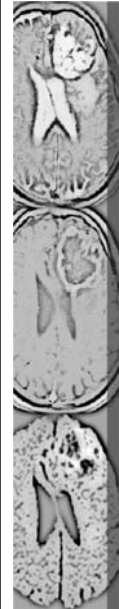




## Magnetic Resonance in Medicine MR Spectroscopy (MRS)

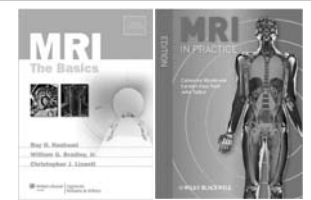
Chia-Feng Lu (盧家鋒), Ph.D.  
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and Radiological Sciences, NYCU  
[alvin4016@nycu.edu.tw](mailto:alvin4016@nycu.edu.tw)



## Content <http://cflu.lab.nycu.edu.tw/>

- Principle of  $^1\text{H}$  MR Spectroscopy (MRS)
  - 磁振頻譜分析
- MRS Pulse Sequences

- MRI The Basics (3rd edition)
  - Chapter 29: MR Spectroscopy
- MRI in Practice, (4th edition)
  - Chapter 12: Functional Imaging Techniques



<http://cflu.lab.nycu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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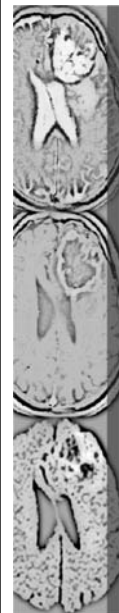
## Principles of $^1\text{H}$ MR Spectroscopy

磁振頻譜分析簡介

<http://cflu.lab.nycu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

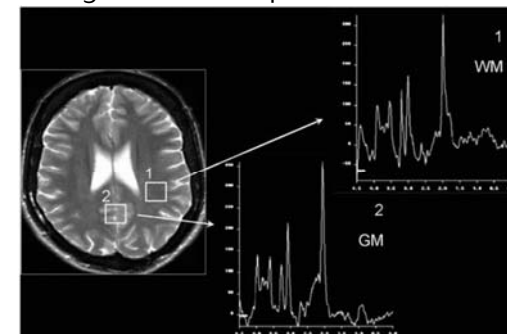
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## What's MR Spectroscopy

- Rather than providing images, it usually provides spectra consisting of individual peaks, the chemical shift of metabolites.



Bio-chemistry information

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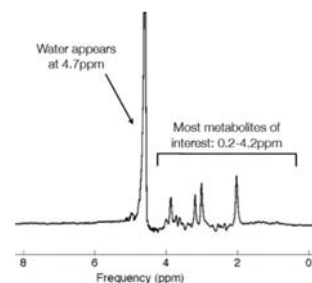
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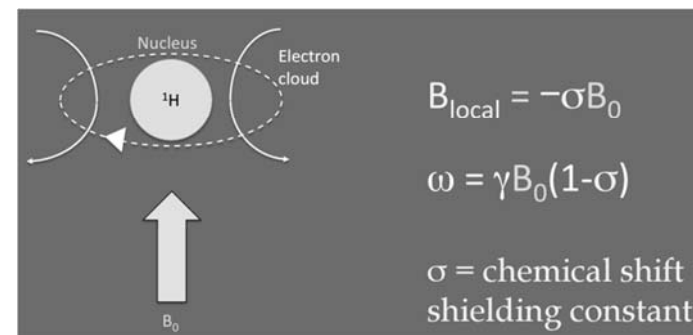
## <sup>1</sup>H Proton Spectroscopy

- Proton spectroscopy is easier to perform and provides much higher SNR than either sodium or phosphorus.
- Proton concentration in water: about 100 M
- Other metabolites: 1~10 mM

Need to suppress the water signal to investigate the signals from metabolites!

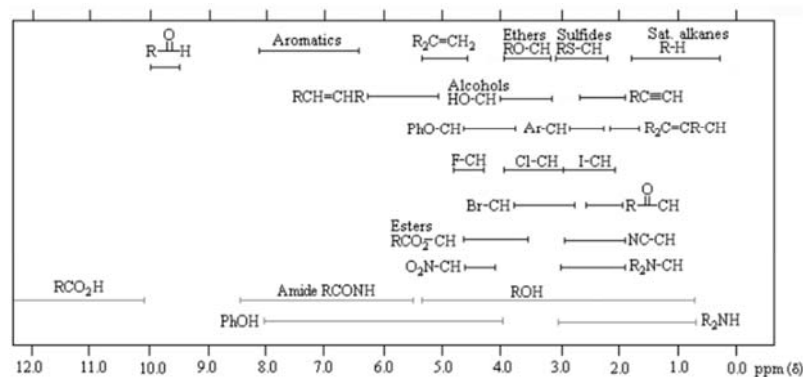


## Chemical Shift & shielding effect



[http://www.mc.vanderbilt.edu/documents/fmri/files/2013\\_Phys352A\\_MRS\(1\).pdf](http://www.mc.vanderbilt.edu/documents/fmri/files/2013_Phys352A_MRS(1).pdf)

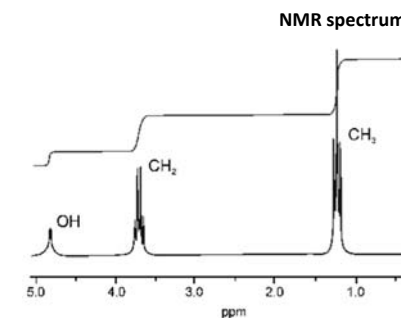
## MRS Spectrum



[http://www.mc.vanderbilt.edu/documents/fmri/files/2013\\_Phys352A\\_MRS\(1\).pdf](http://www.mc.vanderbilt.edu/documents/fmri/files/2013_Phys352A_MRS(1).pdf)

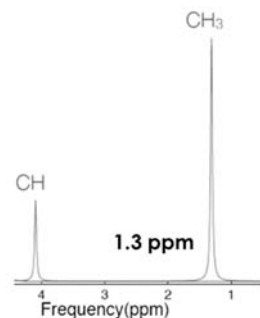
## Spectroscopy

- The area under a given peak is proportional to the number of protons contributing to the peak.
- Ex: ethanol ( $\text{CH}_3\text{—CH}_2\text{—OH}$ ) 乙醇**
  - the area under the methyl ( $\text{CH}_3$ ) peak would be 3 (in relative units);
  - the area under the methylene ( $\text{—CH}_2\text{—}$ ) peak would be 2;
  - and the area under the hydroxyl ( $\text{—OH}$ ) peak would be 1.
- MRS requires a species to be present in at least a 1 mM concentration to be seen.



## Lactate: C<sub>3</sub>H<sub>5</sub>O<sub>3</sub> 乳酸

- Structure:
 
$$\begin{array}{c} \text{OH} \\ | \\ \text{H}_3\text{C} - \text{CH} - \text{COO}^- \end{array}$$
- One methyl group (CH<sub>3</sub>)
  - 3 equivalent protons
- One methane group (CH)
- Shielding:
  - Methyl group: high
  - Methane group: low



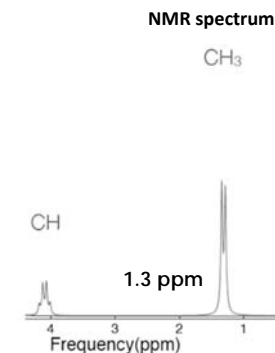
[http://www.mc.vanderbilt.edu/documents/fmri/files/2013\\_Phys352A\\_MRS\(1\).pdf](http://www.mc.vanderbilt.edu/documents/fmri/files/2013_Phys352A_MRS(1).pdf)  
<http://cflu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Lactate: C<sub>3</sub>H<sub>5</sub>O<sub>3</sub> 乳酸

- Structure:
 
$$\begin{array}{c} \text{OH} \\ | \\ \text{H}_3\text{C} - \text{CH} - \text{COO}^- \end{array}$$
- Because the methyl and methane groups share a bond, they are said to be "coupled"
  - Coupling results in peak splitting
  - Splitting or "J-coupling" makes peak identification more difficult.
  - It increases with increasing TE (J-evolution).



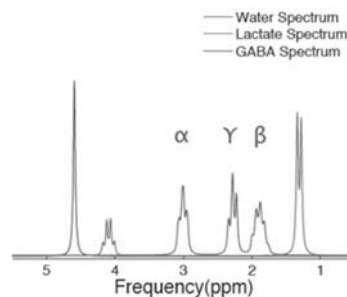
[http://www.mc.vanderbilt.edu/documents/fmri/files/2013\\_Phys352A\\_MRS\(1\).pdf](http://www.mc.vanderbilt.edu/documents/fmri/files/2013_Phys352A_MRS(1).pdf)  
<http://cflu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## γ-aminobutyric acid (GABA)

- Structure
 
$$\begin{array}{c} \gamma \quad \beta \quad \alpha \\ \text{N}^+ - \text{CH}_2 - \text{CH}_2 - \text{CH}_2 - \text{COO}^- \end{array}$$
- Three methylene groups (two equivalent protons per group)
- Coupling:
  - α is coupled to β
  - β is coupled to γ

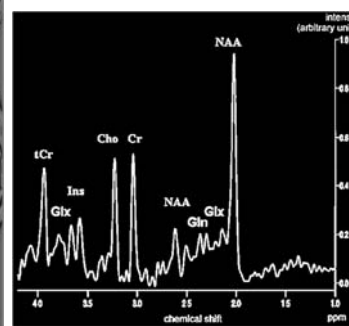


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## MRS peaks in Brain



4.0 ppm ← 1.0 ppm

Metabolite	Major Resonance (ppm)	Effect	Visible only at short TE
Lipids (Lip)	0.9, 1.3	Breakdown of tissue	Y
Lactate (Lac)	1.3	Marker of anaerobic glycolysis	N
N-acetyl aspartate (NAA)	2.0	Marker of neuronal health	N
Glutamate/Glutamine (Glx)	2.1, 3.8	Excitatory neurotransmitter	Y
Choline (Cho)	3.2	Marker of membrane metabolism, cell proliferation	N
Creatine (Cr)	3.0	Marker of cellular energetics	N
Myo-inositol (MI or Ins)	3.5, 3.6	glial cell marker	Y

All ppm are given relative to TMS (tetramethylsilane), 0 ppm.

<http://cflu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Glutamate(Glu)/Glutamine(Gln)

- Glutamate: essential excitatory neurotransmitter
  - Multiplet: 2.04~2.35 ppm
  - Doublet: 3.75 ppm
- Glutamine
  - Multiplets: 2.12~2.46 ppm
  - Triplet: 3.76 ppm
- Overlapping resonance → Glx

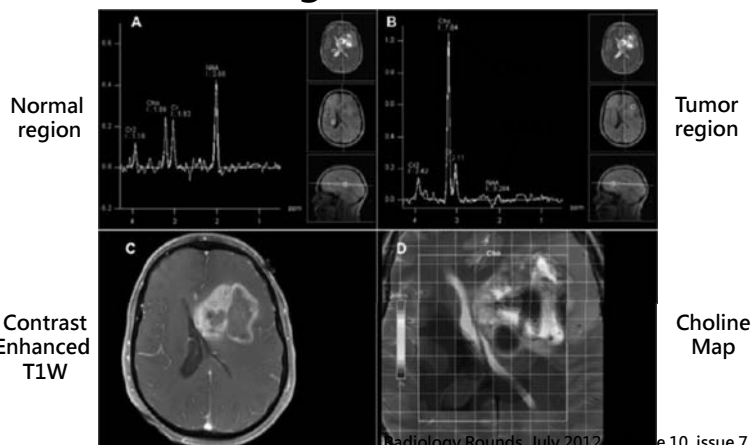
**Glutamate-glutamine cycle**  
 glutamate from neurons → astrocytes  
 → glutamine → synthesis of glutamate

## Clinical Applications of MRS

Disease	Metabolic Changes
Brain tumors	Cho ↑, NAA ↓, Cr ↓, Lac and Lip ↑
Ischemic Stroke	Lac ↑, NAA ↓, Glx ↑, Cr ↓, Cho ↓
Epilepsy	NAA ↓, Lac ↑
Multiple Sclerosis	NAA ↓, Cho ↑, (Cr ↓)
HIV/AIDS	NAA ↓, Cho ↑, MI ↑
Traumatic Brain Injury	NAA ↓, Cho ↑, Lac ↑
Hepatic Encephalopathy	Cho ↓, MI ↓, Glx ↑

Radiology Rounds, July 2012-volume 10, issue 7.

## Brain tumor: glioblastoma



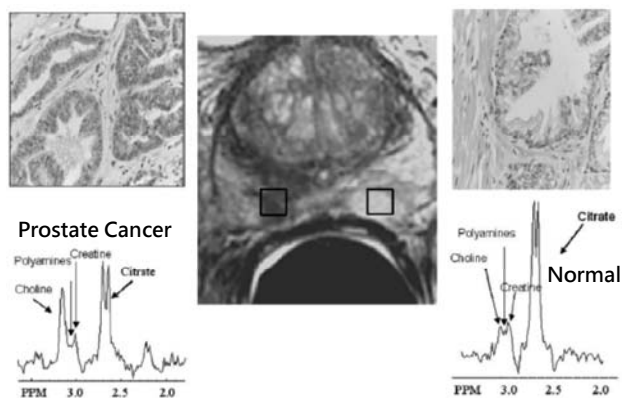
Radiology Rounds, July 2012-volume 10, issue 7.

## Clinical Applications of MRS

Neurodegenerative Disease	Metabolic Changes
Alzheimer	NAA ↓, MI ↑
Parkinson	NAA ↓ (Striatum)
Huntington	NAA ↓, Cho ↑ (Basal ganglia)
Amyotrophic Lateral Sclerosis (ALS)	NAA ↓ (Motor cortex, Brain Stem)

Radiology Rounds, July 2012-volume 10, issue 7.

## MRS for prostate



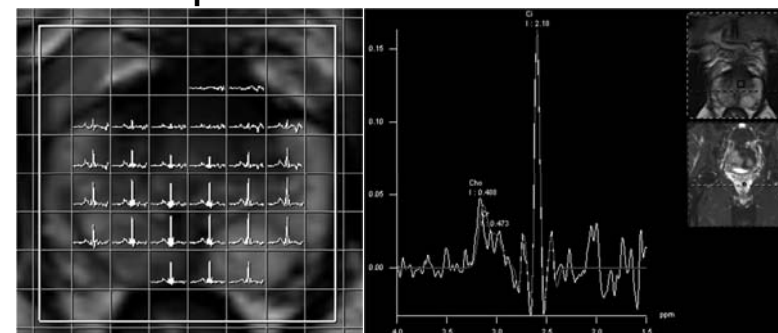
<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## MRS for prostate

- Single voxel spectroscopy (SVS)
- Multi-voxel chemical shift imaging (CSI)



<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## MRS Pulse Sequences

磁振頻譜脈衝程序與分析

<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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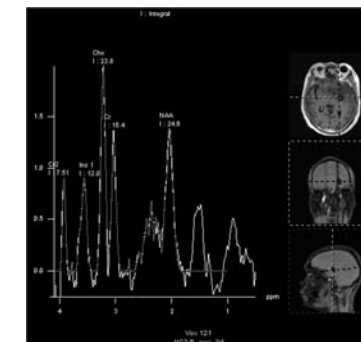
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## MRS Pulse Sequences

- Localization: Covering lesion and normal sites for the comparison.

### Two major sequences

- Point-Resolved Spectroscopy, PRESS
- Stimulated Echo Acquisition Mode, STEAM



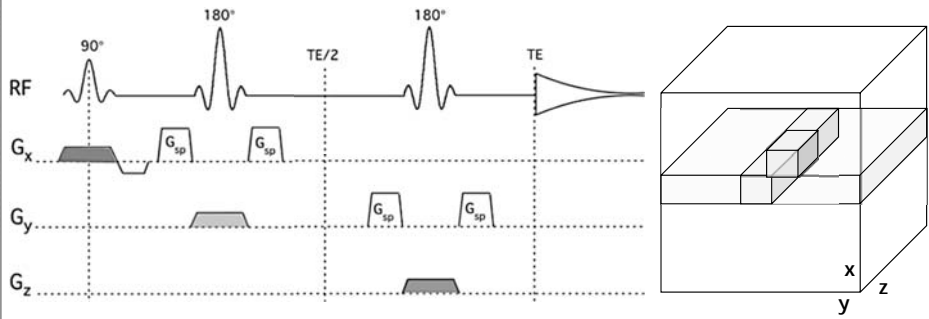
<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Point-Resolved Spectroscopy, PRESS

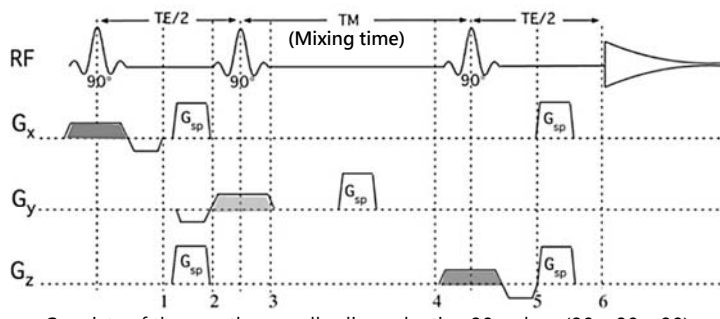


Double spin-echo sequence consisting of three slice selective pulses in orthogonal planes (90 – 180 – 180)

Signal comes from the intersection of the 3 planes!



## Stimulated Echo Acquisition Mode, STEAM

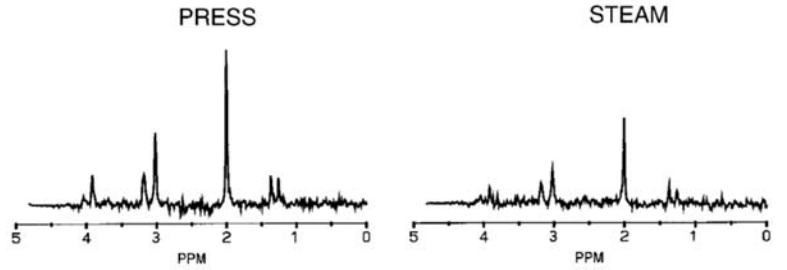


- Consists of three orthogonally slice selective 90 pulses (90 – 90 – 90)
- T2 decay does not occur during TM

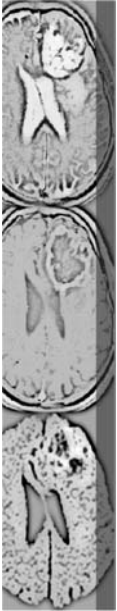


## PRESS vs. STEAM

- Stimulated echo amplitude is only half the size of a PRESS spin echo.



Medical Physics, 29(9), 2177-2197, 2002.



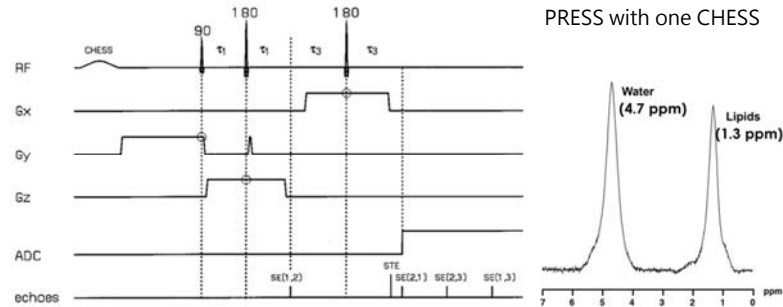
## PRESS vs. STEAM

	PRESS	STEAM	Note
SNR	S	S/2	PRESS SNR 2x STEAM SNR
TE	Short TE difficult (>30 ms)	Short TE possible (~7 ms)	STEAM: Better for metabolites with short T2
SAR	High	Low	90 transmit lower power than 180
Location	Sharp	Sharper	90 pulses have sharper profiles than 180s

[http://www.mc.vanderbilt.edu/documents/fmri/files/2013\\_Phys352A\\_MRS\(1\).pdf](http://www.mc.vanderbilt.edu/documents/fmri/files/2013_Phys352A_MRS(1).pdf)

## Water/Fat Suppression

- Chemical Shift Selection, CHESS  
Frequency-selective presaturation pulse



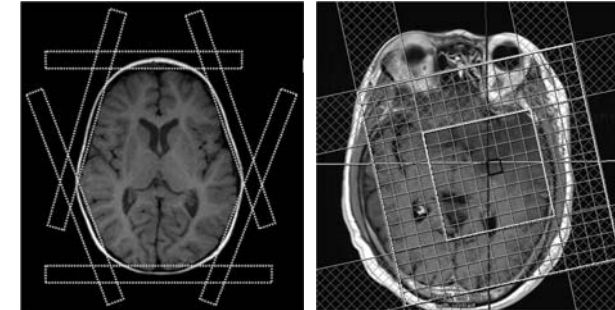
<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Fat Suppression in Brain

- Add spatial saturation bands.
- Outer Volume Suppression, OVS



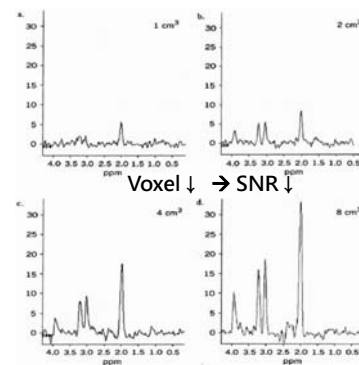
<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Voxel Size of MRS

- MRS (6~15 min) in the brain is generally performed in conjunction with MRI.
- For single voxel techniques, a volume of 8 cc ( $2 \times 2 \times 2 \text{ cm}^3$ ) is generally recommended at 1.5 T.
- Peak height is generally proportional to field strength
  - a smaller voxel can be used at 3 T, reducing partial volume averaging.



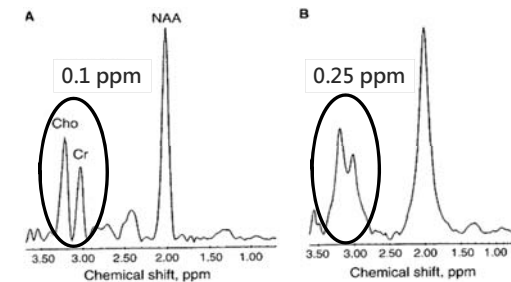
<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Shimming for MRS

- Shimming requirement for MRI is usually less than 5 ppm.
- For MRS, shimming results in improving the uniformity from 1 ppm in the main magnetic field to 0.1 ppm inside the voxel.



<http://cfliu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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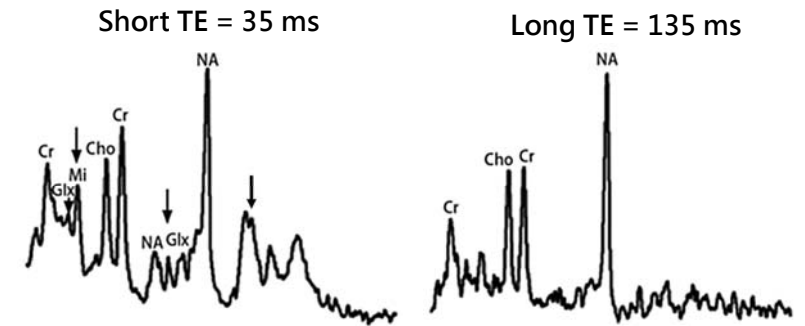
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## TR & TE in MRS

- Most institutions use a TR of 1500 msec and the shortest possible TE of 30 or 35 ms to maximize the SNR.
- This also allows the detection of short T2 species (like myo-inositol and lipid), which would otherwise have already decayed at longer TE.

Peak width is proportional to  $1/T_2$ , thus short T2 species will lead to peak broadening.

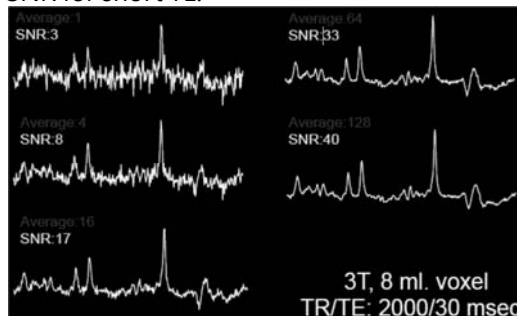
## TE Effects



Ricardo André Amorim Leite et al. Arq. Neuro-Psiquiatr. vol.68 no.1 São Paulo Feb. 2010

## Average & SNR

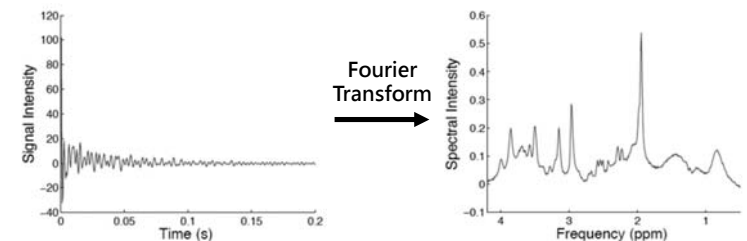
- Another option to increase SNR is to increase the average (NEX).
- Typically, 64~128 averages are demanded to acquire sufficient SNR for short TE.



Quoted from Prof. Tsai, Shang-Yueh's slide

## Fourier Transform

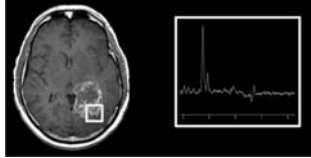
- In the simple MRS experiment, **no frequency-encoding gradients** are applied during the readout for spatial encoding.
- the signal does not contain spatial information, just information of the different resonance frequencies within the sample





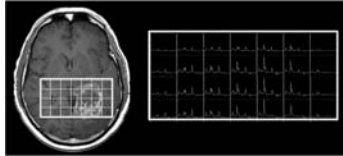
## SVS vs. CSI

### • Single-voxel spectroscopy (SVS)



- Widely used
- Fast and easy
- Limited application for large or inhomogeneous lesion

### • Multi-voxel Chemical Shift Imaging (CSI)



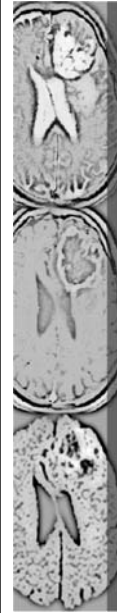
- Time consuming
- Better assessment of large or inhomogeneous lesion
- Better spatial resolution but lower SNR

<http://mriquestions.com/single-v-multi-voxel.html>

<http://cflu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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## Post-Processing

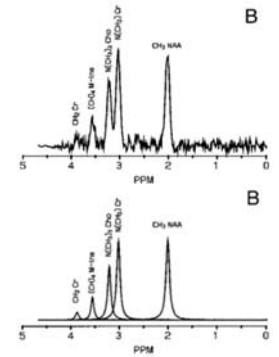
### • FID signal processing

- Water suppression (removing the 4.7 ppm signal)
- Zero filling (Increasing frequency resolution)
- Apodization (noise filtering)

### • Fourier Transform

### • Spectrum processing

- Phase correction
- Baseline correction



<http://cflu.lab.nyu.edu.tw>, Textbook: MRI The Basics, Hashemi et al.

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# THE END

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