

Development of a bimodal paraCEST- ^{19}F contrast agent for MRI

Pierre ERNOTTE

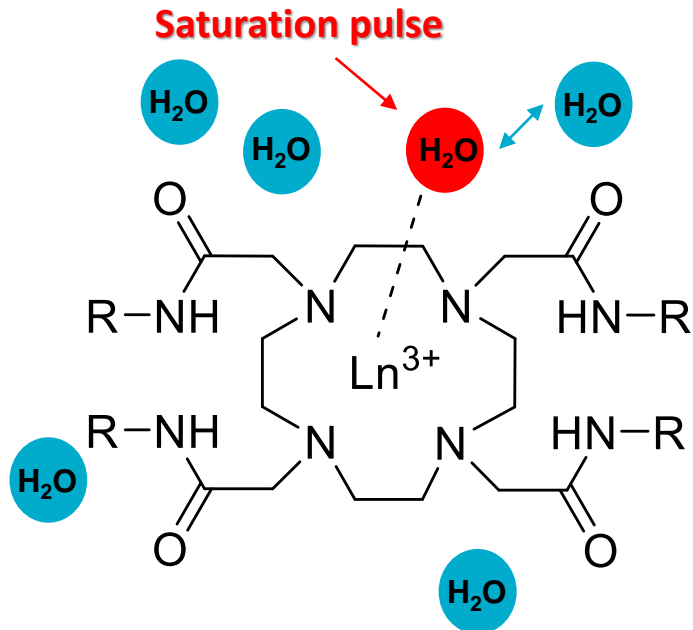
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CEST MRI

Chemical Exchange Saturation Transfer

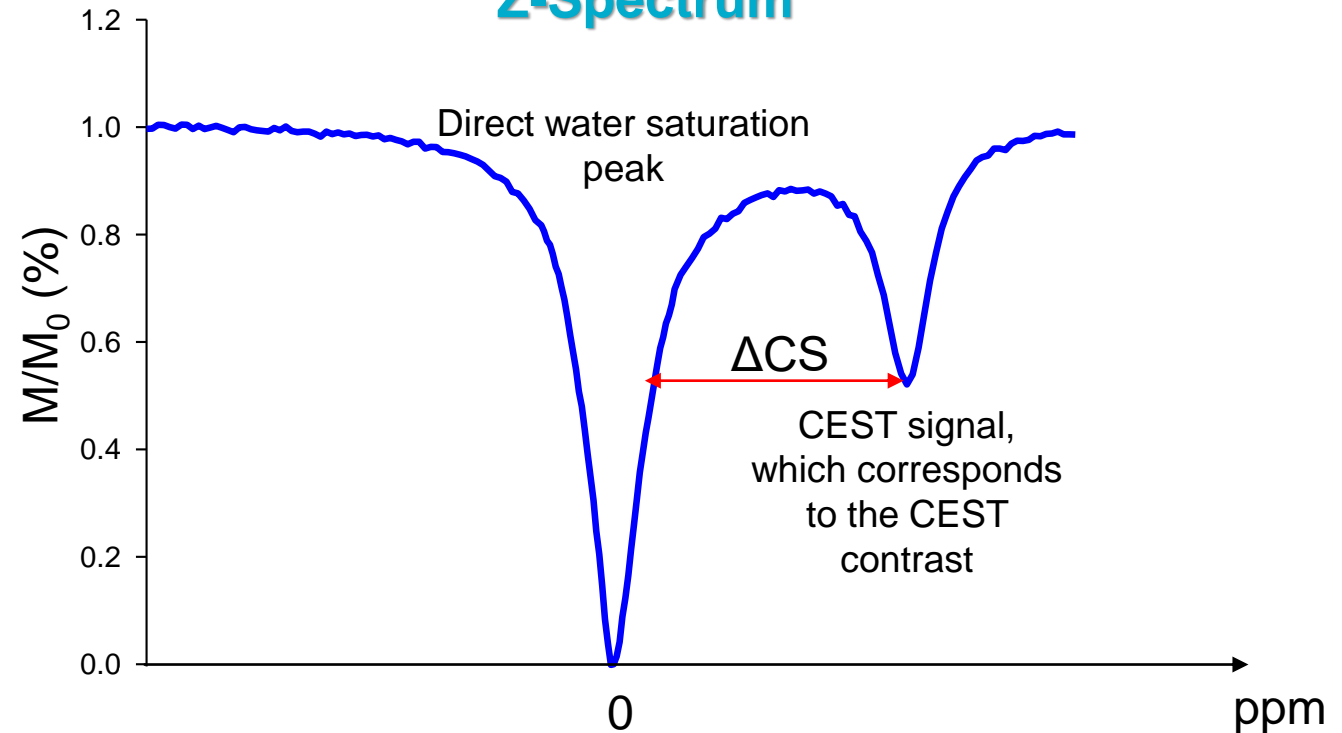


Saturation transferred from innersphere water molecules to bulk water to generate the CEST contrast

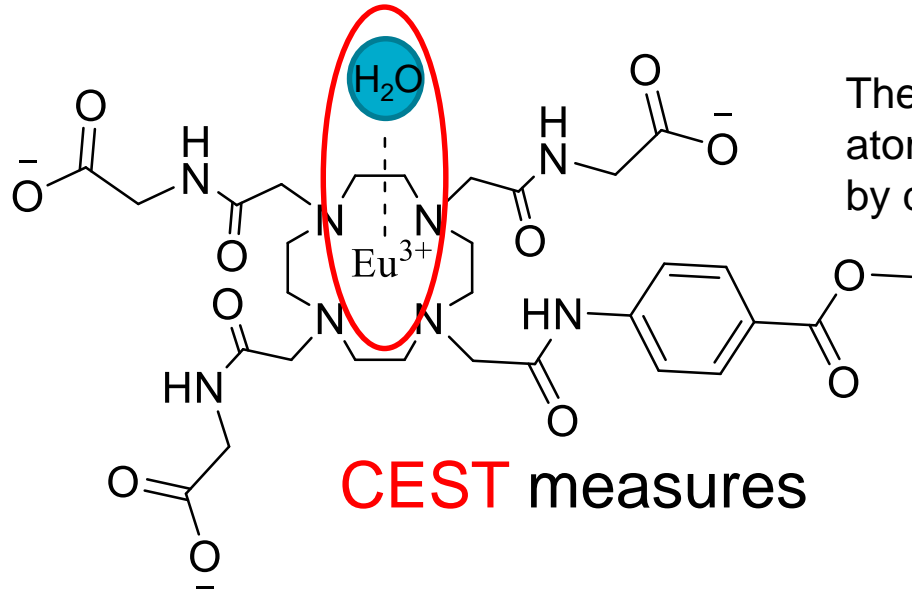
Advantages of paraCEST MRI:

- No need for a pre-contrast image
- Multi-contrast Imaging of several agent injected simultaneously
- Design of agent responsive to different physiological stimuli

Z-Spectrum

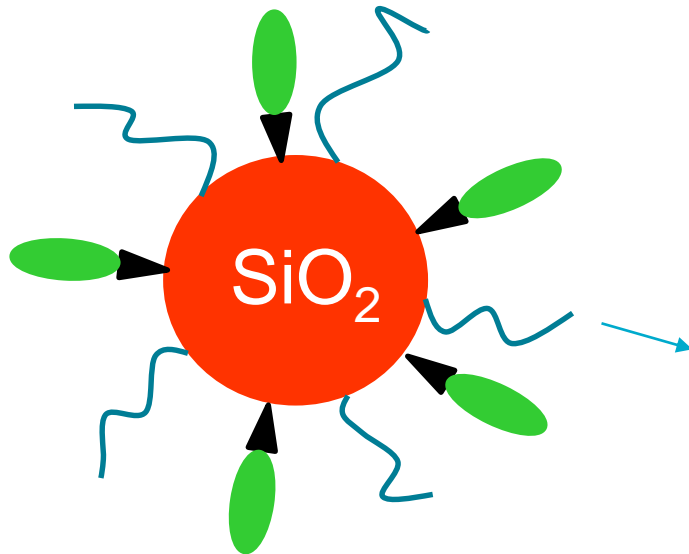
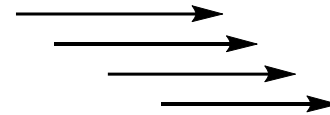


Goal of the thesis



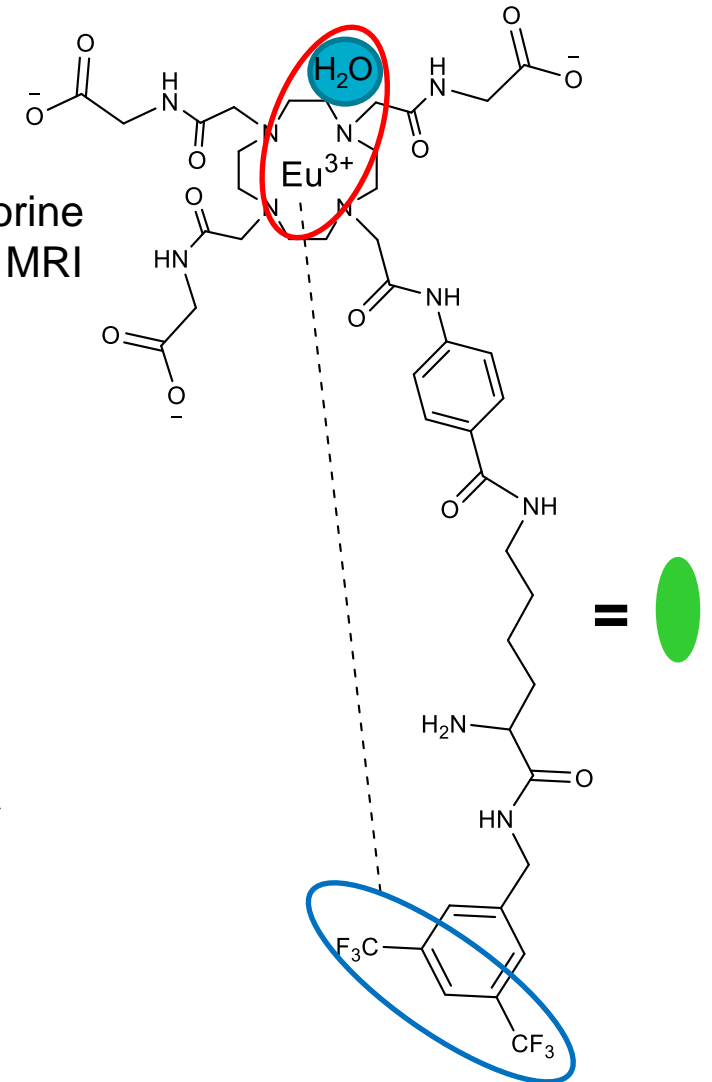
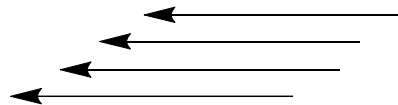
CEST measures

The interaction of a lanthanide with the fluorine atoms increases the sensitivity in fluorine MRI by decreasing the ^{19}F relaxation times



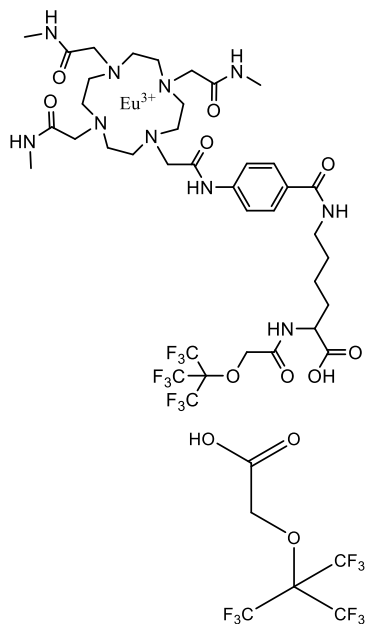
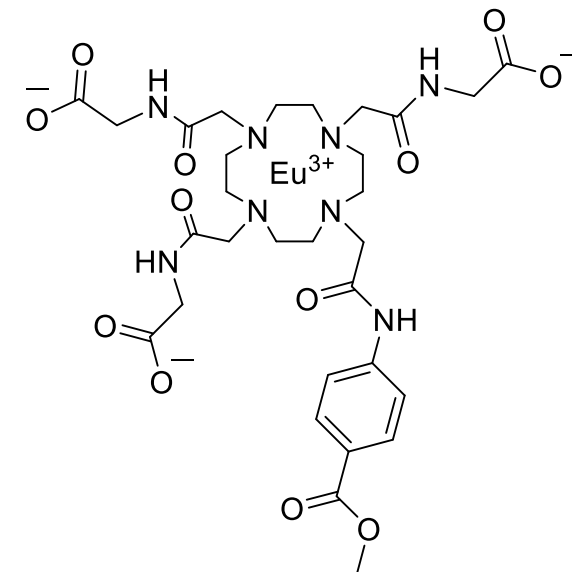
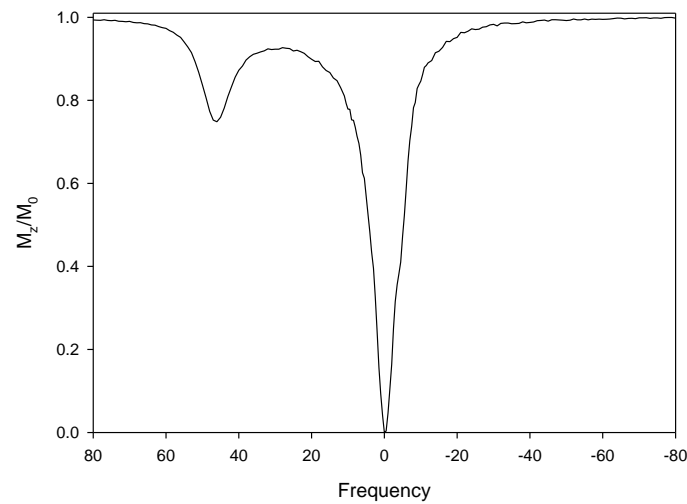
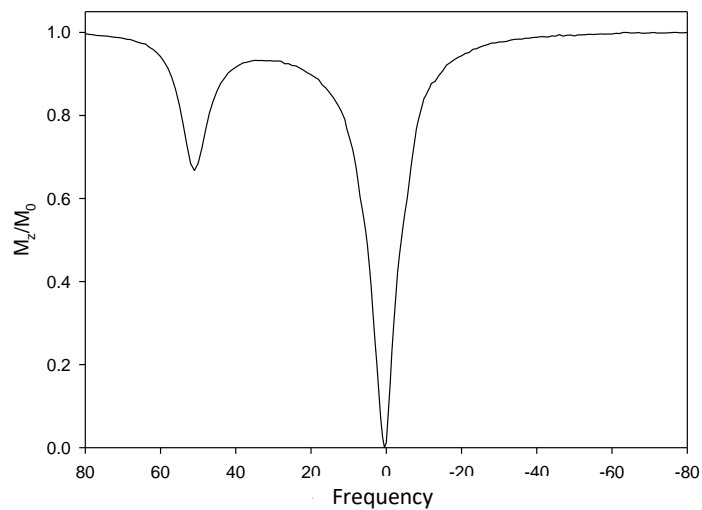
PEG

Grafting on silica nanoparticles to enhance the sensitivity



CEST and ^{19}F relaxation measures

Results



Concentration (mM)	T_1 (ms)	T_2 (ms)
0.25	889,65	463,32
0.125	884,66	473,79
		$\div 2.2$
Concentration (mM)	T_1 (ms)	T_2 (ms)
10	2021	1832
5	2064	1811
		$\div 4$

Conclusions and perspectives

Conclusions:

- Important CEST signal intensity of the presented europium complexes
- Decrease of the ^{19}F relaxation times from the bimodal agent after complexation with europium, which generates an increase of sensitivity
- High loss of water solubility after grafting of the fluorine agent

Perspectives:

- Development of other complexes to increase the water solubility
- Use of different lanthanides to optimize CEST signal and ^{19}F relaxation times
- Creation of nanoplatforms as silica nanoparticles or micelles
- Diseases targeting with a peptide