

Zinc Responsive MRI Contrast Agents

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Magnetic Resonance Imaging (MRI) has been devoted for a long time to obtaining anatomical and functional images. Recently emerging applications in molecular imaging seek information at the molecular level, looking at the biochemical or physiological abnormalities underlying the disease. Unlike anatomic imaging, molecular imaging always requires an imaging probe that is selectively responsive to the parameter to detect. Gd³⁺-based contrast agents are particularly well-adapted for this purpose and most often the changes on the efficacy (relaxivity) are based on changes of the hydration number and/or rotational dynamics of the complexes; these two parameters being the easiest to be tailored by the chemist.¹

Zinc is the second most abundant transition metal ion in humans, and it plays a central role in controlling gene transcription and metalloenzyme function. However, its quantitative distribution and its exact role are not well understood. It has also been shown that disturbances in Zn²⁺ homeostasis is implicated in neurodegenerative diseases (Alzheimer, Parkinson), diabetes, and cancers (prostate, pancreas and breast).² Therefore, monitoring Zn²⁺ *in vivo* by non-invasive technique such as MRI is important in biomedical research to understand its biological role, and to provide earlier diagnosis for specific pathologies.³

We will present the rational development of small molecular zinc responsive contrast agents based on a pyridine unit already used for Gd³⁺ complexation,⁴ to which a zinc complexing unit has been added through a linker,⁵ as well as bioinspired systems based on the zinc finger peptide.⁶ Challenges in terms of selectivity, detection field⁷ and quantification⁸ will also be discussed.

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