

# Factors associated with failure of bony union after conservative treatment of acute cases of unilateral lumbar spondylolysis

Masaki Tatsumura (✉ [tatsumura@md.tsukuba.ac.jp](mailto:tatsumura@md.tsukuba.ac.jp))

Tsukuba university hospital Mito clinical education center / Mito Kyodo General Hospital

<https://orcid.org/0000-0002-3679-9438>

**Hisanori Gamada**

university of Tsukuba

**Shun Okuwaki**

University of Tsukuba

**Fumihiko Eto**

Tsukuba university hospital Mito clinical education and training center, Mito Kyodo general Hospital

**Katsuya Nagashima**

Tsukuba University hospital Mito clinical education and training center, Mito kyodo general Hospital

**Takeshi Ogawa**

Tsukuba University Hospital Mito clinical education and training center, Mito Kyodo general Hospital

**Takeo Mammoto**

Tsukuba University Hospital Mito clinical education and training center, Mito kyodo general Hospital

**Atsushi Hirano**

Tsukuba university hospital Mito clinical education and training center. Mito Kyodo general Hospital

**Masao Koda**

University of Tsukuba

**Masashi Yamazaki**

University of Tsukuba

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## Research article

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# Abstract

**Background:** If bone union is expected, conservative treatment is generally selected for lumbar spondylolysis. However, sometimes conservative treatments are unsuccessful. We sought to determine the factors associated with failure of bony union in acute unilateral lumbar spondylolysis with bone marrow edema including contralateral pseudoarthrosis.

**Methods:** This study targeted unilateral lumbar spondylolysis treated conservatively in high school or younger students. Conservative therapy was continued until the bone marrow edema disappeared on MRI and bone union was investigated by CT. We conducted a univariate analysis of sex, age, pathological stage, lesion level complicating the contralateral bone defect, lesion level, and intercurrent spina bifida occulta, and variables with  $p < 0.1$  were considered in a logistic regression analysis. An item with  $p < 0.05$  was defined as a factor associated with failure of bony union.

**Results:** We found 92 cases of unilateral spondylolysis with bone marrow edema and 66 cases were successfully treated conservatively. Failure of bony union in unilateral lumbar spondylolysis with bone marrow edema was associated with advanced pathological stage ( $p = 0.004$ ), contralateral pseudoarthrosis ( $p < 0.001$ ), and L5 lesion level ( $p = 0.002$ ). The odds ratio was 20.0 (95% CI 3.0–193.9) for advanced pathological stage, 78.8 (95% CI 13–846) for contralateral pseudoarthrosis, and 175 (95% CI 8.5–8192) for L5 lesion level.

**Conclusions:** Conservative therapy aiming at bony union is contraindicated in cases of acute unilateral spondylolysis when the pathological stage is advanced, the lesion level is L5, or there is contralateral pseudoarthrotic spondylolysis.

## Background

Lumbar spondylolysis is a fatigue fracture of pars interarticularis, most of which occur in the 5th lumbar spinal vertebra (L5) [1]. It often occurs in athletes and adolescents and can be fused by conservative treatment [2]. The union rate of lumbar spondylolysis is lower than that of general fatigue fractures. A wide variation in union rate has been reported: 87% in early stage spondylolysis [2], and <56% despite treatment for up to 6 months or more [3]. A possible reason for the substantial variation in union rate between the reports is that there are multiple confounding factors that have a significant impact on bone union. Previous reports revealed several possible factors affecting bone union after conservative treatments for lumbar spondylolysis, including vertebral level [4], stage [4], contralateral condition [4], bilateral spondylolysis [5], and latent spina bifida [6, 7]. Contralateral pseudoarthrosis might be an inhibitory factor for bony union because new spondylolysis tends to occur when a bone defect occurs on the opposite side of the vertebral arch [8]. However, the precise influence of contralateral pseudoarthrosis on bony union rate remains to be elucidated.

Treatment strategies for lumbar spondylolisthesis are controversial. Especially for unilateral surgical treatment, it has been reported that surgery is desirable when symptoms persist for 6 months or longer [9].

In addition, even when surgical treatment is selected, if unilateral lumbar spine separation is present, the surgical results are better than bilateral[10]. On the other hand, if bone fusion can be expected, conservative treatment is also an option. Where bony union cannot be expected by conservative therapy, symptomatic therapy or surgical therapy should be considered rather than continuing exercise or depriving patients of exercise opportunities. Therefore, it is necessary to establish a prognosis of bony union at the time of initial diagnosis of patients with lumbar spondylolysis to determine the treatment strategy. We sought to elucidate the major factors associated with failure of bony union in acute unilateral lumbar spondylolysis with bone marrow edema including contralateral pseudoarthrosis.

## Patients And Methods

### Patients

This study targeted patients with lumbar spondylolysis, which was treated conservatively in our institute from 2014 to 2019. We extracted data from cases in high school students and younger children who were diagnosed with fresh unilateral spondylolysis with bone marrow edema by MRI at the first visit. We excluded cases in which acute lumbar spondylolysis occurred on both sides, and cases in patients who did not consent to conservative therapy.

### Methods

Conservative therapy included using a rigid brace, stopping exercising (including physical class at their school) and physiotherapy until the bone marrow edema disappeared as seen by MRI. Bony union as evaluated with reconstructed sagittal and axial CT images at the time of disappearance of bone marrow edema was the primary outcome of the present study. Failure of bony union was defined as bone discontinuity in reconstructed sagittal and axial CT images.

To determine the factors significantly associated with failure of bony union, we conducted a univariate analysis with bony union at the final visit as the dependent variable and the following items as independent variables: sex, age, pathological stage of spondylolysis in the axial plane of CT [11], contralateral pseudarthrosis, lesion level, and spina bifida occulta. The disease stage was classified as progressive or not (prelysis and early stage) and the level of pathology was separated into L5 and non-L5. Variables with  $p < 0.1$  were included in logistic regression analysis (forward–backward stepwise selection method). Any item with  $p < 0.05$  was defined as a factor associated with bony union failure.

## Result

We included data from 92 patients with cases of unilateral spondylolysis on MRI. Successful bony union was achieved in 66 patients after conservative treatment and the remaining 26 patients failed to obtain bony union, which was diagnosed as pseudarthrosis. The patients included 72 boys and 20 girls with an

average age of 14.4 (range 9–18) years. A prelysis stage was found in 21 patients, early stage was found in 50, and advanced stage in 21. Contralateral pseudarthrosis was found in 25 patients, whereas no pseudarthrosis was found in the remaining 71. The level of spondylolytic lesion was L3 in 9 patients, L4 in 19, and L5 in 64 (Table 1). There were 54 patients with spina bifida occulta.

Univariate analysis revealed that advanced pathological stage ( $p < 0.001$ ), contralateral pseudarthrosis ( $p < 0.001$ ), and L5 lesion level ( $p = 0.001$ ) were significantly associated with bony union failure, whereas age ( $p = 0.40$ ), sex ( $p = 0.85$ ), and spina bifida occulta ( $p = 0.20$ ) were not. The 3 factors showing  $p < 0.1$  in univariate analysis were then analyzed using logistic regression with a forward–backward stepwise selection method. The logistic regression revealed that advanced pathological stage ( $p = 0.004$ , odds ratio (OR) 20.0; 95% CI 3.0–193.9), L5 lesion level ( $p = 0.002$ , OR 175; 95% CI 8.5–8192) and contralateral pseudarthrosis ( $p < 0.001$ , OR 78.8; 95% CI 13–846) (Table 2a) were significantly associated with failure of bony union.

## Discussion

In the present study, we found that lesion level (L5), pathological stage, and contralateral pseudarthrosis were significantly associated with failure of bone union in patients with acute unilateral lumbar spondylolysis and bone marrow edema.

L5 is the most frequent lesion level for lumbar spondylolysis [12]. The union rate of L5 lesions is lower than that for other levels [13, 14]. In the present study, we also found that the union rate at the vertebral level of L5 was significantly lower than that for other vertebral levels. The progressive stage of the disease was previously reported to have a significant negative impact on bony union [15], because according to the stage of disease progression a larger bony defect occurs, which may inhibit bony union.

In addition to the abovementioned known factors that have a significant negative impact on bony union after conservative therapy for lumbar spondylolysis, we demonstrated that unilateral lumbar spondylolysis with contralateral pseudarthrosis is a major risk factor for bony union failure, with an odds ratio of 78.8. Mechanical load can increase significantly when there is pseudarthrosis on the contralateral side of the vertebral arch, resulting in the development of additional fresh spondylolysis [8, 16]. Because contralateral pseudarthrosis disrupts the bone continuity of the vertebral arch, bony union of fresh spondylolysis may be inhibited by the concentration of stress, possibly resulting in a lower rate of bony union (Figure 1).

In the case of fatigue fractures of the extremities, delayed treatment can lead to bony union failure, resulting in a poor prognosis. Therefore, early surgery is indicated according to the prognosis of the stress fractures related to the site and/or stage of the fracture. Surgery may also be considered for stress fractures to shorten rest period and hasten the return to the sports, although they can be healed by conservative treatment. This principal can be indicated for lumbar spondylolysis. As is widely known, conservative treatment is the first choice for fresh lumbar spondylolysis associated with bone marrow edema, and surgery might be considered only when low back pain persists [9, 17].

According to the present results, conservative therapy aimed at bony union should not be applied automatically to all cases. Conservative therapy for acute unilateral spondylolysis should be considered carefully and the patients fully informed of risk factors for bony union failure including advanced stage, a lesion level at L5, and contralateral pseudarthrosis. In such cases, an option is to conduct symptomatic treatment without prolonged external fixation and rest, without aiming for bony union. Another option is to perform direct surgical repair to avoid long rest for attaining bony union earlier.

The limitation of this study is that detailed clinical findings of each case such as the degree of pain could not be confirmed. In addition, it is a retrospective survey and the details of the case where the treatment was dropped out could not be confirmed.

## Conclusion

Conservative therapy aiming at bony union is contraindicated in cases of acute unilateral spondylolysis when the pathological stage is advanced, the lesion level is L5, or there is contralateral pseudoarthrotic spondylolysis.

## Abbreviations

the 5th lumbar spinal vertebra : L5

## Declarations

**Ethics approval and Consent to participate:** All procedures including review of patient record used in this research were approved by the institutional review board of Tsukuba University Hospital Mito Clinical Education and Training Center/ Mito Kyodo General Hospital (No.17-01). Verbal informed consents were obtained from all patients and their parent to participate in this study. An IRB approve the study and the procedure outlined for the verbal consent obtainment.

**Consent for Publication:** Written informed consent for publication was obtained from the patients included in this study.

**Availability of data and material:** The datasets generated during and analyzed during the current study are available from the corresponding author on reasonable request.

**Competing Interests:** The authors declare that there are no relevant competing interest.

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**Author contributions:** MT conceived and designed the study. HG and SO collated and analyzed the data. FE, KN, TO, TM and AH treated the patients. MT, HG and MK drafted the manuscript. MY critically revised the draft manuscript. All the authors approved the final version submitted for publication.

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## Tables

**Table 1: Patient Demographics**

Patient Demographics (n = 92)		
Treatment Result	Union	66
	Pseudoarthrosis	26
Sex	Male	72
	Female	20
Age at First Visit (years old)		14.4 (9-18)
Pathological Stage	Prelysis	21
	early	50
	progressive	21
Contralateral Condition	normal	67
	bone defect	25
Vertebral Level of The Lesion	L3	9
	L4	19
	L5	64
intercurrent SBO	with	54
	without	38

Table 2a: Univariate analysis

Univariate analysis	p-value (#: p<0.1)
Age	0.4
Sex	0.85
L5 or non-L5	0.001 <sup>#</sup>
Spina Bifida Occulta	0.2
CT Pathological Stage	<0.001 <sup>#</sup>
Contralateral Pseudoarthrosis	<0.001 <sup>#</sup>

Table 2b: Stepwise Logistic Regression

Stepwise Logistic Regression	p-value (*: p<0.05)	odds ratio (95% CI)
L5 or non-L5	0.002*	175 (8.5-8192)
CT Pathological Stage	0.004*	20.0 (3.0-193.9)
Contralateral Pseudoarthrosis	<0.001*	78.8 (13-846)

Figures

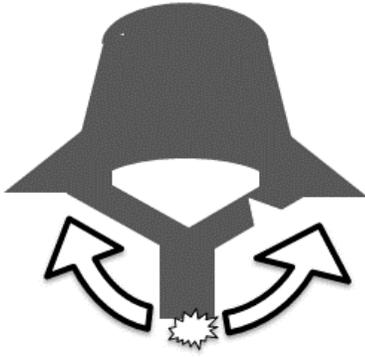


Figure 1a) mechanical stress with contralateral normal

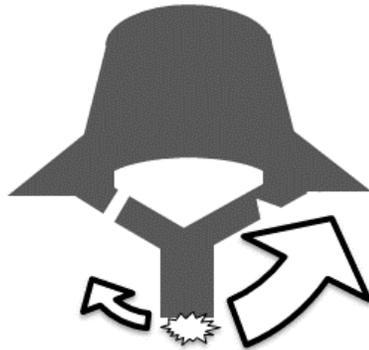


Figure 1b) mechanical stress with contralateral pseudoarthrosis

## Figure 1

The direction and magnitude of the mechanical load are indicated by arrows. (a) In the normal case without contralateral spondylolysis, the mechanical stress from the posterior is evenly distributed to the left and right vertebral arches. (b) When a bone defect occurs due to terminal stage spondylolysis, the mechanical stress from the posterior is concentrated at the site of acute spondylolysis where bone continuity remains. Therefore, stability is not attained and the union rate is low.