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### Whole brain quantitative T2 MRI across multiple scanners with dual echo FSE: Applications to AD, MCI, and normal aging $\stackrel{ ightarrow}{ ightarrow}$

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### ABSTRACT

The ability to pool data from multiple MRI scanners is becoming increasingly important with the influx in 21 multi-site research studies. Fast spin echo (FSE) dual spin echo sequences are often chosen for such studies 22 based principally on their short acquisition time and the clinically useful contrasts they provide for assessing 23 gross pathology. The practicality of measuring FSE-T2 relaxation properties has rarely been assessed. Here, 24 FSE-T2 relaxation properties are examined across the three main scanner vendors (General Electric (GE), 25 Philips, and Siemens). The American College of Radiology (ACR) phantom was scanned on four 1.5 T 26 platforms (two GE, one Philips, and one Siemens) to determine if the dual echo pulse sequence is susceptible 27 to vendor-based variance. In addition, data from 85 subjects spanning the spectrum of normal aging, mild 28 cognitive impairment (MCI), and Alzheimer's disease (AD) was obtained from the Alzheimer's Disease 29 Neuroimaging Initiative (ADNI) to affirm the presence of any phantom based between vendor variance and 30 determine the relationship between this variance and disease. FSE-T2 relaxation properties, including peak 31 FSE-T2 and histogram width, were calculated for each phantom and human subject. Direct correspondence 32 was found between the phantom and human subject data. Peak FSE-T2 of Siemens scanners was consistently 33 at least 20 ms prolonged compared to GE and Philips. Siemens scanners showed broader FSE-T2 histograms 34 than the other scanners. Greater variance was observed across GE scanners than either Philips or Siemens. 35 FSE-T2 differences were much greater with scanner vendor than between diagnostic groups, as no 36 significant changes in peak FSE-T2 or histogram width between normal aged, MCI, and AD subject groups 37 were observed. These results indicate that whole brain histogram measures are not sensitive enough to 38 detect FSE-T2 changes between normal aging, MCI, and AD and that FSE-T2 is highly variable across scanner 39 vendors. 40

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Alzheimer's disease (AD) is a progressive neurodegenerative 45 46 4748

disorder characterized by an insidious onset followed by gradual decline in cognitive function. It is the most common form of dementia and is quickly becoming a global crisis, affecting approximately 10% of individuals over age 65 and nearly 50% of individuals over 85 years of age (Evans et al., 1989). An additional 19% of individuals over age

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65, and 29% over age 85, are estimated to have mild cognitive 51 impairment (MCI) (Lopez et al., 2003). MCI represents the transi- 52 Q8 tional phase between normal aging and probable AD, whereby 53 individuals have diminished memory function, yet maintain normal 54 levels of daily activity and are not demented (Petersen et al., 2001). 55 09 Individuals with the amnestic form of MCI are more likely to develop 56 AD than their normal counterparts; those with amnestic MCI show 57 rates of conversion around 12% per year, whereas normal elderly 58 tend to convert to AD at 1-2% a year (Petersen et al., 1999). With the 59 Q10 development of potential therapeutic interventions it is becoming 60 important to identify potential biomarkers of disease presence as 61 early as possible. 62

Tissue relaxation properties as measured with quantitative T2 MRI 63 have the potential be a valuable resource in early identification of 64 individuals with MCI and AD. T2 is a function of tissue free water 65 properties and the local environment of the nuclei, allowing one to 66

simes Data used in the preparation for this article were obtained from the Alzheimer's Disease Neuroimaging Initiative (ADNI) database (http://www.loni.ucla.edu/ADNI). As such, the investigators within the ADNI contributed to the design and implementation of ADNI and/or provided data, but did not participate in analysis or writing of this manuscript. A complete listing of ADNI investigators can be found at http://www.loni. ucla.edu/ADNI/Collaboration/ADNI\_Citation.shtml.

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examine tissue state and hydration (Jack, 1996). The neuropathology 67 68 of MCI and AD both involve the deposition of neurofibrillary tangles and amyloid plaques. These entities result in neuronal loss and may 69 70 affect the relaxation properties of the surrounding tissue, which in turn may alter the T2 times. Histological studies suggest that iron and 71 water content increase in AD brains, resulting in shortened and 72 prolonged T2 values, respectively (Schenck et al., 2006), as T2 is 73 74sensitive to iron levels in brain tissue (Vymazal et al., 1996; Bartzokis 75et al., 1997). Because quantitative T2 values reflect tissue character-76istics, it may be useful to study neuronal degradation in AD prior to 77 other detectable atrophy.

Most previous research on T2 relaxation in AD has been 78 conducted at a single research center using data collected on a 79 single scanner. Given the current trend for conducting multi-site 80 studies and for individual sites to have multiple scanners, often 81 from different vendors, it is beneficial to examine T2 relaxation 82 properties in a multi-vendor research study. Multi-site investiga-83 tions enable a larger recruitment population than at a single site; 84 however, studies using imaging information obtained at multiple 85 sites have the added difficulty of ensuring consistent results 86 between sites. Of importance to this study are potential differences 87 in the fast (a.k.a. turbo) spin echo (FSE) readouts used with the 2D-FSE 88 89 pulse sequence. It has been shown using voxel-based morphometry (VBM) that the effect of disease on regional brain volume may be greater 90 than the effect of scanner variation in a population of AD and normal 91aged subjects across six scanners (Stonnington et al., 2008). The current 92study will examine if the same holds true for FSE-T2 relaxation. It should 93 94be noted that there are a number of ways to measure T2 and the results of each method are likely to be dependent upon the sequence used. For 95the purposes of clarity in this manuscript we will be referring to T2 as 96 97 FSE-T2 in order to reflect the methodology/sequence that was used to 98 measure it.

99 Prior studies on scanner variability have focused on volumetric measures, primarily derived from T1-weighted pulse sequences. 100 Image uniformity, geometry (Ihalainen et al., 2004; Fu et al., 2006), 101 and signal to noise ratio (Fu et al., 2006) have been shown to vary 102both within and between vendors and platforms; but, no studies 103 104 have examined quantitative T2 using fast spin echo sequences across multiple vendors or platforms. Since no sequences are 105standardized between vendors, it is unknown if the acquired FSE-T2 106 values are similar or different, and to what degree they may be 107 108 different. One relaxometry study that was conducted across three research sites using GE and Siemens platforms has shown minimal 109 variance due to a scanner using the DESPOT2 (driven equilibrium 110 single pulse observation of T2) pulse sequence (Deoni et al., 2008). 111 In the present study we tested if the dual echo FSE pulse sequence 112 113 showed similar accuracy between scanner vendors using both the ACR phantom and human subjects. It should be noted that specific 114 pulse sequence parameters are often slightly different between 115scanner vendors and platforms in multi-site imaging studies in 116 order to produce images with similar image quality and appearance. 117 118 It is unclear how these pulse sequence variations between vendors 119 affect FSE-T2 values. This paper seeks to quantify these potential discrepancies. 120

The specific aim of the current study was to assess the FSE-T2 121values obtained from MRI scans of phantoms and the human brain 122123on either GE, Philips, or Siemens scanners. We used ACR phantom scans from four MRI scanners and human MRI data acquired from 124the Alzheimer's Disease Neuroimaging Initiative (ADNI) database, 125 incorporating multiple platforms from three manufacturers. The 126 human subjects fell into the categories of normal aging (NL), MCI or 127 AD groups. In this context, the purposes of this work were: first, to 128 study possible vendor-dependent systematic differences in FSE-T2 129in the ACR phantom and human subjects, and second, to explore 130possible FSE-T2 histogram signatures of normal aging, mild 131 132 cognitive impairment, and Alzheimer's disease.

### Materials and methods

### **Subjects**

### ACR phantom

The American College of Radiology (ACR) magnetic resonance 136 accreditation phantom was used to test for scanner variability. The 137 ACR MR phantom is designed to test a number of parameters, 138 including geometric distortion, spatial resolution, slice thickness and 139 position, interslice gap, estimation of image bandwidth, low contrast 140 detectability, image uniformity, signal to noise ratio, slice offset, and 141 landmark. For more detailed information on this phantom, refer to 142 http://www.aamp.org/meetings/99AM/pdf/2728-58500.pdf. The ACR 143 phantom was used because it is widely available and already part of 144 the certification system for scanners. It should be noted that the ADNI 145 phantom is also used each time a subject is scanned in the ADNI study, 146 but it is only scanned with the MP-RAGE sequence and there are no 147 data available with this phantom using the FSE sequence. We asked 148 the specific sites listed in the Acknowledgments to run the ACR 149 phantom for us as an additional scan. 150

### ADNI subjects

Data from 85 subjects (age range = 60–91, average age = 75.47, 44 152 females, 41 males,  $N_{\text{NL}} = 32 N_{\text{MCI}} = 26$ ,  $N_{\text{AD}} = 27$ ) across three vendors 153 (27 Philips, 29 GE, 29 Siemens) was selected from the ADNI database 154 (http://www.loni.ucla.edu/ADNI). MMSE scores for each subject were 155 obtained (MMSE<sub>NC</sub>=24-30, MMSE<sub>MCI</sub>=17-29, MMSE<sub>AD</sub>=18-29) 156 (Table 1). Data from 18 ADNI study sites across Canada and the United 157 States were chosen at random for the current study. Multiple scanner 158 brands from each vendor were accepted in the study: GE: Signa Excite, 159 Signa HDx; Philips: Achieva, Intera, Gyroscan Intera, Intera Achieva; 160 Siemens: Sonata, Symphony. 161

All participants in the ADNI underwent a battery of neuropsycho- 162 logical tests, including the MMSE (Folstein et al., 1975), the CDR-Sum 163 of Boxes (Morris, 1993), and the Global dementia scale (GD-scale) 164 (Auer and Reisberg, 1997; Reisberg et al., 1988). Subjects were 165 clinically assessed for cognitive status and classified as: (a) normal 166 controls with normal cognition and memory, CDR 0, and MMSE 167 between 24 and 30; (b) amnestic MCI with memory complaint 168 verified by a study partner, memory loss measured by education- 169 adjusted performance on the Logical Memory II subscale of the 170 Wechsler Memory Scale-Revised (Wechsler, 1987), preserved activ- 171 ities of daily living, CDR 0.5, MMSE between 24 and 30, and absence of 172 dementia at time of baseline MRI scan; or (c) probable AD with 173 memory complaint validated by an informant, abnormal memory 174 function for age and education level, absence of depression, impaired 175 activities of daily living, diminished cognition, CDR>0.5, and MMSE 176 between 20 and 26. 177

### Alzheimer's Disease Neuroimaging Initiative

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The ADNI is a 5-year non-randomized natural history non- 179 treatment study utilizing data from multiple study centers across 180 the United States and Canada. The primary aim of the ADNI is to 181 examine if serial MRI, PET, biological markers, and clinical and 182 neuropsychological assessments can be combined to analyze the 183 progression of MCI to early AD. The ADNI is a public-private 184

#### Table 1 t1.1 Subject gender breakdown by vendor and diagnostic group (female/male). t1.2

Subject enrollment table					
	Philips	GE	Siemens	Total	t1.4
Normal	6/3	5/13	4/4	15/20	t1.5
MCI	2/5	7/6	7/4	16/15	t1.6
AD	4/7	6/4	6/4	16/15	t1.7
Total	12/15	18/23	17/12	47/50	t1.8

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### t2.1 Table 2

ACR phantom results from 4 scanners. Siemens shows approximately 30 ms prolonged peak T2 compared to GE and Philips. GE showed 10 ms variance between scanners.

t2.2 t2.3	ACR phantom peak T2 and histogram width							
t2.4		GE HDx	GE Signa Excite	Philips Intera	Siemens Avanto			
t2.5	Peak T2 (ms)	145	135	140	170			
t2.6	T2 histogram width	50	50	50	55			

partnership, launched in 2003, funded by the National Institute on
 Aging, the National Institute of Biomedical Imaging and Bioengi neering, the Food and Drug Administration, private pharmaceutical
 companies, and non-profit organizations.

One of the main goals of the ADNI is to develop optimized methods and uniform standards for the acquisition of multicenter MRI and PET data on normal control subjects and patients with AD and MCI. For more information, refer to http://www.adni-info.org.

193 Image acquisition

For the phantom scans, MRI scanning was performed on four 1.5 T scanners: GE Signa Excite, GE HDx, Philips Intera, and Siemens Avanto. For the ADNI subjects, 1.5 T scanners from General Electric (GE), Philips Medical Systems, and Siemens Medical Solutions were used for examination of tissue from below the base of the cerebellum through the top of the head.

For both the ACR phantom and study participants, the dual fast/ 200 turbo spin echo pulse sequence used was acquired in the straight axial 201 202 plane with the following parameters: effective echo time  $(TE1_{eff}) =$  $10.04-13 \text{ ms}, \text{TE2}_{\text{eff}} = 95.22-103 \text{ ms}, \text{ repetition time (TR)} = 3000 \text{ ms},$ 203 echo train length (ETL) (turbo factor) = 7-16 (GE = 16, Philips = 10, 204Siemens =  $7^*$  one Siemens subject had an ETL of 13), echo 205206 spacing = 12-12.7 ms, slice thickness = 3.0 mm, slice gap = 0 mm, 207pixel spacing = 0.9375 mm, matrix size =  $228-256 \times 256$ . Effective TE 208was consistent across Philips platforms, but for both GE and Siemens it varied within vendor. Matching parameters between platforms do not 209produce the same image quality, thus small variations in pulse 210 sequence parameters are often incorporated in large-scale studies. 211 More specific parameter values for each research site can be found at 212 http://www.loni.ucla.edu/ADNI/Research/Cores. For simplicity, the 213 pulse sequence will be referred to throughout as FSE, although both 214

Q14 215 Philips and Siemens call it turbo spin echo (Table 2).

216 Data processing and segmentation

Images from the two echo times, for both phantoms and human subjects, were separated using EFilm (Merge Healthcare, Milwaukee, WI), providing two separate image stacks, one from TE1, the other from TE2. Header information was separated from the images using Image J (http://www.rsbweb.nih.gov/ij/). Refer to Fig. 1 for a flow diagram of the post-processing steps between the directly-acquired images and the histograms.

The datasets were then analyzed with an in-house computer program using MathCAD 2001i (PTC, Needham, MA) software to generate quantitative-MRI maps (Jara et al., 2006; Suzuki et al., 2006; Jensen et al., 2001). FSE-T2 quantitative maps were generated on a pixel-by-pixel basis from the two T2-weighted dual echo FSE datasets, 228 according to the mono-exponential function:  $T2 = (TE_2 - TE_1)/ln(S_1/S_2)$ , 229 where  $S_1$  and  $S_2$  were the signals obtained at  $TE_1$  and  $TE_2$ , respectively. 230 Fig. 2 shows typical FSE-T2 quantitative maps for NL subjects, MCl 231 subjects, and patients with probable AD. 232

The brain was segmented using a dual-clustering segmentation 233 algorithm in an in-house MathCAD program (Suzuki et al., 2006; Jara 234 et al., 2006; Jensen et al., 2001). The scans were individually analyzed 235 to obtain the best segmentation of the whole brain and eliminate the 236 inclusion of fat and extra-cranial matter (Fig. 3). Overall the program 237 was able to segment brain tissue from each vendor with no 238 appreciable differences in ease. 239

Histograms of FSE-T2 relaxation times were generated from the 240 FSE-T2 maps using MathCAD (Fig. 4). The histogram shows an 241 approximately monomodal curve with an asymmetrical tail repre-242 senting meninges and extra-ventricular CSF. The main peak repre-243 sents the T2 relaxation time of both gray and white matter (MacKay 244 et al., 2006; Jara et al., 2006). 245

Statistics

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Phantom data were visually analyzed for differences in peak FSE-T2 247 and histogram width. Statistical analysis was performed on the human 248 data using Excel, Datadesk version 6.1, and SPSS 13. Differences in 249 gender composition between groups were assessed using  $\chi^2$  test. 250 Multiple analysis of variance (MANOVA) was used to examine 251 differences between diagnostic groups and platform vendor on peak 252 FSE-T2, FSE-T2 full width at half maximum (T2-width), and volumes for 253 each segment. Scheffe post-hoc analysis was performed on the data. The 254 overall interaction between scanner, diagnosis, and dependent variables 255 was examined with MANOVA with further pairwise comparisons using 256 Scheffe post-hoc analysis. *F*-tests were used to examine variance 257 between scanner vendors for peak FSE-T2 and histogram width. 258 ANOVA with Scheffe post-hoc analysis was performed within scanner 259 vendor to test for differences between NL, MCI, and AD. 260

Partial correlation analysis was performed to examine relation- 261 ships between peak FSE-T2, FSE-T2 histogram width, and neuropsy- 262 chological test scores while controlling for scanner vendor. 263

### Results

### ACR phantom

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Phantom histograms show differences in peak FSE-T2 between 266 manufacturers ( $GE_{HDx} = 145$  ms,  $GE_{Signa Excite} = 135$  ms, Philips<sub>Intera</sub> = 267 140 ms, Siemens<sub>Avanto</sub> = 170 ms). Histograms of the Siemens scan show 268 approximately 30 ms prolonged peak FSE-T2 compared to the GE and 269 Philips counterparts. Histograms from GE scans show the same average 270 value as the Philips histograms (140 ms), but an approximately 10 ms 271 difference was noted between the 2 GE scanners. Also, Siemens 272 histograms were broader by 5 ms as compared to either GE or Philips 273 (histogram width,  $GE_{HDx} = 50$  ms,  $GE_{Signa Excite} = 50$  ms, Philips<sub>Intera</sub> = 274 50 ms, Siemens<sub>Avanto</sub> = 55 ms). 275



Fig. 1. Flow chart showing the post-processing data analysis used to obtain histograms from the directly-acquired images.

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Fig. 2. Representative T2 quantitative MR maps. a) T2 map of a 79 year old male control subject on a GE Signa Excite platform. b) T2 map of a 77 year old female subject with MCI scanned on a Siemens Sonata platform. c) Shows a T2 map from a 75 year old male AD subject scanned with a Siemens Avanto platform.

### 276 ADNI subjects

Age and gender were not significantly different amongst diagnostic groups or scanner vendors (p>0.05). Average histograms were created according to vendor and subject population (Fig. 5).

### 280 Peak FSE-T2

Overall analysis showed a significant difference between scanner vendors (F 2, 73 = 146.37, p<0.0001). Follow up comparisons revealed Siemens peak FSE-T2 was 15–29 ms (p<0.0001) longer than both GE and Philips across the three diagnostic groups (Fig. 5). GE peak values were on average 4 ms prolonged over Philips values; however, this did not reach significance possibly due to the larger degree of variance seen across GE subjects.

288 Within GE platforms no significant differences were found for peak 289FSE-T2. The directionality of change in GE indicated a decrease in FSE-T2 290from NL to MCI, an increase in FSE-T2 from MCI to AD, and an increase in FSE-T2 from NL to AD. Within Philips platforms, no significant 291differences were found for peak FSE-T2. The directionality of change 292 in Philips shows an increase in peak FSE-T2 from NL to MCI, a decrease in 293 peak FSE-T2 from MCI to AD, and a decrease in peak FSE-T2 from NL to 294 AD. Within Siemens platforms a significant difference was found for 295 peak FSE-T2 between NL and MCI (p > 0.05), but not between NL and AD 296subjects. The directionality of peak FSE-T2 change in Siemens indicates a 297decrease from NL to MCI, an increase from MCI to AD, and a decrease 298299from NL to AD.

Tests of variance indicate that Philips ( $s^2 = 7.764$ ) scanners show significantly lower variance than either GE ( $s^2 = 38.239$ ) (p < 0.001) or Siemens ( $s^2 = 30.988$ ) (p < 0.001) for peak FSE-T2. GE and Siemens variance for peak FSE-T2 was not significantly different (p > 0.05).

### 304 FSE-T2 histogram width

The width of the histogram reflects the homogeneity of tissue composition. Width was measured at half maximum for each FSE-T2





histogram. Overall analysis showed significant scanner differences (*F* 2, 307 73 = 9.483, *p*<0.0001). ANOVA revealed a significant difference 308 between Siemens and both Philips and GE (*p*<0.005). 309

### Correlations

Peak FSE-T2 and FSE-T2 width were assessed for correlations with 311 age, CDR, MMSE, and GD scale scores. Peak FSE-T2 was found to 312 significantly correlate with age (r = 0.370, p < 0.01). FSE-T2 histogram 313 width was significantly correlated with GD scale (r = 0.232, p < 0.05). 314

### Discussion

The specific goals of this work were to identify potential vendor- 316 dependent systematic differences in quantitative FSE-T2 maps of the 317 ACR phantom and human brain and to study FSE-T2 histogram 318 properties across the spectrum of normal aging, MCI, and AD. 319 Significant overall differences were found between scanner vendors 320 across the FSE-T2 histogram-derived parameters in both phantom 321 and human studies. Follow up analysis showed that Siemens had 322 higher FSE-T2 peak values and broader histograms than GE and 323 Philips. Measurements were not statistically significant between 324 diagnostic groups when accounting for scanner vendor, which 325 decreased the effective sample size per group to between 7 and 18 326 subjects. The small sample size within vendors may partially account 327 for there not being a significant difference between peak FSE-T2 of AD, 328 MCI, and normal aging subjects. 329

The trends between NL, MCl, and AD, suggest a scanner vendor- 330 disease interaction effect, such that the trend for FSE-T2 between 331 normal aging, MCl, and AD was inconsistent between vendors (i.e. 332 normal aging from one vendor produced prolonged T2 compared to 333 AD, while in another vendor we observed the opposite trend). These 334 interactions are of greatest concern and will need to be verified with a 335 larger sample. If true, this suggests that combining data from different 336 vendors in one analysis, even when using co-factors, will end up 337 masking the underlying effect. 338

Peak FSE-T2 was shown to correlate with age, consistent with the 339 results of a previously published study (Laakso et al., 1996). 340 Histogram width correlated with GD scale. Histogram width reflects 341 water environment inhomogeneity (Whittall et al., 1997), indicating 342 that brain tissue becomes more heterogeneous as the severity of 343 dementia measured by the GD scale increases. 344

Variability between sites indicates that there was a larger degree of 345 variance between GE and Siemens sites. The variability between sites 346 using GE scanners was also observed in the phantom scans. Although 347 GE was the only vendor for which the phantom was scanned on more 348 than one platform. Additional phantom scans on Philips and Siemens 349 platforms would be useful to help confirm the degree of variance 350 between scanners. 351

Peak FSE-T2, for both phantom and human subjects, was at least 352 20–30 ms prolonged with Siemens' histograms compared to GE and 353

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Philips. The FSE pulse sequences used in the ADNI were not
standardized between research sites, consistent with protocols of
many other large-scale studies including the Framingham Heart Study
(DeCarli et al., 2005), MIRAGE (Cuenco et al., 2008), and many clinical
drug trials.

This study poses a few limitations related to pulse sequence 359 parameters and scanner hardware and software. Because the 360 sequences were not completely standardized between all platforms, 361 some scans were acquired with different effective TE or ETL. The signal 362 intensity with T2 is primarily controlled by echo time. Effective TE 363 may differ between vendors based on the k-space acquisition scheme, 364 which may have induced some of the observed scanner-related 365 variance. Much of the inaccuracy of FSE-T2 is due to stimulated echo, 366 which is affected by both ETL (turbo factor) and echo spacing. The 367 368 observed difference in FSE-T2 may be inherent to the vendor-specific scheme used to acquire k-lines with the fast spin echo readout, slice 369 profile, and phase encoding order. Other factors, such as the coils, B0 370 and B1 inhomogeneities (Majumdar et al., 1986b; Poon and Henkel- 371 man, 1992), RF pulse imperfections (Majumdar et al., 1986a), and 372 temperature variations, could also contribute to the scanner-related 373 variance. 374

Whole brain histogram-derived FSE-T2 measures may not be 375 sensitive enough to detect AD-related changes; however, T2 has been 376 shown to regionally differ in AD and MCI compared to normal aging 377 (Englund et al., 1987; Kirsch et al., 1992; Laakso et al., 1996; Pitkanen 378 et al., 1996; Parsey and Krishnan, 1998; Wang et al., 2004; Schenck 379 et al., 2006; Arfanakis et al., 2007). A potential area for future research 380 is to examine T2 relaxation times using voxel-based relaxometry 381 (VBR), which has been used to show T2 changes in autism (Hendry 382 et al., 2006), epilepsy (Pell et al., 2004; Pell et al., 2008) and multiple 383

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**Fig. 5.** Quantitative ICM T2 histogram averaged within scanner vendor: Philips, GE, and Siemens. The slight bimodal nature of the GE curve represents those subjects from site 005, which had average values approximately 20 ms lower than other GE sites. a) Normal control subjects: Philips and GE peak at 97.35 and 98.37 while Siemens peaks at 119.69 ms. The width of the spectrum is approximately 50.6 for Philips, GE, and Siemens respectively. b) MCI subjects: Philips peak T2 = 94.29 ms, GE peak = 99.38 ms, while Siemens peaks at 119.09 ms. The width of the spectrum is 52, 53, and 55 U for Philips, GE, and Siemens respectively. c) AD subjects: Philips peak = 95.68 ms, GE peak = 102.08, and Siemens peak = 116.75 ms. The width of the spectrum is 51, 53, and 54 for Philips, GE, and Siemens respectively.

# system atrophy of the cerebellar type (Specht et al., 2005; Minneropet al., 2007).

Future studies that seek to utilize quantitative FSE-T2 measures 386will need to standardize the pulse sequence across scanners or devise 387 a post-processing method to standardize measures. Alternatively, 388 small fluid-filled objects with known T2 values could be scanned 389 alongside each subjects' head to provide reference signal (House et al., 390 2006). Using other sequences may also help us to understand some of 391 the differences between how each vendor handles the processing of 392 T2 based imaging, but since these are not generally used in multi-site 393 studies it is difficult to say how this will help us to understand the 394 differences we have found using the FSE sequence. 395

396This study used MRI and neuropsychological test ADNI data across397NL, MCI and AD subjects. MRI data acquired with GE, Philips, and398Siemens scanners to examine which properties of FSE-T2 quantitative

MRI may be useful for the classification of MCI and early AD. 399 Significant quantitative FSE-T2 differences were found between 400 vendors in peak FSE-T2 and histogram width. The results herein 401 suggest that FSE-T2 histogram measures can vary significantly with 402 scanner vendor. Specifically, Siemens data consistently produced 403 higher peak FSE-T2 values and broader histogram widths than either 404 GE or Philips. The second purpose was to examine T2 histograms 405 within normal aging, MCI, and AD over the whole brain. Few 406 significant differences were found between diagnostic groups and 407 the observed trends were inconsistent amongst the represented 408 vendors, suggesting a potential scanner-disease interaction. The 409 differences in scanner overshadowed the potential influence of 410 subject diagnostic group on FSE-T2 measures. Significant correlations 411 between peak FSE-T2 and FSE-T2 histogram width with global scale of 412 dementia and measures of memory and cognitive functioning were 413 observed. 414

To the authors' knowledge, a multi-site study involving quantita- 415 tive FSE-T2 datasets from GE, Philips, and Siemens has not been 416 reported in previous literature. The results obtained in this study 417 should serve to encourage increased quality control for measures of 418 FSE-T2 related scans in large-scale studies utilizing data from multiple 419 scanner platforms. They also point out potential differential effects of 420 scanner brand that may not be adequately controlled by adding a co-421 variate to a statistical analysis. 422

### Conclusion

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The aim of this study was to assess the utility of FSE-T2 424 quantitative MRI of the brain for the diagnosis of AD and its early 425 manifestations. The results indicate that FSE-T2 measures can vary 426 significantly between scanner platforms and that FSE-T2 quantitative-427 MRI image processing algorithms which include the platform specific 428 magnetization dynamic effects during the FSE readouts are needed for 429 reconciling multi-platform FSE-T2 measurements. Despite these 430 differences, overall FSE-T2 relaxation properties were related to the 431 global dementia status of the subjects. 432

<b>Incited</b> ref	ferences
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Mueller et al., 2005	435
Petersen et al., 2005	436

### Acknowledgments

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