

# The Age and Gender Dependence of Total Transverse Relaxation Rate in Normal Human Brain at 3 T

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## Introduction

The utilizations of MRI and *in vivo* MRS at 3 T for clinical examinations and research have seen a steady increase. The transverse relaxation plays an important role in human brain MRI. Total transverse relaxation rate  $R_2^*$  ( $1/T_2^*$ ) has been shown to be useful in the measurement of tissue iron content. In this study, by using a high-resolution 3D  $R_2^*$  mapping method, multi gradient-echo slice excitation profile imaging (mGESEPI) [1], we systemically characterized anatomical distribution of the  $R_2^*$  variability with age and gender in the human brain at 3 T.

## Methods

### Human Subjects

Thirty-nine healthy normal volunteers (22 males and 17 females,  $24.5 \pm 10.8$  years of age, range 9 to 50 years) participated in the study. There was no significant age distribution difference between the two gender groups ( $p = .45$ ). Participants had no history of neurologic or psychiatric diseases. All subjects and parents of the subjects under 18 years of age gave informed written consent prior to participation.

### MRI Protocol

The mGESEPI method [1] was used to scan the brain on a Bruker MedSpec S300 3 T system with a TEM head coil for RF transmission and reception. Imaging parameters were: TR / TE / FA = 360 ms / 8 ms / 50°, bandwidth = 100 kHz, 12 echoes, 5 10-mm-thick axial slabs centered at the level of hippocampus with no gap between slabs, FOV =  $25 \times 25 \times 1 \text{ cm}^3$ , matrix =  $256 \times 192 \times 16$ .

### Data Processing and Analysis

The  $R_2^*$  maps were generated using linear regression with a home-developed software for quantitative MRI written with Interactive Data Language. The  $R_2^*$  maps from all the subjects were normalized to the Montreal Neurological Institute brain template [2] using SPM2 [3]. The resultant resolution of the  $R_2^*$  map was  $1 \times 1 \times 2.5 \text{ mm}^3$ . Both voxel-based and region of interest-based statistical analyses were performed.

## Results

The voxel-based analysis showed an evident age dependence of  $R_2^*$  in the basal ganglia, amygdala, thalamus, and red nucleus (Figure 1). Regression analyses revealed that an increase of  $R_2^*$  followed a similar logarithmic trend in most of these structures, i.e., a rapid increase of  $R_2^*$  from 9 to 20 years of age followed by a slower rate of increase from age 20 to 30 years. The rate of increase was further reduced after 30 years of age. Figure 2 shows  $R_2^*$  of the left amygdala as a function of age. No significant difference was observed between the two gender groups (Ancova,  $T < 3.35$ ) with age as a covariate. Table 1 shows the heterogeneity of  $R_2^*$  distribution in some selected brain structures of 30 to 50 years old healthy normal human. The magnitude of  $R_2^*$  change versus age varied with brain structures. No significant lateralization effect was observed (paired t-test,  $p > .25$ ).

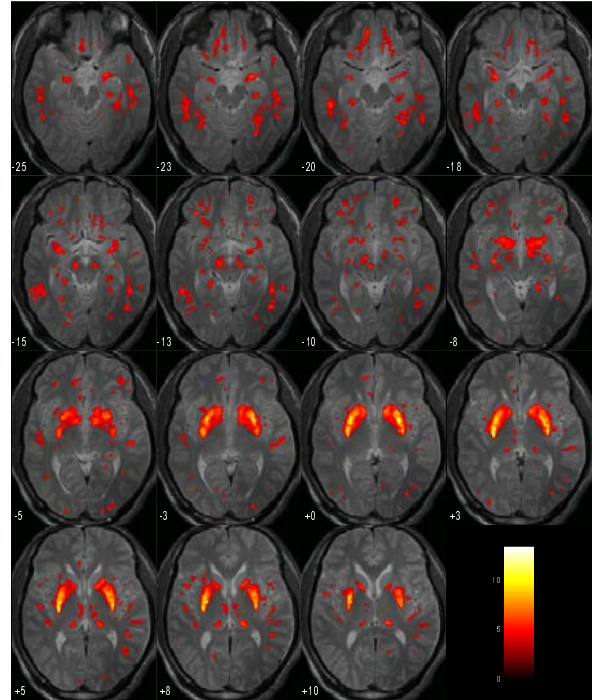
## Discussion & Conclusion

The age dependence of  $R_2^*$  in the brain exhibited a complex distribution with respect to brain anatomy. The  $R_2^*$ -age regression followed a pattern similar to the one of brain tissue iron concentration obtained by a previous postmortem study [4]. This indicates that tissue iron plays an important role in the changes of brain  $R_2^*$  during normal development and aging and, conversely,  $R_2^*$  mapping has the sensitivity in the detection of age-related iron changes in the brain *in vivo*. There was no significant gender difference on  $R_2^*$  distribution and its change with age.

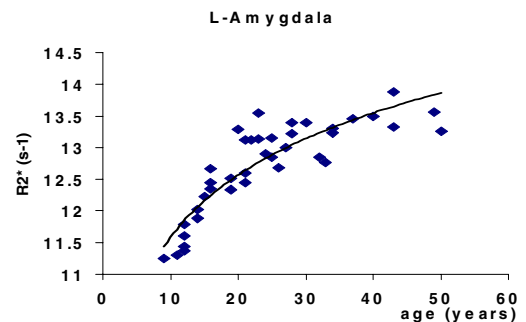
## References

1. Yang QX, et al. Magn Reson Med. 1998;39:402-409.
2. Collins DL, et al. IEEE Trans Med Imaging 1998;17:463-468.
3. Friston KJ, et al. Human Brain Mapp 1994;1:153-171.
4. Hallgren B, Sourander P. J Neurochem 1958;3:41-51.

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**Figure 1.** The anatomic structures showing positive linear correlations between  $R_2^*$  and age (simple regression, uncorrected,  $p < .001$ , extended threshold = 6). The brighter the color, the higher the correlation coefficient.



**Figure 2.** The age dependence of  $R_2^*$  at left amygdala from 39 healthy normal subjects ( $R^2 = .79$ ).

	Amygdala	Putamen	Ant Thalamus	Pos Thalamus
Left	13.32/.31	14.28/.51	12.42/.39	12.75/.58
Right	13.30/.34	14.24/.49	12.43/.32	12.89/.54

**Table 1.**  $R_2^*$  (mean/std,  $s^{-1}$ ) of selected brain structures (age 30 to 50 years). Ant, anterior; Pos, posterior.