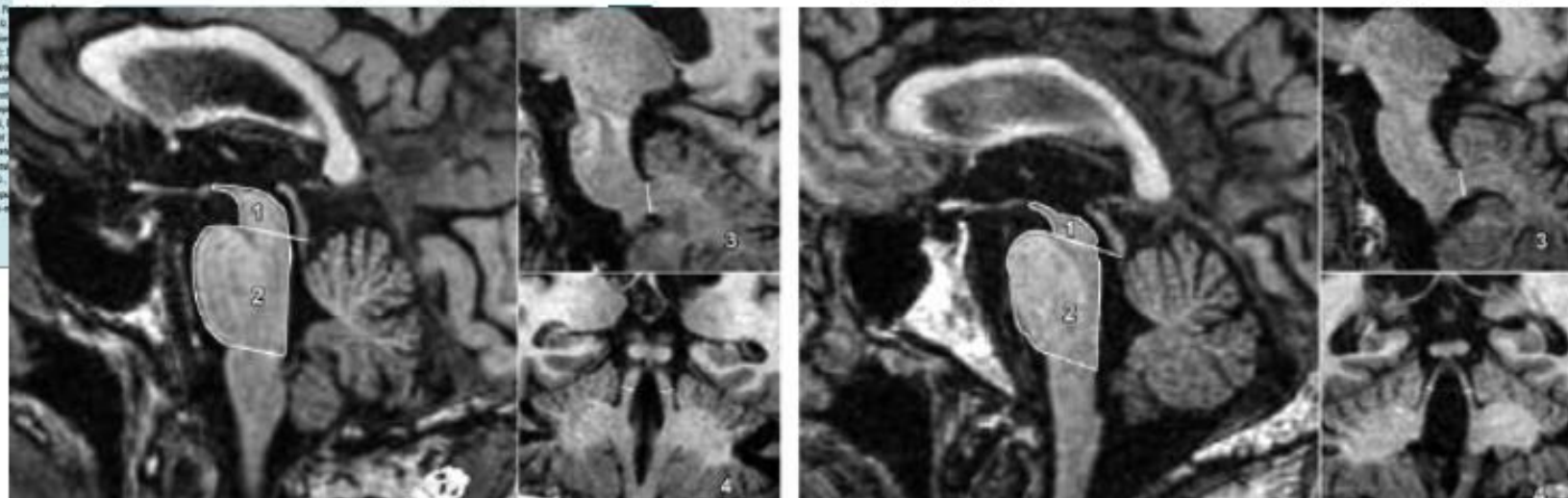
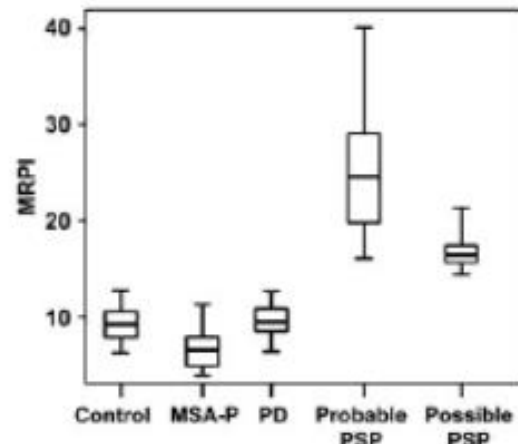
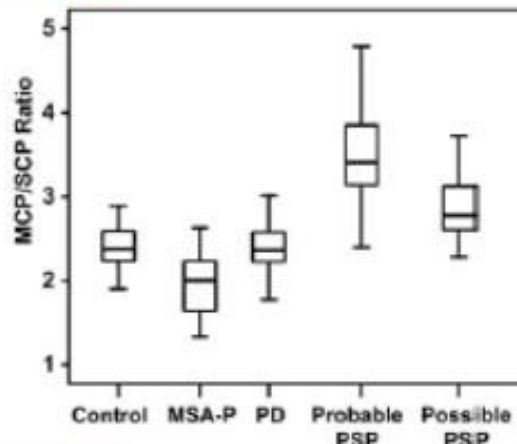
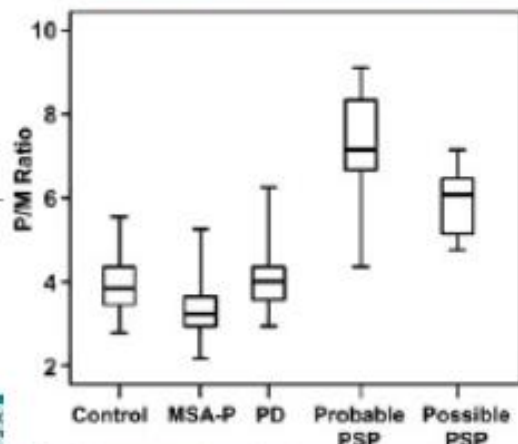


C767
W1807

MR Imaging Index for Differentiation of Progressive Supranuclear Palsy from Parkinson Disease and the Parkinson Variant of Multiple System Atrophy¹

Radiology: Volume 246: Number 1—January 2008

Purpose: To prospectively assess sensitivity and specificity of mag-



Aldo Quattrone, MD
Giuseppe Nicoletti, MD
Domenico Massimo, MD
Francesca Fenu, MD
Francesca Condini, PhD
Pierfrancesco Pugliese, MD
Pierluigi Lanza, MD
Paolo Barone, MD
Ludovico Morgante, MD
Mario Zappalà, MD
Umberto Agaglia, MD
Oliviero Galati, STC

¹ From the Institute of Neurology (A.Q.), Department of Neurology (P.F.), Magna Graecia University, Catanzaro, Calabria, Italy; Institute of Sciences, National Research Council, P.le Marconi, Genova, Genova, Italy (A.Q., G.O.); Department of Neurological Sciences, Naples Federico II, Naples, Italy (P.B.); Department of Neurology and Institute of Neurology, University of Messina, Messina, Italy (L.M.); Department of Neurosciences, University of Bari, Bari (M.Z.), and Regional Hospital, Ospedale Riuniti Medico Neurologico, Catanzaro, Italy (P.A.). Received October 12, 2007; accepted February 20, 2008; final version 14. Address correspondence to A.Q., quattrone@uniba.it.

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Magnetic Resonance-Transcranial Ultrasound Fusion Imaging: A Novel Tool for Brain Electrode Location

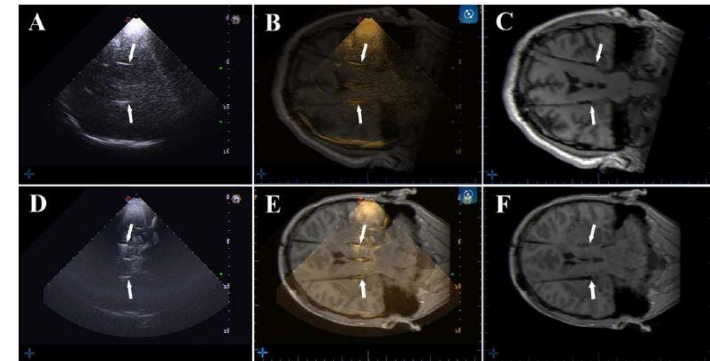
Uwe Walter, MD,^{1*} Jan-Uwe Müller, MD,² Johannes Rösche, MD,¹ Michael Kirsch, MD,³ Annette Grossmann, MD,⁴
Reiner Benecke, MD,¹ Matthias Wittstock, MD,^{1†} and Alexander Wolters, MD^{1†}

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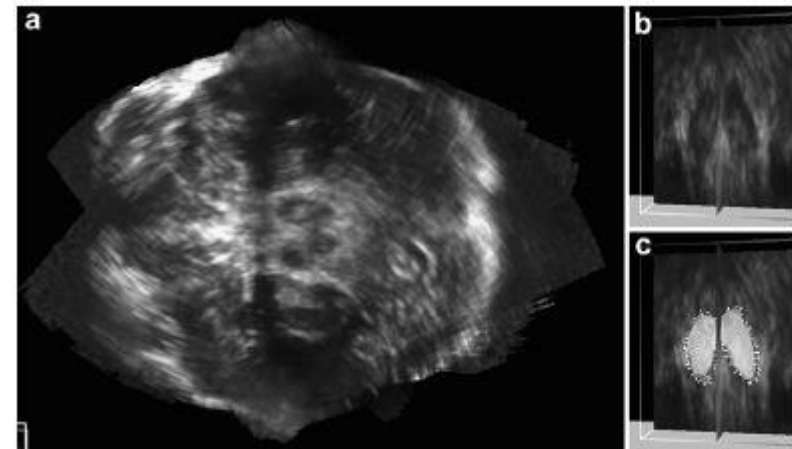
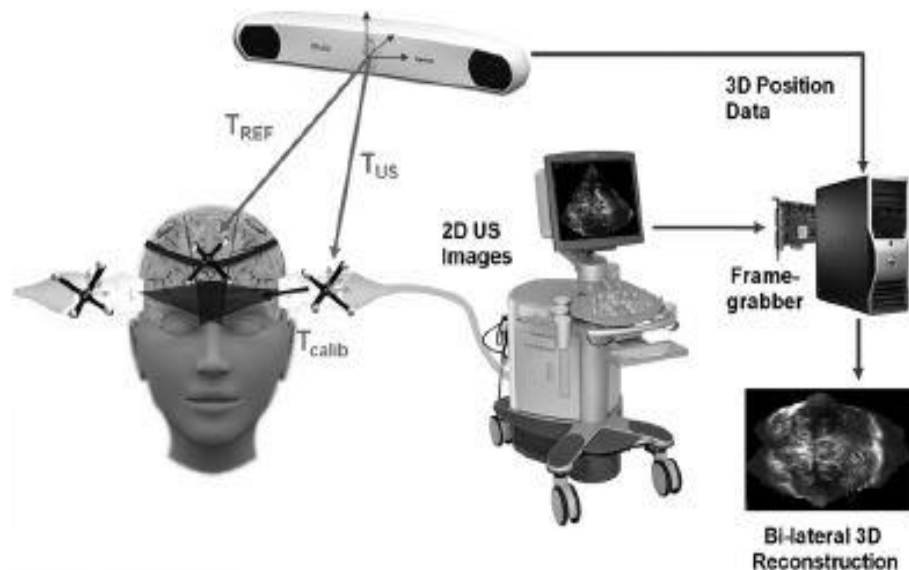


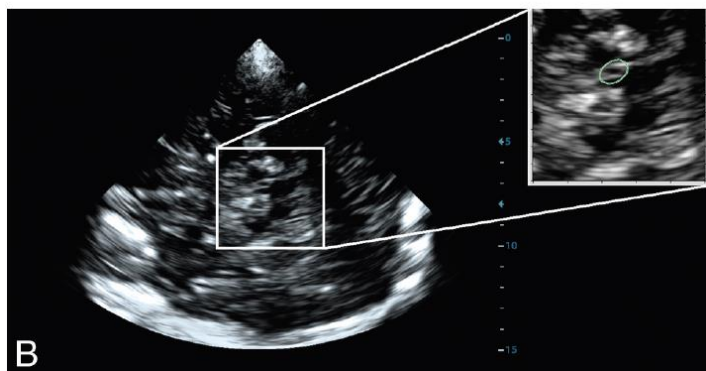
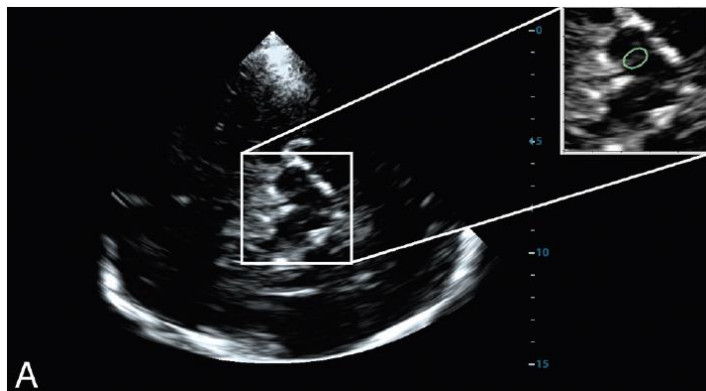
THREE-DIMENSIONAL SONOGRAPHIC EXAMINATION OF THE MIDBRAIN FOR COMPUTER-AIDED DIAGNOSIS OF MOVEMENT DISORDERS

ANNIKA PLATE,^{*} SEYED-AHMAD AHMADI,[†] OLIVIER PAULY,^{††} TASSILO KLEIN,[†] NASSIR NAVAB,[†]
and KAI BÖTZEL^{*}

^{*}Department of Neurology, Ludwig-Maximilians-Universität München, Munich, Germany; [†]Computer Aided Medical Procedures (CAMP), Technische Universität München (TUM), Garching at Munich, Germany; and ^{††}Institute of Biomathematics and Biometry, Helmholtz Zentrum München, Germany

3-D sonographic examination of the midbrain ● A. PLATE *et al.*

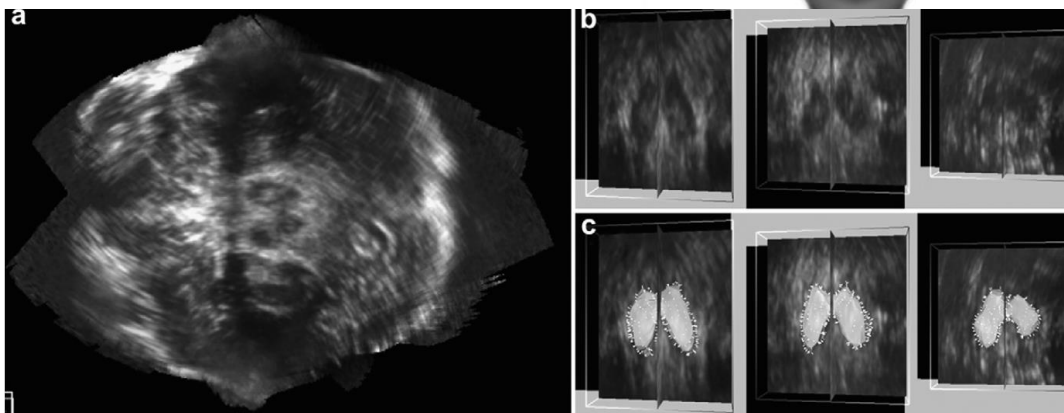
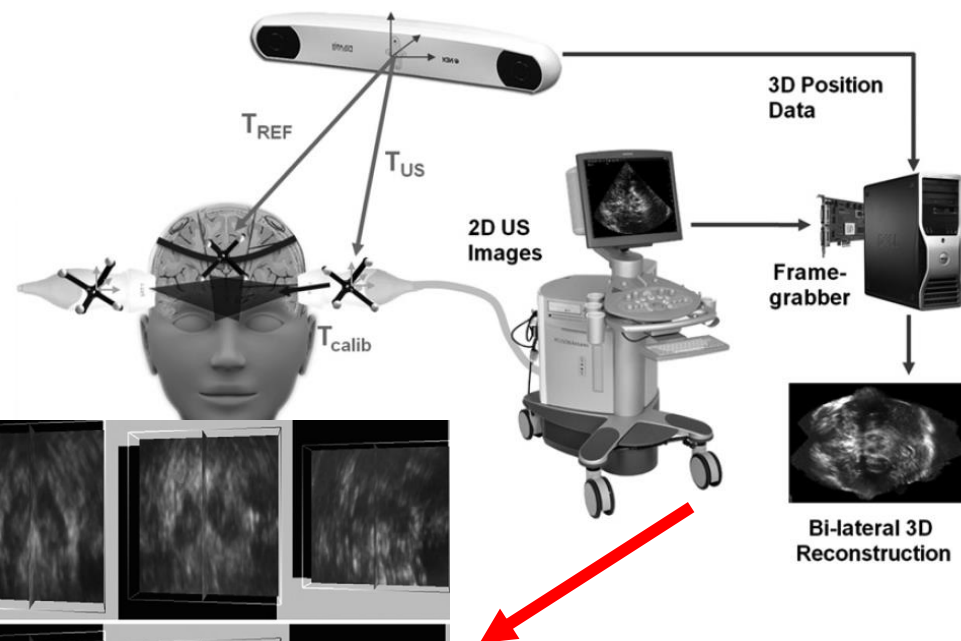




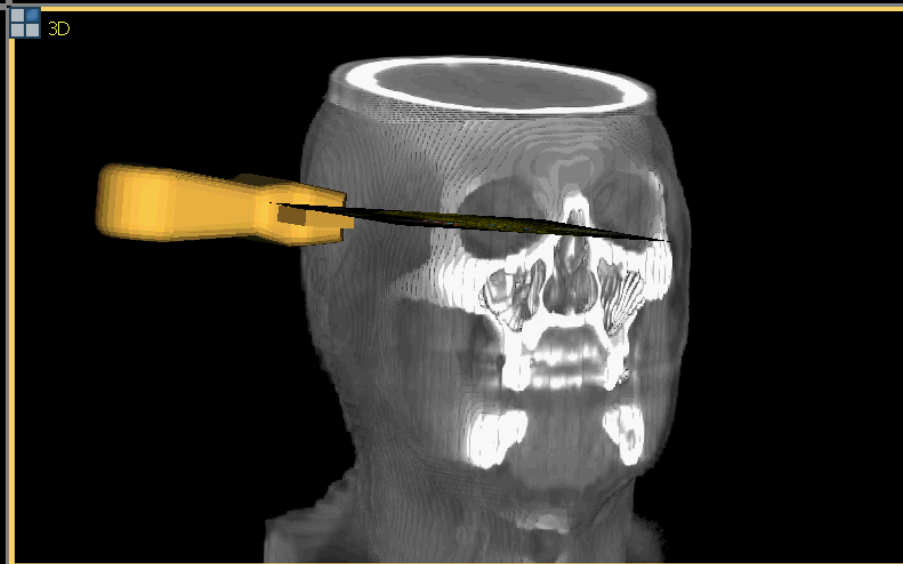
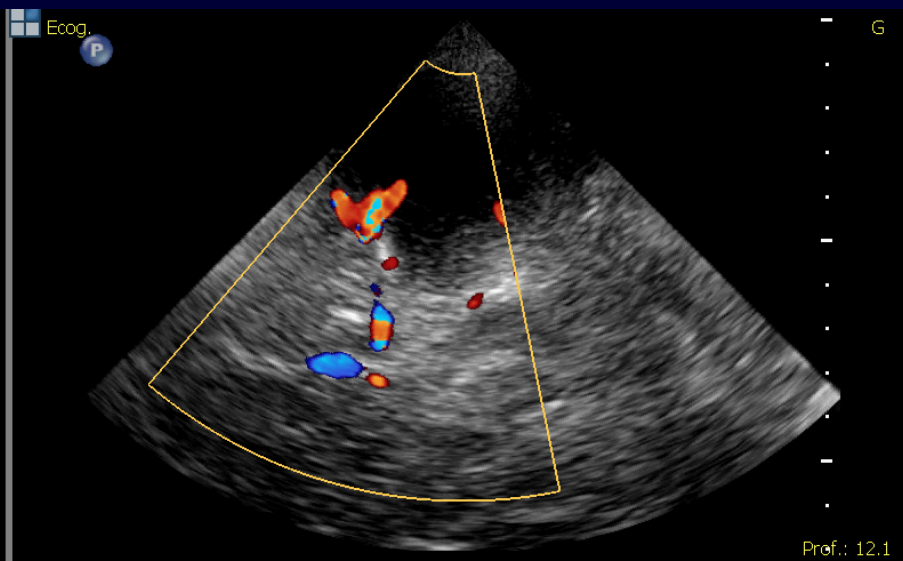
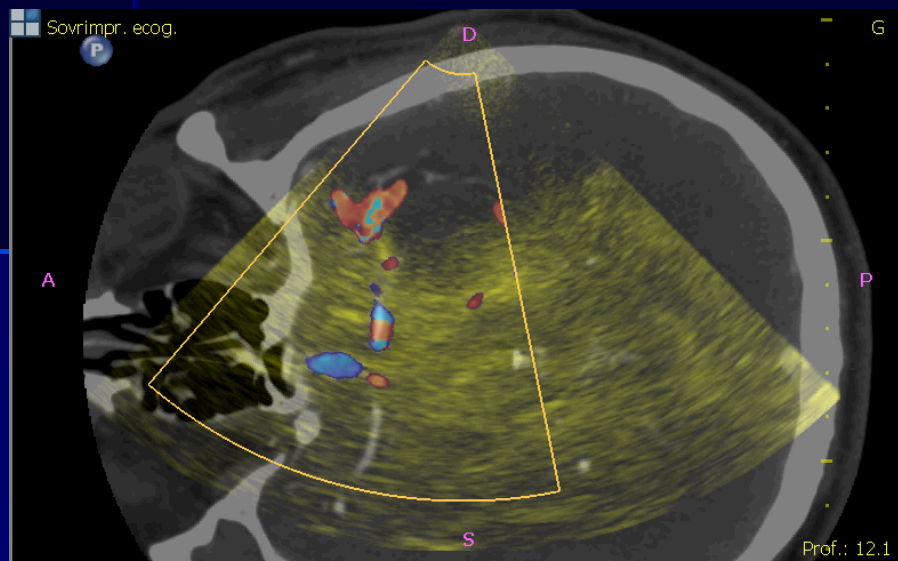
Skoloudik 2014 AJNR

- active contour algorithms
- 3D SN detection based
(random forests vs principal component analyses of artificial neural networks)

still dependent on the image's quality
and on operator's skills).

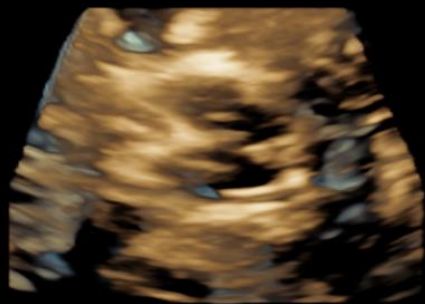
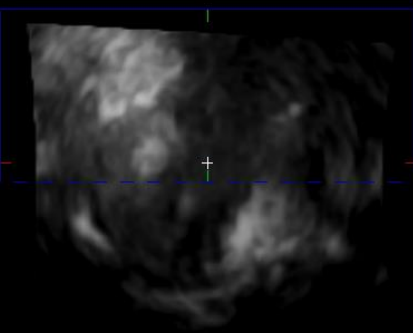
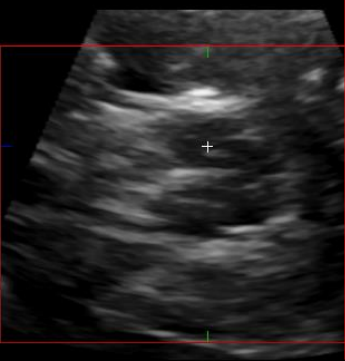
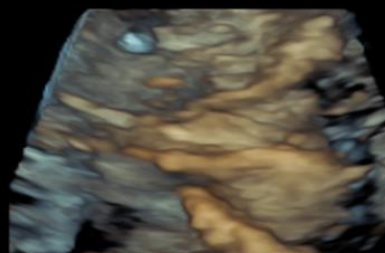
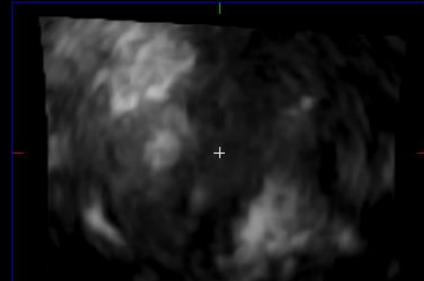
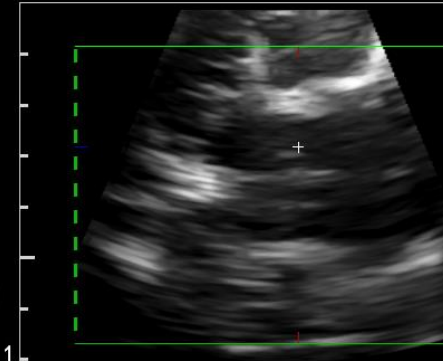
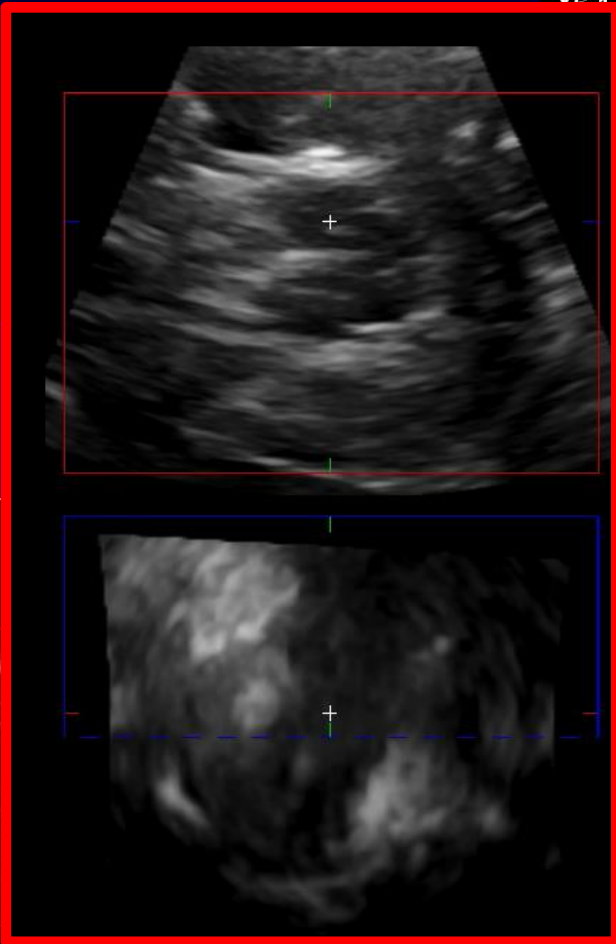


IL CONCETTO DI «FUSION» IN NEUROLOSONOLOGIA



TCD
X6.4

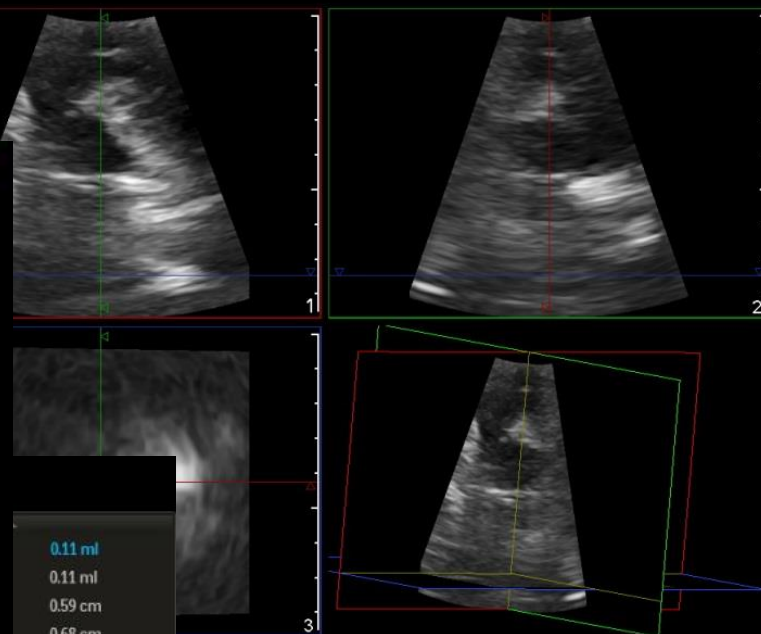
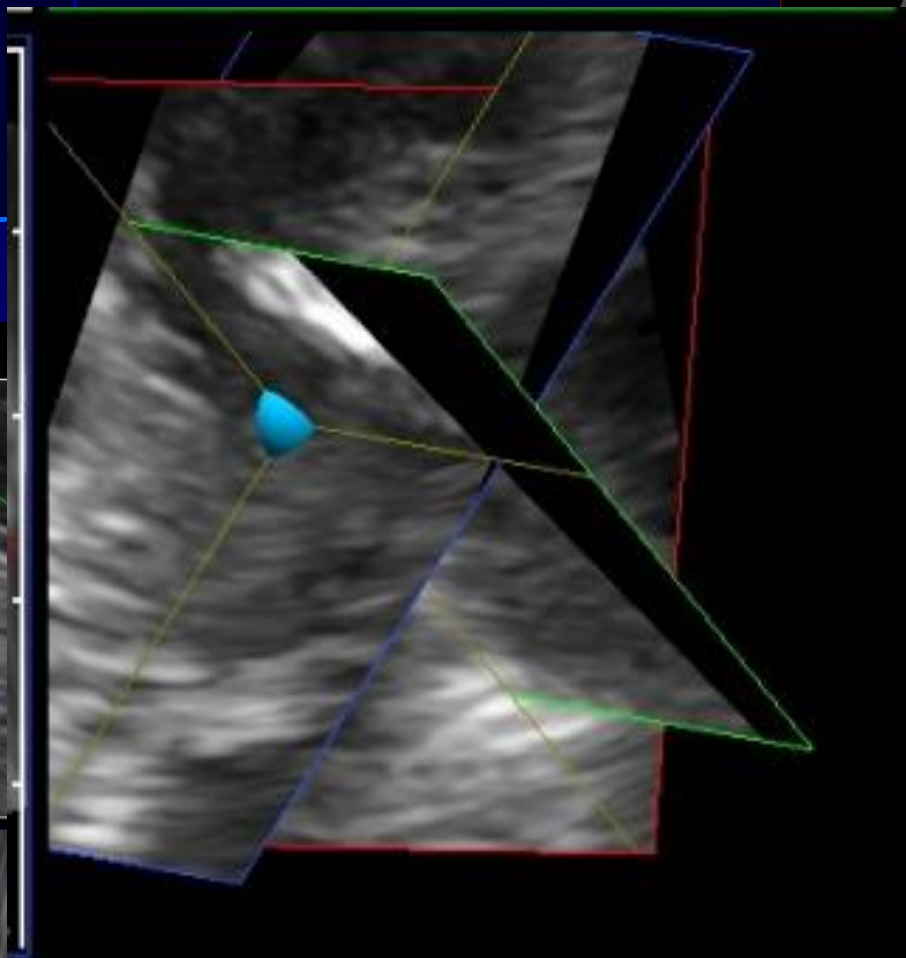
1150.3 M



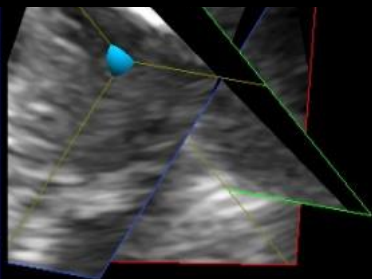
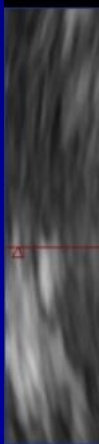
3

4

4



0.11 ml
0.11 ml
0.59 cm
0.68 cm
0.54 cm



3



ARTERIAL STUDY
VENOUS STUDY
PARENCHYMAL STUDY





Dr.Ragno M.

C.Sala Sig.ra Rossi M.

Dr. S.Sanguigni

Dr.T.Carboni

Dr.R.Gobbato

Dr.ssa C.Paci

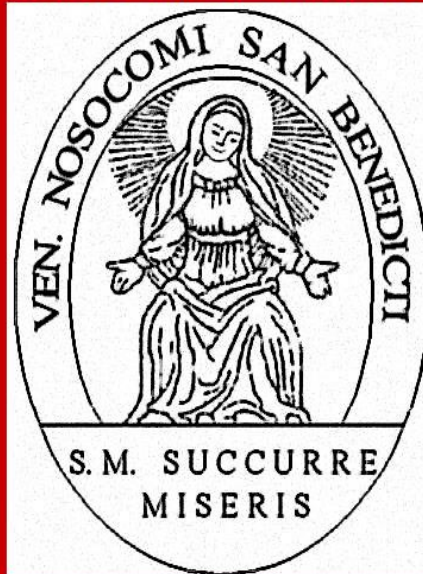
Dr.G.D'Andreamatteo

Dr.F.Di Marzio

Dr. E. Puca

Dr.ssa S.Renzi

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UNITA' OPERATIVA DI NEUROLOGIA

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Tnfp:

S.Sobrini

V.Urbano

A.Brandimarti

L.Iannelli



San Benedetto del Tronto

GRAZIE PER L'ATTENZIONE