

# Influence of Age, BMI, Gender and Lumbar Level on T1ρ Magnetic Resonance Imaging of Lumbar Discs in Healthy Asymptomatic Adults

## Einfluss von Alter, Geschlecht, BMI und lumbalem Level auf T1ρ-MRT-Bildgebung lumbaler Bandscheiben gesunder asymptomatischer Erwachsener

### Authors

Raphael Gübitz<sup>1</sup>, Tobias Lange<sup>2</sup>, Georg Gosheger<sup>2</sup>, Walter Heindel<sup>3</sup>, Thomas Allkemper<sup>3</sup>, Christoph Stehling<sup>4</sup>, Joachim Gerss<sup>5</sup>, Christian Kanthak<sup>6</sup>, Tobias L. Schulte<sup>7</sup>

### Affiliations

- 1 Department of Radiology and Neuroradiology, Asklepios Hospital Altona, Hamburg, Germany
- 2 Department of Orthopaedics and Tumor Orthopaedics, University Hospital Münster, Münster, Germany
- 3 Department of Clinical Radiology, University Hospital Münster, Münster, Germany
- 4 Clinic for Radiology and Neuroradiology, Sankt-Barbara Hospital Ham-Heessen, Hamm, Germany
- 5 Institute of Biostatistics and Clinical Research, University of Münster, Münster, Germany
- 6 Fraunhofer MEVIS, Institute for Medical Image Computing, Bremen, Germany
- 7 Department of Orthopaedics and Trauma Surgery, St. Josef Hospital, Ruhr-University Bochum, Bochum, Germany

### Key words

T1ρ, spine, magnetic resonance imaging, lumbar disc, asymptomatic adults

received 07.03.2017

accepted 06.06.2017

### Bibliography

DOI <https://doi.org/10.1055/s-0043-115898>

Published online: 1.9.2017 | Fortschr Röntgenstr 2018; 190: 144–151 © Georg Thieme Verlag KG, Stuttgart · New York, ISSN 1438-9029

### Correspondence

Herr Dr. Raphael Gübitz  
Radiology, Asklepios Klinik Altona, Paul-Ehrlich-Straße 1,  
22763 Hamburg, Germany  
Tel.: ++49/40/18 18 81 18 11  
[r.guebitz@asklepios.com](mailto:r.guebitz@asklepios.com)

### ZUSAMMENFASSUNG

**Ziel** Ermittlung der Größenordnung der T1ρ Werte der lumbalen Bandscheiben in gesunden asymptomatischen Probanden bei 1,5T. Zusätzlich wurde der Einfluss von Alter, body

mass index (BMI), Geschlecht und lumbaler Level auf die T1ρ Relaxation.

**Material und Methoden** In der prospektiven Studie wurden 81 freiwillige Probanden zwischen 20 und 80 Jahren eingeschlossen und in drei Altersgruppen unterteilt (A, 20–39 Jahre; B, 40–59 Jahre; C, 60–80 Jahre). Alle Probanden wurden in einem 1,5 T MRT untersucht und sagittale T1ρ Bilder akquiriert. Die ermittelten T1ρ Relaxationszeiten wurden korreliert mit Alter, BMI, Geschlecht und lumbalem Level jeweils bezogen auf die gesamte Bandscheibe, den Annulus fibrosus und den Nucleus pulposus.

**Ergebnisse** Das Alter zeigte einen signifikanten Einfluss auf die T1ρ Relaxationszeit in allen lumbalen Leveln wobei zunehmendes Alter mit abnehmenden Relaxationszeiten verbunden war. Darüber hinaus zeigte sich ein signifikanter Unterschied zwischen den Altersgruppen A vs. C und B vs. C ( $P=0,0008$  und  $P=0,0149$ ). Kein signifikanter Unterschied bestand zwischen den T1ρ Relaxationszeiten von Männern und Frauen ( $P>0,05$ ). Der BMI zeigte eine signifikant negative Korrelation mit der T1ρ Relaxationszeit ( $P<0,0001$ ). Hinsichtlich des lumbalen Levels zeigte sich eine signifikante Abnahme der Relaxationszeiten von L 1/2 zu L5/S1 ( $P=0,0013$ ).

**Schlussfolgerung** Steigendes Alter korrelierte signifikant mit zunehmender Degeneration lumbaler Bandscheiben bei asymptomatischen Probanden, insbesondere ab einem Alter von 60 Jahren. Ein hoher BMI korrelierte ebenfalls signifikant mit zunehmender Degeneration. Die unteren lumbalen Bandscheiben zeigten insgesamt eine fortgeschrittenere Degeneration als die oberen.

### Kernaussagen

- Steigendes Alter vermindert die T1ρ Relaxationszeit der Bandscheiben signifikant ( $P<0,05$ )
- Geschlecht zeigt keinen signifikanten Einfluss auf die T1ρ Relaxationszeit ( $P>0,05$ )
- BMI korreliert signifikant negativ mit der T1ρ Relaxationszeit ( $P<0,01$ ).
- Signifikant niedrigere Relaxationszeiten der unteren gegenüber oberen LWS ( $P<0,01$ ).

## ABSTRACT

**Purpose** To assess the T1 $\rho$  range of lumbar intervertebral discs in healthy asymptomatic individuals at 1.5 T and to investigate the influence of age, body mass index (BMI), gender, and lumbar level on T1 $\rho$  relaxation.

**Materials and Methods** In a prospective study, a total of 81 volunteers aged 20–80 years were included in this study and divided into three age groups (A: 20–39y; B: 40–59y; C: 60–80y). All of the volunteers underwent magnetic resonance imaging (MRI) at 1.5 T with acquisition of sagittal T1 $\rho$  images. The calculated T1 $\rho$  relaxation times were correlated with age, BMI, gender, and lumbar level relative to the total disc, the annulus fibrosus, and the nucleus pulposus.

**Results** Age had a significant influence on T1 $\rho$  relaxation times at all lumbar levels, with increasing age being associated with reduced relaxation times. There was also a significant difference between age groups A vs. C and B vs. C ( $P=0.0008$  and  $P=0.0149$ , respectively). No significant differences in T1 $\rho$  relaxation time were observed between men and women ( $P>0.05$ ). BMI showed a significant negative correlation with T1 $\rho$  relaxation times ( $P<0.0001$ ). Analysis of the lumbar level

revealed a significant decrease in relaxation times from L1/2 to L5/S1 ( $P=0.0013$ ).

**Conclusion** Increasing age correlated significantly with advanced lumbar disc degeneration in asymptomatic individuals, particularly in those aged 60 or older. Increasing BMI correlated significantly with increasing degeneration. The lower discs showed more degeneration than the upper ones.

### Key Points

- Increasing age significantly reduces the T1 $\rho$  relaxation time in the intervertebral discs ( $P<0.05$ )
- Gender does not significantly influence T1 $\rho$  relaxation times ( $P>0.05$ )
- BMI shows a significant negative correlation with T1 $\rho$  relaxation times ( $P<0.01$ )
- Significantly shorter relaxation times in lower lumbar spine vs. upper lumbar spine ( $P<0.01$ )

### Citation Format

- Gübitz R, Lange T, Gosheger G et al. Influence of Age, BMI, Gender and Lumbar Level on T1 $\rho$  Magnetic Resonance Imaging of Lumbar Discs in Healthy Asymptomatic Adults. *Fortschr Röntgenstr* 2018; 190: 144–151

## Introduction

Magnetic resonance imaging (MRI) is the gold standard for examining the lumbar discs in routine clinical practice [1]. Degenerative changes are frequently found, with growing prevalence in patients of increasingly advanced age. However, these changes do not necessarily always correlate with the patients' clinical symptoms. Signs of degeneration may even be found in asymptomatic adults. For surgeons and physicians, it is crucial to correlate the radiographic findings with the clinical symptoms in order to develop specific treatment strategies. It is therefore important to have information about the prevalence of degenerative changes in healthy asymptomatic individuals.

One of the earliest, important studies on abnormal MRI of the lumbar spine in asymptomatic individuals, published by Boden et al. in 1990, presented the MRI results for 67 adults and identified degenerative changes in approximately one-third of them [2]. The authors concluded that there is a need for strict correlation between MRI changes, patient age, and symptoms before any treatment decisions are made [2]. Other studies showed that degenerative changes can be found to some extent in asymptomatic subjects [3, 4].

The study by Boden et al. was based only on qualitative MRI data. Today, quantitative imaging techniques are available [5]. Among these, T1 $\rho$  imaging appears to be the most promising method to become available for routine use in the study of disc pathology [6]. The T1 $\rho$  relaxation time describes the spin-lattice relaxation in the rotating frame that occurs after application of a spin-lock pulse. This spin-lock technique makes it possible to observe slow-motion processes such as interactions between water and extracellular matrix molecules [7]. Early studies showed

a significant correlation between the T1 $\rho$  relaxation time and the loss of proteoglycan molecules in cartilage [8]. In the spine, T1 $\rho$  relaxation times showed relevant correlations with the amount of proteoglycans in the extracellular matrix of the lumbar disc, with longer relaxation times indicating greater amounts of proteoglycans [7, 9, 10]. In addition, T1 $\rho$  relaxation times correlated significantly with the water content in the intervertebral lumbar disc [7]. In comparison with T2 mapping, earlier studies suggest that T1 $\rho$  might be more sensitive to early degenerative changes, especially in the annulus fibrosus [11, 12]. T1 $\rho$  therefore appears to be a promising noninvasive tool for detecting early disc degeneration in vivo.

The purpose of the present study was to systematically evaluate the differences in T1 $\rho$  data for the lumbar intervertebral discs in asymptomatic individuals of different ages. In addition, it was investigated whether body mass index (BMI), gender, and lumbar level influence T1 $\rho$  relaxation times. In the study, T1 $\rho$  was used as a noninvasive indicator for disc degeneration and the ability of T1 $\rho$  to quantify disc degeneration was assumed [11–13].

## Materials and Methods

This prospective study was approved by the local ethics committee (ref. no. 2013–025-f-S) and supported by a research grant from the German Spine Society. All of the participating volunteers provided written informed consent prior to examination.

### Volunteers

This study included healthy asymptomatic individuals between 20 and 80 years old. The exclusion criteria were: any current lumbar

back pain or pain radiating into the leg(s) at the time of examination; a medical history including any lumbar back pain or radiating pain requiring any type of medical treatment; any history of lumbar back pain or radiating pain lasting more than 24 hours; and any inability to work due to the above symptoms. Based on the fact that degeneration increases with age, three groups were defined in order to evaluate age-related effects: group A: 20–39y; group B: 40–59y; group C: 60–80y. In addition, the volunteers' sex, height, weight, and body mass index (BMI) were recorded. Each individual underwent a standardized physical examination and structured interview by an experienced orthopedic spine surgeon before inclusion in the study. In addition, each volunteer completed a standardized questionnaire regarding medical history focusing on back pain and spine pathologies.

## Magnetic Resonance Imaging

All of the images were acquired using a 1.5-T clinical MRI system (Achieva; Philips Healthcare, Best, Netherlands) with an eight-channel coil. Sagittal T1 $\rho$  images with 14 slices in the lumbar spine were acquired for all subjects. The spin-lock times were 0, 10, 20, 40 and 80 ms. Other imaging parameters were as follows: repetition time msec/echo time msec 14/7; flip angle 15°; field of view 200 × 200 × 42 mm; acquisition matrix 248 × 215; and specific absorption rate < 0.2 W/kg.

## Image Analysis

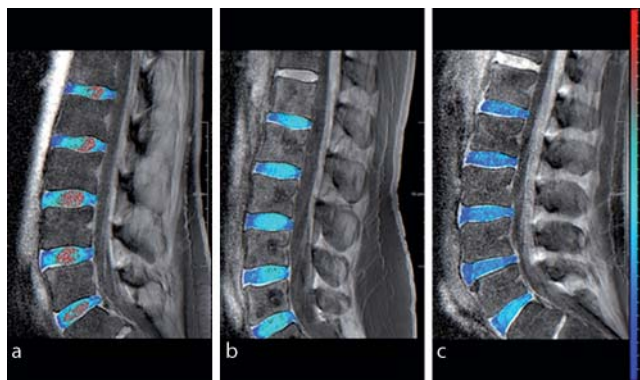
The T1 $\rho$  images were analyzed using specially developed software (Fraunhofer MEVIS, Institute for Medical Image Computing, Bremen, Germany). With the help of this software program, T1 $\rho$  maps were calculated on a pixel-by-pixel basis using a monoexponential decay mode:  $M(SLT) = M_0 \times \exp(-SLT / T1\rho)$ .  $M_0$  and  $M(SLT)$  denote the equilibrium magnetization, and T1 $\rho$  refers to the prepared magnetization for each slice thickness (SLT). T1 $\rho$  maps were generated from these data for each section by fitting all pixel intensity data as a function of SLT, using a Levenberg–Marquardt fitting algorithm.

T1 $\rho$  segmentation was performed on sagittal T1 $\rho$  maps for each disc from L1/2 to L5/S1 in each individual by manually outlining the total intervertebral disc and the nucleus pulposus. This made it possible to investigate the intervertebral disc in general, as well as to differentiate between the annulus fibrosus and the nucleus pulposus (an example is shown in ► Fig. 1). Finally, three different mean relaxation times were calculated for each disc.

To account for potential interobserver variability in outlining the whole intervertebral disc or the nucleus pulposus and annulus fibrosus, intervertebral discs in 10% of the study population were segmented by two readers and the calculated T1 $\rho$  relaxation times were compared.

## Statistics

The influence of age, BMI, gender, and lumbar level on T1 $\rho$  relaxation was investigated, with differentiation between the total disc, the annulus fibrosus, and the nucleus pulposus. Pearson's correlation coefficients were calculated and Student's *t*-tests were carried out where appropriate. A multivariate linear mixed model was established, with the outcome variable of T1 $\rho$  relaxation



► Fig. 1 Examples of T1 $\rho$  maps for each group (a: group A; b: group B; c: group C), with segmentation of all five lumbar discs. Color coding: from dark blue, representing low relaxation times, to red, representing high relaxation times.

time and explanatory variables of age group, BMI, lumbar segment (L1/2–L5/S1) and the three anatomic structures (total disc, nucleus pulposus, annulus fibrosus). Correlations between individual measurements taken from the same individual were accounted for by means of a subject-specific random intercept. Heterogeneous variances in measurements at the three different anatomic structures were taken into account by appropriately partitioning the variance–covariance matrix of the error term. The linear mixed model was fitted using the restricted maximum likelihood method. *P* values were regarded as significant at  $P \leq 0.05$ . No adjustment for multiple testing was performed. An overall significance level was not determined and cannot be calculated.

Power analysis showed that with a sample size of at least 24–30 individuals per group, differences in the mean T1 $\rho$  relaxation times  $\geq 10$  ms could be detected with 80% power in a (two-sided) Student's *t*-test at  $\alpha = 5\%$ , assuming a standard deviation of 10–20 ms.

Interobserver variability in the calculated T1 $\rho$  relaxation times from the two readers was investigated by calculating the mean relative difference as well as the intraclass correlation coefficient (ICC).

Descriptive and inferential statistical analyses were carried out using IBM SPSS Statistics for Windows, version 22 (IBM Corporation, Armonk, New York, USA) and SAS 9.4 for Windows (SAS Institute Inc., Cary, North Carolina, USA).

## Results

A total of 81 volunteers were included (41 women and 40 men): group A (age 20–39y),  $n = 24$  (13 women, 11 men); group B (age 40–59y),  $n = 30$  (15 women, 15 men); group C (age 60–80y),  $n = 27$  (13 women, 14 men). The baseline data for all of the volunteers are presented in ► Table 1. They all underwent the MRI examinations successfully, with no complications, and all of the data proved to be of sufficiently high quality to provide robust T1 $\rho$  results.

► **Table 1** Baseline Data for All 81 Volunteers.

group	age (years)			height (cm)			weight (kg)			BMI (kg/m <sup>2</sup> )		
	mean	min.	max.	mean	min.	max.	mean	min.	max.	mean	min.	max.
a	29.2	20.0	39.0	177.3	163.0	200.0	72.0	55.0	93.0	22.8	18.8	28.4
b	47.6	40.0	58.0	175.3	155.0	194.0	74.7	55.0	95.0	24.4	19.4	36.2
c	66.3	60.0	78.0	171.4	160.0	187.0	77.3	46.0	115.0	26.1	17.1	36.3

group a (age 20–39), n = 24 (13 women, 11 men); group a (age 40–59), n = 30 (15 women, 15 men); group c (age 60–80 years), n = 27 (13 women, 14 men); SD, standard deviation.

The pre-test revealed a mean difference in the calculated T1p relaxation times between the two readers of 2% for the total disc (max. difference 5%), 6% for the annulus fibrosus (max. difference 11%), and 1% for the nucleus pulposus (max. difference 7%). The intraclass correlation coefficients (ICCs) were 0.989 for the total disc, 0.937 for the annulus fibrosus, and 0.980 for the nucleus pulposus. The interobserver variability was thus very low.

The results for T1p relaxation times are listed in ► **Table 2**, ► **Fig. 2–4**, with differentiation between groups A–C, the various lumbar segments (L1/2–L5/S1), and the three anatomic structures (total disc, nucleus pulposus, and annulus fibrosus). Student’s t-test did not show any significant differences with regard to T1p relaxation times between women and men for each group, segment, or anatomic region.

Analysis of the lumbar level showed a significant decrease in the T1p relaxation times between L1/2 and L5/S1 (P = 0.0013). With regard to the different parts of the intervertebral disc, there were significant differences between the total disc, the nucleus pulposus, and the annulus fibrosus (P < 0.001 for all combinations). All in all, the T1p relaxation time was 9.6 ms lower on average in the annulus fibrosus than in the total disc and 25.2 ms lower than in the nucleus pulposus. In addition, the T1p relaxation time in the nucleus pulposus was a mean of 15.6 ms higher than in the total disc.

► **Table 3** shows the influence of age as a continuous variable (in years) on T1p relaxation times, using a Pearson correlation test. In relation to the three age groups (A–C), significant differences were detected between groups A and C (P = 0.0008) and between groups B and C (P = 0.0149) using a linear mixed model with inclusion of lumbar level, age group, and disc part. Additionally including BMI in the model still led to a significant difference between groups A and C (P = 0.037), while the difference between groups B and C became borderline significant (P = 0.0651).

With regard to BMI, the linear mixed model showed a significant negative correlation between BMI and T1p relaxation time (P < 0.0001). ► **Table 4** presents correlations between BMI and T1p relaxation times relative to the different age groups, lumbar levels, and parts of the intervertebral discs. A significant impact, with high Pearson correlation coefficients of BMI on T1p relaxation times, was found here for several combinations of segment and age group.

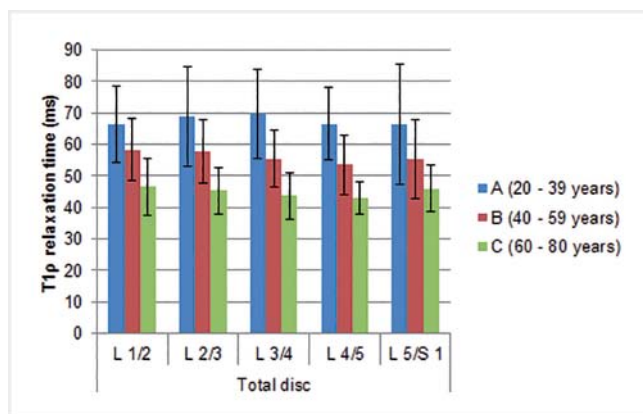
## Discussion

The aims of this study were to establish T1p data for the intervertebral discs in asymptomatic individuals and to evaluate the influence of age, BMI, gender, and lumbar level on T1p relaxation times.

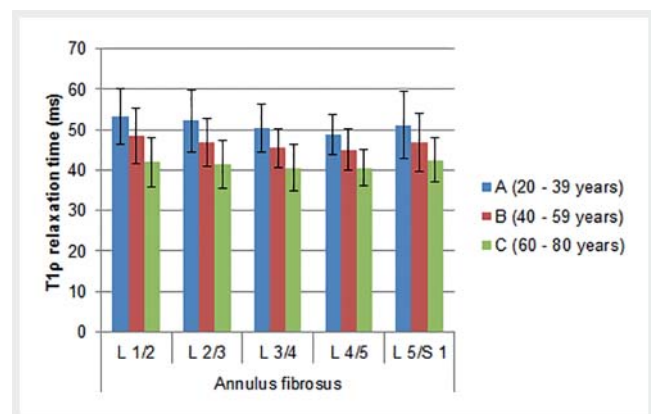
To the best of the authors’ knowledge, this is the first study that has systematically investigated T1p imaging at 1.5 T in asymptomatic individuals between the ages of 20 and 80. According to Boden et al., three age groups were defined: A: 20–39y; B: 40–59y; and C: 60–80y [2]. Means and standard deviations showed a homogeneous distribution of ages across the groups in the study. Half of the individuals examined were women and half were men, to allow gender effects to be studied.

► **Table 2** T1 $\rho$  Relaxation Times (in ms) for All Lumbar Segments (L1/2 to L5/S1) for the Total Disc, Annulus Fibrosus, and Nucleus Pulposus.

	L1/2		L2/3		L3/4		L4/5		L5/S1	
	mean	SD	mean	SD	mean	SD	mean	SD	mean	SD
total disc T1 $\rho$ relaxation times (ms)										
group a	66.51	12.10	69.05	15.86	69.78	14.09	66.52	11.50	66.40	19.24
group b	58.32	9.89	57.82	10.04	55.48	8.91	53.61	9.47	55.36	12.68
group c	46.51	8.90	45.27	7.54	43.73	7.36	42.85	5.21	46.03	7.55
annulus fibrosus T1 $\rho$ relaxation times (ms)										
group a	53.24	6.87	52.26	7.70	50.35	6.03	48.82	5.02	51.11	8.37
group b	48.51	6.94	46.98	5.92	45.48	4.77	45.05	5.19	46.86	7.32
group c	42.00	6.12	41.50	5.91	40.54	5.77	40.59	4.44	42.49	5.37
nucleus pulposus T1 $\rho$ relaxation times (ms)										
group a	90.38	21.42	96.87	34.69	98.86	27.12	94.69	22.64	94.94	39.02
group b	74.30	15.77	73.57	17.70	70.21	15.82	66.80	16.18	69.20	21.91
group c	54.00	13.67	51.64	10.89	49.33	10.34	46.99	8.20	51.48	12.64



► **Fig. 2** T1 $\rho$  relaxation times (in ms with standard deviation) in all five lumbar discs for the three age groups (group A: blue; group B: red; group C: green) for the total disc.



► **Fig. 3** T1 $\rho$  relaxation times (in ms with standard deviation) in all five lumbar discs for the three age groups (group A: blue; group B: red; group C: green) for the annulus fibrosus.

## Age

Increasing age was associated with lower T1 $\rho$  relaxation times, with a highly significant and uniform impact (► **Table 3**). The results in the annulus fibrosus show the shortest relaxation times and those in the nucleus pulposus the longest. The values for the total discs are in between, as mixtures of the two anatomical structures. Published T1 $\rho$  studies of the spine have reported similar results, with decreasing relaxation times with increasing age [7, 11, 13–15].

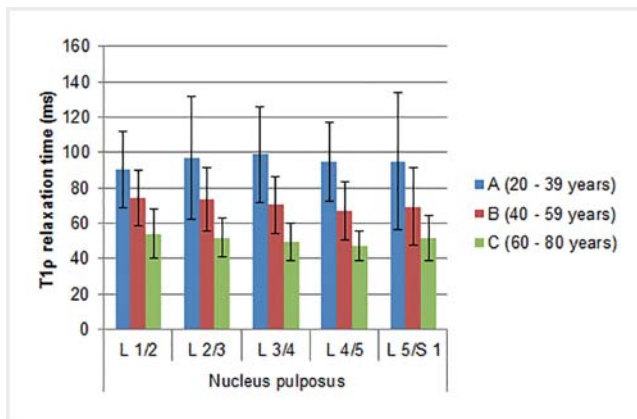
Johannessen et al. studied T1 $\rho$  relaxation times at 1.5T in the nucleus pulposus of seven fresh frozen cadaveric human lumbar spine sections, with a mean age of 51.6 years [7]. They found decreasing T1 $\rho$  relaxation times with increasing age and degeneration, and also observed a strong linear correlation between T1 $\rho$  and the proteoglycan content.

In a study including 11 healthy volunteers at 3 T, Blumenkrantz et al. found a significant correlation between the individuals' age and the T1 $\rho$  relaxation times measured [14]. The volunteers' mean age was 31.3 years, with a range of 23–60 years. The significant influence of age was evident for the T1 $\rho$  relaxation time in the nucleus pulposus as well as the annulus fibrosus, but the correlation in the annulus was lower than that in the nucleus.

Another study on the influence of age on T1 $\rho$  relaxation time was carried out by Filippi et al. at 3 T using parallel-transmission MRI [15]. They studied 34 individuals (mean age: 38.4 for men and 36.5 years for women) with no history of back pain, who were further divided into four age groups (20–29, 30–39, 40–49, 50–59y). A significant moderate negative correlation was observed between T1 $\rho$  and age in all of the age groups. There was also a statistically significant difference in T1 $\rho$  relaxation

► **Table 3** Influence of Age (in Years) on T1 $\rho$  Relaxation Times, using a Pearson Correlation Test.

		L1/2	L2/3	L3/4	L4/5	L5/S1
total disc	pearson correlation	-0.650	-0.698	-0.786	-0.770	-0.589
	p value (two-sided)	0.000	0.000	0.000	0.000	0.000
annulus fibrosus	pearson correlation	-0.611	-0.599	-0.640	-0.598	-0.484
	p value (two-sided)	0.000	0.000	0.000	0.000	0.000
nucleus pulposus	pearson correlation	-0.687	-0.687	-0.804	-0.806	-0.644
	p value (two-sided)	0.000	0.000	0.000	0.000	0.000



► **Fig. 4** T1 $\rho$  relaxation times (in ms with standard deviation) in all five lumbar discs for the three age groups (group A: blue; group B: red; group C: green) for the nucleus pulposus.

times between all of the studied age groups, a finding that is also consistent with the present results.

The influence of age on T1 $\rho$  relaxation times in the lumbar intervertebral discs in individuals with disc degeneration and clinical symptoms was evaluated in a study by Blumenkrantz et al. [11]. Sixteen individuals with a mean age of 40.2 years were studied with imaging at 3 T. There was a significant negative correlation between T1 $\rho$  relaxation times and age and a significant positive correlation between age and Pfirrmann grade.

In a mixed group of asymptomatic individuals and patients with low back pain, Wang et al. also found a significant reduction in T1 $\rho$  relaxation times along with increasing age [13]. With regard to the different parts of the intervertebral disc, the reduction was more pronounced in the nucleus pulposus than in the annulus fibrosus.

In contrast to the above studies as well as the present results, Zhou et al. did not find a significant influence of age on the T1 $\rho$  relaxation time [16]. They examined 80 patients with low back pain at 1.5 T. While there was a significant negative correlation between Pfirrmann grade and T1 $\rho$  relaxation time, age was not found to have a significant influence on T1 $\rho$  times. One reason for the divergent results between the findings published by Zhou et al. and the present study might be the fact that Zhou et al. only studied patients with low back pain and corresponding degeneration. Their cohort also only comprised patients between 20 and

43 years old, with a mean age of 31.6 years, potentially diminishing the observed effect of age on T1 $\rho$  relaxation times.

## BMI

When the effect of BMI was investigated, a significant negative correlation was found between BMI and T1 $\rho$  relaxation time. Further analyses relative to different age groups, lumbar levels, and the parts of the intervertebral disc showed that this correlation was particularly strong in participants in age group C (60–80y).

The only published study to date on the effect of BMI on T1 $\rho$  relaxation times is that of Zobel et al., which did not find any significant correlations between BMI and T1 $\rho$  values [17], in contrast to the results of the present study. However, it is important to note that Zobel et al. conducted their study in 63 healthy young individuals with a mean age of 22.95 years. As the most prominent correlation between BMI and T1 $\rho$  relaxation times was found in age group C in the present study, it might be possible that the young age of the population included in the study by Zobel et al. led to the lack of any significant influence of BMI.

With regard to studies using standard clinical MRI to assess intervertebral disc degeneration, there are no clear results in the literature on the influence of BMI. Some studies have not identified any significant influence of BMI on disc degeneration [18, 19], but the majority of studies have reported a correlation between BMI and disc degeneration [20–22].

## Gender

The results of the present study did not show any significant effect of gender on relaxation times. This finding corresponds with the results published by Filippi et al., who also did not identify any statistically significant differences in T1 $\rho$  values between men and women in a study at 3 T including 34 healthy volunteers [15]. In contrast, in a T1 $\rho$  study including healthy young adults, Zobel et al. found significantly lower relaxation times in women in the L3/4 and L4/5 discs [17]. A trend toward more severe disc degeneration in women was also reported in the study by Wang et al., who used a modified Pfirrmann grading system to assess disc degeneration [23]. They also found significant differences between men and women at the L3/4 and L4/5 levels.

Consistent with the findings of the present study, other reports on intervertebral disc degeneration have not identified a signifi-

► **Table 4** Correlations between BMI and T1 $\rho$  relaxation times relative to age group, lumbar level, and disc part.

		L1/2	L2/3	L3/4	L4/5	L5 / S1
total disc						
group a	pearson correlation	-0.310	-0.412	-0.636	-0.557	-0.328
	p value (two-sided)	0.141	0.046	0.001	0.005	0.117
group b	pearson correlation	-0.337	-0.373	-0.235	-0.306	-0.438
	p value (two-sided)	0.069	0.042	0.212	0.100	0.018
group c	pearson correlation	-0.475	-0.565	-0.630	-0.614	-0.382
	p value (two-sided)	0.012	0.002	0.000	0.001	0.054
annulus fibrosus						
group a	pearson correlation	-0.388	-0.294	-0.500	-0.441	-0.309
	p value (two-sided)	0.061	0.164	0.013	0.031	0.142
group b	pearson correlation	-0.351	-0.415	-0.343	-0.419	-0.544
	p value (two-sided)	0.057	0.023	0.063	0.021	0.002
group c	pearson correlation	-0.519	-0.624	-0.674	-0.657	-0.486
	p value (two-sided)	0.006	0.001	0.000	0.000	0.012
nucleus pulposus						
group a	pearson correlation	-0.385	-0.492	-0.732	-0.540	-0.355
	p value (two-sided)	0.063	0.015	0.000	0.006	0.089
group b	pearson correlation	-0.298	-0.291	-0.160	-0.236	-0.372
	p value (two-sided)	0.109	0.119	0.397	0.209	0.047
group c	pearson correlation	-0.429	-0.507	-0.564	-0.465	-0.135
	p value (two-sided)	0.026	0.007	0.002	0.017	0.510

cant influence of gender on the degenerative process using either clinical MRI [18] or MRI evaluation of fresh cadaveric spines [24].

### Lumbar level

A significant but small decrease in the T1 $\rho$  relaxation times from levels L1/2 to L5 / S1 was observed in the present study. This finding corresponds to the results reported by Vadalá et al., who noted a significant decrease in T1 $\rho$  values in the nucleus pulposus from the upper to the lower lumbar levels when evaluating asymptomatic weightlifters for early intervertebral disc degeneration [25]. This trend toward lower T1 $\rho$  relaxation times in the caudal segments of the spine was also described by Blumenkrantz et al. in a study including 16 patients suffering from intervertebral disc degeneration with clinical symptoms [11].

Studies using standard MRI techniques have also provided evidence for increased disc degeneration in the lower parts of the lumbar spine, supporting the findings of the present study [26–28]. The different loading environment of the discs in the different levels is one the well-known major reasons [29].

### Limitations

Limitations of this study include the fact that only a single radiologist evaluated all of the T1 $\rho$  images. This is justified, however, as the described preliminary test showed a very low level of interobserver variability. In addition, studies in the literature have shown

excellent reproducibility results and high levels of interobserver reliability [6, 16, 30]. It would not have been possible to control for the minimal bias in light of the fact that none of the included individuals had ever suffered from low back pain, even though each individual was examined and interviewed by an orthopedic spine surgeon. T1 $\rho$  imaging was not compared with any other imaging modalities. The main reason for this decision was to keep the total examination time for the volunteers (who underwent a standardized physical examination, structured interview and a standardized questionnaire in addition to MRI) as short as possible to ensure their willingness to take part in the study. The lack of a morphologic parameter such as the Pfirrmann Grade is justified as studies in the past have shown high correlations between T1 $\rho$  and degeneration status in symptomatic patients [6, 11, 12]. Finally, the ability of T1 $\rho$  to quantify disc degeneration was assumed [11–13].

### Conclusion

The results of the present study, which is one of the largest studies on T1 $\rho$  imaging of the spine and includes volunteers up to the age of 80 years, confirm those of earlier investigations indicating that some degree of disc degeneration may be regarded as “normal” in asymptomatic individuals. Age showed a significant and uniform impact on T1 $\rho$  relaxation times. Furthermore, increasing

BMI and a more distal location of the affected discs also proved to influence disc degeneration, whereas gender showed no significant impact. The results of the study underline the possible application of T1 $\rho$  imaging in the evaluation process in patients with intervertebral disc degeneration as it is a quantitative method that isn't susceptible to the radiologist's experience as in the case of qualitative or semi-qualitative evaluation. Still, a close clinical correlation between symptoms and imaging findings is strongly required. Although the acquired data are not entirely transferable to other MRI systems, the results may still serve as baseline data for further studies or as comparative values for other populations.

### Conflict of Interest

The authors declare that they have no conflict of interest.

### Acknowledgements

The authors would like to thank the German Spine Society for providing a research grant in order to enable this investigation to be carried out. Furthermore, the authors would like to thank Hendrik Kooijmann for his assistance in establishing the T1 $\rho$  sequences.

### References

- [1] Berns DH, Blaser SI, Modic MT. Magnetic resonance imaging of the spine. *Clin Orthop Relat Res* 1989; 244: 78–100
- [2] Boden SD, Davis DO, Dina TS et al. Abnormal magnetic-resonance scans of the lumbar spine in asymptomatic subjects. A prospective investigation. *J Bone Joint Surg Am* 1990; 72: 403–408
- [3] Jensen MC, Brant-Zawadzki MN, Obuchowski N et al. Magnetic resonance imaging of the lumbar spine in people without back pain. *N Engl J Med* 1994; 331: 69–73
- [4] Weishaupt D, Zanetti M, Hodler J et al. MR imaging of the lumbar spine: prevalence of intervertebral disk extrusion and sequestration, nerve root compression, end plate abnormalities, and osteoarthritis of the facet joints in asymptomatic volunteers. *Radiology* 1998; 209: 661–666
- [5] Brayda-Bruno M, Tibiletti M, Ito K et al. Advances in the diagnosis of degenerated lumbar discs and their possible clinical application. *Eur Spine J* 2014; 23 (Suppl. 3): S315–S323
- [6] Auerbach JD, Johannessen W, Borthakur A et al. In vivo quantification of human lumbar disc degeneration using T(1rho)-weighted magnetic resonance imaging. *Eur Spine J* 2006 (Suppl. 3): S338–S344
- [7] Johannessen W, Auerbach JD, Wheaton AJ et al. Assessment of human disc degeneration and proteoglycan content using T1rho-weighted magnetic resonance imaging. *Spine (Phila Pa 1976)* 2006; 31: 1253–1257
- [8] Akella SV, Regatte RR, Gougoutas AJ et al. Proteoglycan-induced changes in T1rho-relaxation of articular cartilage at 4T. *Magn Reson Med* 2001; 46: 419–423
- [9] Mulligan KR, Ferland CE, Gawri R et al. Axial T1rho MRI as a diagnostic imaging modality to quantify proteoglycan concentration in degenerative disc disease. *Eur Spine J* 2014; 24: 2395–2401
- [10] Zuo J, Joseph GB, Li X et al. In vivo intervertebral disc characterization using magnetic resonance spectroscopy and T1 $\rho$  imaging: association with discography and Oswestry Disability Index and Short Form-36 Health Survey. *Spine (Phila Pa 1976)* 2012; 37: 214–221
- [11] Blumenkrantz G, Zuo J, Li X et al. In vivo 3.0-tesla magnetic resonance T1rho and T2 relaxation mapping in subjects with intervertebral disc degeneration and clinical symptoms. *Magn Reson Med* 2010; 63: 1193–1200
- [12] Wang YX, Zhao F, Griffith JF et al. T1rho and T2 relaxation times for lumbar disc degeneration: an in vivo comparative study at 3.0-Tesla MRI. *Eur Radiol* 2013; 23: 228–234
- [13] Wang YX, Griffith JF, Leung JC et al. Age related reduction of T1rho and T2 magnetic resonance relaxation times of lumbar intervertebral disc. *Quant Imaging Med Surg* 2014; 4: 259–264
- [14] Blumenkrantz G, Li X, Han ET et al. A feasibility study of in vivo T1rho imaging of the intervertebral disc. *Magn Reson Imaging* 2006; 24: 1001–1007
- [15] Filippi CG, Duncan CT, Watts R et al. In vivo quantification of T1rho in lumbar spine disk spaces at 3 T using parallel transmission MRI. *Am J Roentgenol* 2013; 201: W110–W116
- [16] Zhou Z, Jiang B, Zhou Z et al. Intervertebral disk degeneration: T1rho MR imaging of human and animal models. *Radiology* 2013; 268: 492–500
- [17] Zobel BB, Vadala G, Del Vecovo R et al. T1rho magnetic resonance imaging quantification of early lumbar intervertebral disc degeneration in healthy young adults. *Spine (Phila Pa 1976)* 2012; 37: 1224–1230
- [18] Farshad-Amacker NA, Hughes AP, Aichmair A et al. Determinants of evolution of endplate and disc degeneration in the lumbar spine: a multifactorial perspective. *Eur Spine J* 2014; 23: 1863–1868
- [19] Kanayama M, Togawa D, Takahashi C et al. Cross-sectional magnetic resonance imaging study of lumbar disc degeneration in 200 healthy individuals. *J Neurosurg Spine* 2009; 11: 501–507
- [20] Takatalo J, Karppinen J, Taimela S et al. Body mass index is associated with lumbar disc degeneration in young Finnish males: subsample of Northern Finland birth cohort study 1986. *BMC Musculoskelet Disord* 2013; 14: 87
- [21] Hangai M, Kaneoka K, Kuno S et al. Factors associated with lumbar intervertebral disc degeneration in the elderly. *Spine J* 2008; 8: 732–740
- [22] Samartzis D, Karppinen J, Mok F et al. A population-based study of juvenile disc degeneration and its association with overweight and obesity, low back pain, and diminished functional status. *J Bone Joint Surg Am* 93: 662–670
- [23] Wang YX, Griffith JF, Ma HT et al. Relationship between gender, bone mineral density, and disc degeneration in the lumbar spine: a study in elderly subjects using an eight-level MRI-based disc degeneration grading system. *Osteoporos Int* 2011; 22: 91–96
- [24] Siemionow K, An H, Masuda K et al. The effects of age, sex, ethnicity, and spinal level on the rate of intervertebral disc degeneration: a review of 1712 intervertebral discs. *Spine (Phila Pa 1976)* 2011; 36: 1333–1339
- [25] Vadala G, Russo F, Battisti S et al. Early intervertebral disc degeneration changes in asymptomatic weightlifters assessed by t1rho-magnetic resonance imaging. *Spine (Phila Pa 1976)* 2014; 39: 1881–1886
- [26] Evans W, Jobe W, Seibert C. A cross-sectional prevalence study of lumbar disc degeneration in a working population. *Spine (Phila Pa 1976)* 1989; 14: 60–64
- [27] Ong A, Anderson J, Roche J. A pilot study of the prevalence of lumbar disc degeneration in elite athletes with lower back pain at the Sydney 2000 Olympic Games. *Br J Sports Med* 2003; 37: 263–266
- [28] Videman T, Battie MC, Gill K et al. Magnetic resonance imaging findings and their relationships in the thoracic and lumbar spine. Insights into the etiopathogenesis of spinal degeneration. *Spine (Phila Pa 1976)* 1995; 20: 928–935
- [29] Keorochana G, Taghavi CE, Lee KB et al. Effect of sagittal alignment on kinematic changes and degree of disc degeneration in the lumbar spine: an analysis using positional MRI. *Spine (Phila Pa 1976)* 2011; 36: 893–898
- [30] Allkemper T, Sagmeister F, Cicinnati V et al. Evaluation of fibrotic liver disease with whole-liver T1rho MR imaging: a feasibility study at 1.5 T. *Radiology* 2014; 271: 408–415