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#### ORIGINAL RESEARCH

# Reliability of Hounsfield Unit for Assessing Asymmetrical Vertebral Bone Mass in Adult Degenerative Scoliosis

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**Objective:** Hounsfield Unit (HU) has been used to investigate the asymmetrical vertebral bone mass in patients associated with adult degenerative scoliosis (ADS). Therefore, there is an inevitable need to evaluate the performance of HU values in ADS subjects.

**Methods:** A total of 162 patients (81 ADS patients and 81 non-ADS patients) aged  $\geq$ 50 years undergoing the CT examination were reviewed. The HU values of the lumbar vertebral body (including total, convex side, and concave side) at bilateral pedicle plane were obtained and compared. The paired *t*-test, chi-squared test, independent samples *t*-test, and interclass correlation coefficient (ICC) were used for statistical analyses.

**Results:** The HU values were significantly different between the convex and concave sides of the lumbar vertebral body (P < 0.01). The total prevalence of osteoporosis (OP) in ADS patients was higher than that of non-ADS patients. The prevalence of OP in female or male of ADS patients was higher than that of non-ADS patients, respectively. Intra- and inter-rater reliability were very strong (both >0.8) for measuring asymmetrical vertebral bone mass in ADS patients.

**Conclusion:** HU value was a high reproducibility method for evaluating the vertebral bone mass in ADS patients. The HU values at the concave sides were significantly higher than that of convex sides at the lumbar vertebral body on the pedicle plane. The prevalence of OP in ADS patients was higher than that of non-ADS patients, especially for females associated with ADS. Moreover, the static asymmetric load did not enhance the bone mass at the concave side compared with the left/right side of non-ADS patients. **Keywords:** adult degenerative scoliosis, bone mass, Hounsfield units, reliability

## Introduction

Adult degenerative scoliosis (ADS) was believed to be a predominant spinal disorder disease in recent years.<sup>1</sup> It may cause low back pain or disability in elderly patients. Hospital admissions for ADS have increased in recent years with the aging population. Osteoporosis (OP) was a common degenerative condition in aging people,<sup>2</sup> and the prevalence of OP was higher in ADS patients compared with other patients.<sup>3</sup> Given that a part of ADS patients may need fixation surgery, OP conditions in ADS patients should be taken into consideration by the surgeon before surgery. ADS itself produces asymmetrical mechanics on the convex and concave sides of the spine, which can aggravate the procedure of ADS. Accumulating evidence has shown that sustained biomechanical imbalance of scoliosis may contribute to asymmetric remolding of vertebral microstructure.<sup>4,5</sup> However, much of our understanding of asymmetric loading-induced bone remodeling comes from data collected on adolescent idiopathic scoliosis patients (AIS),<sup>5</sup> and microstructural changes and remolding of AIS vertebral body between

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concave and convex sides have been demonstrated by several studies.<sup>6,7</sup> Whether there was an ultrastructural difference between the concave and convex side in the vertebral body of ADS spine has not been well elucidated until now.

The gold standard method for determining osteoporosis was the dual-energy x-ray (DXA) absorptiometry.<sup>2</sup> Yet, degenerative lumbar conditions often lead to falsely increasing scores of DXA measurements of lumbar bone mineral density (BMD).<sup>8</sup> Moreover, the bone microstructure of the vertebral body cannot be precisely evaluated by DXA.<sup>9</sup> As an alternate measurement to DXA, the Hounsfield units (HU) value, measuring CT attenuation of tissue, has been proposed to evaluate the BMD in recent years.<sup>10–12</sup> The HU measurement can avoid degenerative conditions and cortical bone in 3-dimensional CT images. Surgeons can manually set the region of interest (ROI) to measure the HU value of vertebral trabecular bone, which is more sensitive to detecting the bone mass of the vertebral body than DXA method. Therefore, the HU value may be a useful tool to evaluate the bone mass difference between the concave and convex side of the vertebral body of ADS spine. In recent years, Wang et al found that HU measurement was valuable for evaluating the asymmetrical vertebral bone mass.<sup>13</sup> However, no studies have so far addressed the issue of the reliability of HU values on bone mass associated with ADS. Hence, the aim of this study was to evaluate the performance of HU value on the vertebral bone mass and characterize the bone mass features of lumbar vertebrae in ADS patients by comparing with non-ADS patients. The results we got here prove that HU value was a valuable and reliable tool to evaluate the bone quality of vertebral body. Special attention should be taken by the surgeon when facing the elderly ADS patients.

## **Materials and Methods**

### Patient Cohort

After IRB approval (2021PHB238), we retrospectively reviewed all patients aged 50 years or older in our hospital from June 2015 to June 2020, and finally include 81 ADS and 81 pared-non-ADS patients according to the inclusion and exclusion criteria. The inclusion criteria were as follows: (1) patients aged 50 years and older; for ADS patients (2) with diagnosis of ADS based on physical examination and whole spinal radiography (Cobb angle  $\geq 10^{\circ}$ );<sup>14,15</sup> (3) the upper end vertebrae (the vertebrae with the greatest inclination at the rostral side of spine) should be upon the L1 vertebrae; for non-ADS patients (4) with diagnosis of lumbar spinal stenosis or lumbar disc herniation based on CT or MR examinations without lumbar scoliosis (Cobb angle  $<10^{\circ}$ ). (5) all included patients underwent three-dimension CT scanning at hospital. While, exclusion criteria were (1) patients with spinal spondylolysis, tuberculosis, tumor, trauma, and skeletal hyperostosis; (2) patients with history of lumbar spinal surgery; (3) patients with metabolic bone disease or taking medications such as chronic glucocorticoids; (4) patients without lumbar CT in our hospital. The informed consents were obtained from all included patients.

## HU Value

All included patients underwent three-dimension reconstructive CT (GE discovery, a tube voltage of 120kV). The parameters of CT had a slice thickness of 1.25 mm with 0.625 mm intervals. The coronal and sagittal planes were acquired from the 3D reconstruction of lumbar CT. The supine position was adopted by patients, and lumbar spine should be ensured in a neutral position. Picture archiving and communication systems was used to get the HU value with the method shown in Figure 1: an oval ROI of 400 mm<sup>2</sup> was placed at an axial image on the pedicle plane; two oval ROI of 75 mm<sup>2</sup> were respectively positioned at the bilateral regions of vertebral body also on the pedicle plane. The ROI should focus on trabecular bone as possible and avoid cortical bone, hemangiomas, and dense bone islands. As shown in Figure 2, a 59 years old male patient was diagnosed as ADS, the HU values were obtained from CT images, and we found that the HU values on the concave side was higher than that of convex side. Measurements were performed by one independent observer, and mean values were calculated. The HU value of L1 vertebral body  $\leq$ 110HU with specificity (90%) was used to determine the osteoprosis.<sup>10,11,16,17</sup>

After training of ROI drawing, two observers (observer 1: with 10 years of experience in degenerative spinal surgery, and observer 2: with 6 years of experience in spinal surgery) independently measured the data of HU value of L1-L5 in each patient. The measurement results of these two observers were used to assess the inter-rater reliability, and the measurement was repeated by the first observer with a washout period of at least 2 weeks, in order to assess the intra-rater reliability. The information of patients was blinded to each other's results.

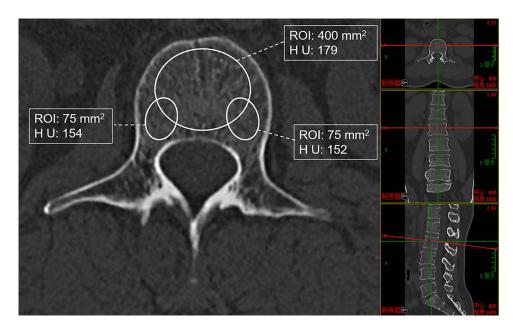


Figure 1 A 53-year old female went to hospital for low back pain, and the final diagnosis was T12 vertebral fracture. One oval ROI of 400 mm<sup>2</sup> was placed at the middle region of axial image and two oval ROI of 75 mm<sup>2</sup> were placed at bilateral regions of vertebral body also on the pedicle plane.

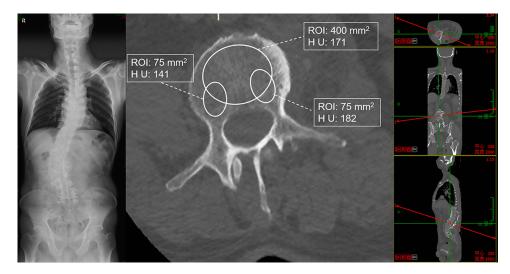


Figure 2 A 59 years old male, whose Cobb angle was 38.5°, went to hospital for low back pain, and the final diagnosis was ADS. The upper end vertebrae was T11. The lower end vertebrae was L5. The HU values on the concave side was higher than that of convex side (182 vs 141).

## Statistical Analysis

All numeric results were recorded as mean  $\pm$  SD. The HU values in non-ADS patients were presented as mean  $\pm$  SD, and the HU values on the left and right sides of lumbar vertebral body in non-ADS patients were compared with paired *t*-test. The chi-squared test was used to compare the prevalence between ADS and non-ADS patients. The HU values on the convex and concave sides of lumbar vertebral body in ADS patients and those on left and right sides in non-ADS patients were compared with two independent samples *t*-test. The HU values were compared among different lumbar vertebral bodies in ADS patients with one-way analysis of variance. The intra-rater and inter-rater reliability of the HU values was evaluated by the two-way intraclass correlation coefficient (ICC). The ICC values were 0–1.00, and values closer to 1.00 represent better reliability. They were defined as very weak (0.01 to 0.19), weak (0.20–0.39), moderate (0.40–0.59), strong (0.60–0.79) or very strong (≥0.080).<sup>18</sup> Moreover, the 95% confidence intervals of the ICC were calculated along

with Cronbach's  $\alpha$ . Statistical analysis was conducted using SPSS for Windows (version 24.0, Statistical Product and service Solutions, Chicago, IL). P value <0.05 was considered statistically significant.

## Results

A total of 162 patients were included in the final analysis. The basic information of ADS and non-ADS patients is summarized in Table 1. There were no significant differences in the terms of age, height, weight, and BMI, between ADS patients and non-ADS patients (p > 0.05). The mean HU values of each vertebrae in ADS patients are lower than that of non-ADS patients, but there is no statistically significant difference between them. Strong to very strong reliability was obtained during the measurements of all the HU values. Both inter-rater and intra-rater ICCs were larger than 0.80, and all the ICC values, 95% CI, and Cronbach's  $\alpha$  are shown in Tables 2 and 3. The strong reliability of HU values suggests a reliable indicator for evaluating the bone mass of vertebrae in ADS patients.

	ADS (n=81)	Non-ADS (n=81)	P value
Age	62.12±7.76	61.77±7.55	0.644
Gender			
Male	36	36	/
Female	45	45	/
Height (cm)	166.58±8.25	167.22±8.57	0.628
Weight (kg)	67.84±11.65	66.38±11.75	0.429
BMI (kg/m <sup>2</sup> )	24.34±3.09	23.62±3.04	0.137
HU value			
LI	127.83±38.32	135.32±38.22	0.215
L2	124.44±40.98	129.07±39.63	0.466
L3	121.99±40.62	126.96±37.83	0.421
L4	127.19±38.16	130.63±39.33	0.572
L5	I 32.09±44.57	138.01±45.15	0.402

Table I Basic Information of ADS and Non-ADS Patients

Note: P value with 2 independent samples t-test.

Abbreviations: BMI, body mass index. HU, Hounsfield unit.

Table 2Inter- and Intra-RaterReliability (Intra-Class Correlation Coefficients, ICC) with 95%Confidence Interval (CI) and Cronbach's Alpha of the HU Values Gained from Non-ADS Patients

Variable	Inter-Rater (n=81)		Intra-rater (n=81)			
	ICC	CI 95%	Cronbach $\alpha$	ICC	CI 95%	Cronbach $\alpha$
LI Total	0.845	[0.769, 0.898]	0.916	0.949	[0.922, 0.967]	0.947
LI Left	0.841	[0.764, 0.895]	0.914	0.953	[0.929, 0.970]	0.967
LI Right	0.833	[0.752, 0.889]	0.909	0.943	[0.913, 0.963]	0.971
L2 Total	0.854	[0.782, 9.904]	0.921	0.954	[0.930, 0.970]	0.976
L2 Left	0.858	[0.787, 0.906]	0.923	0.959	[0.936, 0.973]	0.979
L2 Right	0.855	[0.783, 0.904]	0.922	0.952	[0.927, 0.969]	0.976
L3 Total	0.821	[0.767, 0.887]	0.901	0.949	[0.922, 0.967]	0.974
L3 Left	0.868	[0.803, 0.913]	0.930	0.961	[0.940, 0.975]	0.980
L3 Right	0.850	[0.776, 0.901]	0.919	0.953	[0.928, 0.970]	0.976
L4 Total	0.853	[0.781, 0.903]	0.921	0.955	[0.930, 0.971]	0.977
L4 Left	0.850	[0.776, 0.901]	0.919	0.962	[0.941, 0.975]	0.980
L4 Right	0.882	[0.822, 0.923]	0.937	0.960	[0.938, 0.974]	0.979
L5 Total	0.887	[0.830, 0.926]	0.940	0.964	[0.945, 0.977]	0.982
L5 Left	0.913	[0.868, 0.943]	0.955	0.975	[0.962, 0.984]	0.987
L5 Right	0.912	[0.866, 0.942]	0.954	0.971	[0.955, 0.981]	0.985

Variable	Inter-Rater (n=81)			Intra-Rater (n=81)		
	ICC	CI 95%	Cronbach $\alpha$	ICC	CI 95%	Cronbach $\alpha$
LI Total	0.852	[0.779, 0.902]	0.920	0.949	[0.922, 0.967]	0.974
LI Concave	0.851	[0.778, 0.902]	0.920	0.946	[0.918, 0.965]	0.972
LI Convex	0.876	[0.814, 0.919]	0.934	0.962	[0.942, 0.976]	0.981
L2 Total	0.870	[0.805, 0.915]	0.931	0.956	[0.932, 0.971]	0.977
L2 Concave	0.856	[0.784, 0.905]	0.922	0.948	[0.920, 0.966]	0.973
L2 Convex	0.882	[0.822, 0.922]	0.937	0.963	[0.943, 0.976]	0.981
L3 Total	0.872	[0.808, 0.916]	0.932	0.994	[0.990, 0.996]	0.997
L3 Concave	0.877	[0.815, 0.919]	0.935	0.957	[0.933, 0.972]	0.978
L3 Convex	0.885	[0.827, 0.925]	0.939	0.965	[0.946, 0.977]	0.982
L4 Total	0.856	[0.785, 0.905]	0.922	0.993	[0.989, 0.995]	0.996
L4 Concave	0.866	[0.799, 0.911]	0.928	0.952	[0.927, 0.969]	0.976
L4 Convex	0.835	[0.754, 0.891]	0.910	0.959	[0.937, 0.974]	0.979
L5 Total	0.869	[0.803, 0.914]	0.930	0.963	[0.943, 0.977]	0.981
L5 Concave	0.900	[0.849, 0.935]	0.948	0.966	[0.948, 0.978]	0.983
L5 Convex	0.828	[0.745, 0.886]	0.906	0.964	[0.944, 0.975]	0.984

Table 3 Inter- and Intra-Rater Reliability (Intra-Class Correlation Coefficients, ICC) with 95%Confidence Interval (CI) and Cronbach's Alpha of the HU Values Gained from ADS Patients

The gender-related distribution of osteoporosis diagnosed based on HU value was shown in Table 4. The total prevalence of osteoporosis (OP) in ADS patients was higher than that of non-ADS patients. There is no statistically difference regarding the prevalence of OP in female or male of ADS patients.

As shown in Table 5, the total HU values on the convex side (L1-4) were significantly lower than that on the concave side (p < 0.05). The HU values on the convex side of each vertebral body were significantly lower than those on the

	Prevalence	P value <sup>§</sup>	
	ADS	Non-ADS	]
Female	51.11% (23/45)	37.77% (17/45)	0.203
Male	33.33% (12/36)	25.00% (9/36)	0.437
Total	43.21% (35/81)	32.10% (26/81)	0.144
P value <sup>§</sup>	0.109	0.221	

**Table 4** Gender-Related Distribution of Osteoporosis Diagnosed on HUValues in ADS and Non-ADS Patients

**Notes**: The criterion for osteoporosis diagnosed on CT scan was the HU value of LI < 110. <sup>§</sup>The chi-squared test was used to compare the prevalence.

Table 5 HU Values in Different Side of Each Vertebral Body

	AD	S	Non-ADS		
	Concave	Convex	Left	Right	
LI	129.67±37.49 <sup>§</sup>	105.95±41.51	122.75±37.68	123.79±36.09	
L2	129.99±38.23§	105.83±42.24	128.1±39.81	127.15±39.57	
L3	128.41±41.52 <sup>§</sup>	102.62±42.95	128.98±41.17	3 .  ±40.28	
L4	135.64±39.66 <sup>§</sup>	105.99±40.12	135.69±41.47	138.17±43.04	
L5	129.75±46.17	124.01±43.51	141.74±51.36	144.67±50.58	

Note: §P< 0.05, compared with the convex side, using paired t-test.

concave side in ADS patients (all p < 0.05). While, there were no differences of HU values between the left and right side of each vertebral body in non-ADS patients (p > 0.05). In addition, there were no significant differences in HU values between the concave side in ADS patients and the left/right side in non-ADS patients.

## Discussion

In this study, we performed an in vivo bone mass investigation of ADS. Intra-rater and inter-rater analysis showed that HU methods might be a useful tool to evaluate the vertebral body mass in elder patients according to the strong reliability. The female objectives have a higher prevalence of OP than males in ADS patients and non-ADS patients, and the prevalence of OP in ADS patients was higher than that of non-ADS patients. We here also reported a significant difference in HU values between the convex and concave sides of the lumbar vertebral body on the pedicle plane, which suggests that asymmetrical loading might lead to irregular cancellous bone remodeling of the vertebral body.

The Hounsfield unit value is a relative quantitative measurement of radio density based on the CT images.<sup>19</sup> HU value has been used to evaluate the bone mineral density in recent studies due to obtaining easily from CT images without added cost or radiation.<sup>12</sup> Therefore, many studies have recommended the use of Hounsfield unit (HU) measurements in computed tomography (CT) images to assess BMD, and a positive correlation between HU value and DXA value was also confirmed.<sup>3,10,12,13,20</sup> In addition, several advantages of HU measurements have been proposed, such as avoiding areas with obvious degenerative conditions and removing the interferences of cortical bone when compared with the DXA method. To the best of our knowledge, this study was the first time to assess the reliability of HU value on the vertebral bone mass associated with ADS. In this study, we obtained the HU values of the L1-L5 vertebral body from ADS patients and pared-non-ADS patients. All the results of ICC were larger than 0.8, both in the dimensions of intraclass and inter-class. The previous study has shown HU measurement has excellent inter-and intra-rater reliability in assessing bone quality.<sup>21</sup> However, few studies have focused on the asymmetrical vertebral bone mass distribution in the same vertebrae of ADS patients. The HU measurement could be used as a reproductivity spinal instrumentation for evaluating the bone quality associated with ADS.

Osteoporosis was a common degenerative condition associated with decreased bone mass and strength, which occurred in the aging population. This pathological condition was the most frequent cause of bone fractures (such as spine, proximal femur, distal forearm fractures) in the elderly.<sup>2</sup> The prevalence of OP in the Chinese general aging population ranges from 30% to 40% among females and from 10% to 20% among males.<sup>22,23</sup> While in the current study, we found that the prevalence of OP in ADS and non-ADS female patients was 51.11% and 37.77%, respectively, and the prevalence of OP in ADS and non-ADS male patients was 33.33% and 25.00%, respectively. The results of our study were higher than previous studies. Several reasons may explain this different trait. First, the traditional gold standard method for determining OP, DXA, used in previous studies, usually measured the BMD of lumbar vertebral bodies (L1-L4) and hips. However, the degenerative spine conditions may lead to a lower prevalence of OP using the DXA measurements of lumbar BMD.<sup>8,11</sup> In recent years, HU value detected by CT has been broadly studied to evaluate the bone mineral density because acquiring the HU values by CT can avoid vertebral degenerative changes and cortical bone and directly focus on the cancellous bone of vertebral body, which was more sensitivity than DXA.<sup>10–12</sup> The other reason for the difference may be the lack of physical activity related to spine degenerative diseases. Low back and leg pain, claudication, and neural symptom are common among the ADS and non-ADS patients. These pain conditions may directly lead to decreased physical activity, for that the lack of physical activity was associated with bone mass loss and development of OP. In addition, our study showed that the prevalence of OP in ADS patients was higher than that of non-ADS patients. Zou et al<sup>3</sup> reported that the incidence of OP in ADS patients was higher than in patients with other lumbar degenerative conditions. Furthermore, lumbar scoliosis was a useful clinical marker for OP.<sup>24</sup> A possible cause of this group difference may be that asymmetric loading in ADS patients may enhance the progression of bone resorption.<sup>14</sup>

According to the Wolff's law, it was reported that mechanical loading may influence the bone mass distribution.<sup>25</sup> Bones can adjust their micro-structure to ensure that they were strong enough to withstand the load which they were carrying without too much microdamage or fracture.<sup>26</sup> This functional adaptation to habitual mechanical load led to bone mass changes through the process of bone remodeling.<sup>27</sup> To date, much of the cognizing on mechanics-induced bone

remodeling mainly comes from research focused on young and healthy individuals.<sup>28</sup> However, little information clarified the process of bone remodeling in aging populations who suffered from the asymmetric load. The remodeling progression of bone was different between young and elderly patients' micro-architecture, and it has been shown that aging changed the activity of bone mechano-adaptation.<sup>29</sup> In the current study, we focused on ADS patients whose spine deformity naturally led to asymmetric loading on the vertebral body. There were significant differences in bone mass between the concave and convex sides of the vertebral body on the pedicel plane. The results of this study supported that the process of bone remodeling occurred in the vertebral body of ADS patients. In our previous finite element analysis study.<sup>30</sup> we found that the mechanic load on the concave side was more extensive than that of the convex side in the major curve of the spine. The upper end vertebrae and above suffered from the opposite asymmetric load (the mechanic load at the concave side was larger than that of convex side). Thus, the HU values at the concave side were higher than those of convex side in the major curve vertebrae. Several clinical and experimental studies have confirmed the connection between the mechanical environment and bone structure under sports<sup>31</sup> or bed rest conditions.<sup>32</sup> Moreover, weight-bearing or gymnastic activities can enhance the bone mineral density in human and animal.<sup>33,34</sup> However, our results demonstrated that the static asymmetric load did not enhance the bone mass at the concave side compared with the left/right side of non-ADS patients. Aging-related changes may play an important role in the mechanic-induced process of bone remodeling,<sup>29</sup> but the underlying mechanism has yet to be identified in future research.

For spinal surgeons, OP conditions in ADS patients should be placed in a relatively important position. Pedicle screw fixation is the gold standard method to stabilize the spine and get a rigid fixation to correct the deformity.<sup>35</sup> Non-rigid fusion after fixation surgery often led to non-fusion of the surgical segments, which in turn resulted in screw loosening. The quality of bone tissue was considered as one of the predictors for the occurrence of screw loosening.<sup>36</sup> It was reported that elderly patients had a higher incidence of screw loosening,<sup>36</sup> and screw loosening was more common in osteopenia and osteoporotic conditions.<sup>37</sup> Seller et al<sup>38</sup> reported that bone mineral density was a good predictor for the pullout resistance of pedicle screw. Furthermore, cancellous, not cortical bone, affected the stability and anchorage of pedicle screws.<sup>6</sup> Here, we evaluated the local cancellous bone mass of lumbar vertebrae in ADS patients using the HU values and found that bone mass on the concave side of lumbar vertebrae was higher than that of the convex side. In addition, the prevalence of OP in ADS patients was higher than in non-ADS patients. Thus, special attention should be taken by the surgeon when they are facing elderly ADS patients. Although our study did not include the information about the incidence of screw loosening, it would be important to arouse the attention of surgeons to focus on the OP in ADS patients before surgery. It would be interesting to investigate the cut-off HU values of screw augmentation in the future study.

There are several limitations in our study. Firstly, this retrospective study may cause selection bias. Secondly, the number of patients was relatively small in a single center. A further larger sample prospective study may strengthen the reliability of our findings. Thirdly, the present study did not include the correlation between the incidence of screw loosening and asymmetric bone mass of the vertebral body, and it would be interesting to investigate the relationships in future studies.

In conclusion, HU measurement could be a reliable method in assessing the asymmetrical vertebral bone mass in ADS patients. OP was common among ADS patients compared with non-ADS patients. The static asymmetric load did not enhance the bone mass at the concave side compared with the left/right side of non-ADS patients. The HU values at the concave sides were significantly higher than that of convex side at the lumbar vertebral body on the pedicle plane. Surgeons should carefully evaluate the bone quality of ADS patients to avoid screw loosening.

### Abbreviations

ADS, adult denigrative scoliosis; OP, osteoporosis; CT, computer tomography; HU, Hounsfield unit; ROI, the region of interest.

### **IRB** Approval Statement

This study was approved by the institutional review board of Peking University people's hospital (2021PHB238). The informed consents were obtained from all included patients. This study was also conducted in accordance with the Declaration of Helsinki.

## **Data Sharing Statement**

Yes.

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# Disclosure

The authors have no personal, financial, or institutional interest in any of the drugs, materials, or devices described in this article.

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