1	The Prevalence and Risk Factors for Delayed Union of the Superior Pubic Ramus at 1 Year After
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24	https://link.springer.com/article/10.1007/s00264-017-3706-9
25	
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27	Disclosure statement: No benefits in any form have been received or will be received from a
28	commercial party related directly or indirectly to the subject of this article. The authors received no
29	financial support for the research, authorship, and/or publication of this article.
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32 Abstract

33 Purpose: Curved periacetabular osteotomy (CPO) has been developed for the treatment of acetabular 34 dysplasia. While several studies have reported its good clinical results, the complications of CPO 35 include delayed union and nonunion of the superior pubic ramus. The purpose of this study is to 36 investigate the prevalence of delayed union of the pubis 1 year after CPO, and to determine the risk 37 factors for this complication.

38 Methods: The study examined 113 hips that underwent CPO between 2008 and 2012. Delayed union 39 was assessed based on the anteroposterior radiography 1 year after CPO. A superior pubic ramus union 40 group (U group) and a delayed union group (D group) were retrospectively compared regarding patient 41 characteristics, clinical evaluations, and radiographic parameters.

- 42 Results: Delayed union rate was 16.8%. The D group contained a significantly greater proportion of
- 43 smokers (p < 0.001). The gap at the pubic osteotomy site on CT coronal images was significantly larger
- 44 in the D group (p < 0.001), and the cut-off value for the risk of nonunion was a larger than 5.1 mm.
- 45 Multivariate regression analysis indicated that smoking (OR 10.7, 95% CI 2.1–55.4) and a gap at the
- 46 superior pubic ramus > 5.1 mm (OR 16.5, 95% CI 3.7–73.7) were significantly associated with delayed
- 47 union as independent risk factors.
- 48 Conclusion: The prevalence of delayed union 1 year after CPO was 16.8%. Smoking and a gap larger
- 49 than 5.1 mm at the pubic osteotomy site are risk factors for delayed union after CPO.

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51 Key words
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periacetabular osteotomy; complication; delayed union and nonunion; superior pubic ramus; risk factor
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55 Introduction

56Acetabular dysplasia of the hip is known to cause of osteoarthritis of the hip [1], which accounts for 57more than 70% of hip osteoarthritis cases in Japan [2]. Periacetabular osteotomy (PAO) has been 58developed as a surgical treatment for symptomatic acetabular dysplasia, and good clinical results have 59been reported [3-9]. However, the complications of PAO include superior pubic ramus nonunion, inferior 60 pubic ramus stress fracture, posterior column fracture, and lateral cutaneous nerve injuries [3–15]. Several 61 studies have reported the prevalence of delayed union and nonunion of the superior pubic ramus after 62 PAO to be around 1.0 to 17.0% [7-9,11-14]. However, all these reports did not mention the evaluation 63 time. In addition, to our knowledge, no reports have described the risk factors for this complication. The 64 purpose of this study was to assess the prevalence of delayed union of the superior pubic ramus on 65 radiographs at 1 year after CPO, and also to investigate the risk factors correlated with delayed union.

66

67 Materials and Methods

68 Between the period of April 2008 and January 2012, CPO was performed in consecutive 128 hips in 69 120 patients for the treatment of acetabular dysplasia of the hip. To assess the osteotomy sites, pelvic 70computed tomography (CT) scans were routinely performed at 1 week postoperatively, and 71anteroposterior (AP) and false-profile radiographs of the hip were assessed at preoperative and 1 year 72postoperatively. CT scans could not be performed in seven hips in seven patients, and an additional eight 73hips in eight patients were lost to follow-up. Finally, 113 hips in 105 patients were included in this study 74(follow-up rate 88.3%). The mean age of the patients at the time of surgery was 38.3 (range, 15–68) 75years, and the 113 hips comprised 10 hips in male patients and 103 hips in female patients. The mean 76follow-up duration was 51.2 (range, 12.0-81.0) months. The mean period until radiographic follow-up at 1 year postoperatively was 12.3 (range, 10.7–13.4) months. The mean body mass index (BMI) was 22.1
(range, 16.8–31.7) kg/m². The preoperative Tönnis grades [16] were grade 0 in 72 hips, grade 1 in 40
hips, and grade 2 in 1 hip.

80 CPO, which is a modified Bernese PAO developed by Ganz et al. [3,4] has been performed at our 81 hospital since 1995. The surgical indications for CPO included acetabular dysplasia associated with hip 82 pain and discomfort that caused some limitation of daily activities, lateral center-edge angle (LCEA) [17] 83 of less than 20° on AP radiographs, and improvement in joint congruency in the abducted position on AP 84 radiographs. All surgical procedures were performed by three surgeons. CPO was undertaken using the 85 modified Smith-Peterson approach. The c-shaped osteotomy was performed from the proximal part of the 86 anterior inferior iliac spine to the distal part of the quadrilateral surface. Osteotomy of the superior pubic 87 ramus was carried out just medial to the iliopubic eminence, and was performed at an inclination of 30° to 88 the horizontal line for medialization of the femoral head [18]. Bone grafting was not performed at the 89 osteotomy area of the superior pubic ramus. Active motion exercises were initiated on the first 90 postoperative day. Partial weight-bearing (10 kg) was allowed on the third postoperative day, and full 91 weight-bearing was allowed from 8 weeks postoperatively.

92 Clinical evaluations were performed using the Harris hip score (HHS) both preoperatively and at 1 year 93 postoperatively. The pain component of the HHS was also investigated. The lifestyle habits and 94 comorbidities of the patients were examined by reading their electronic charts retrospectively. Patients 95 who were smokers at the time of CPO or had a history of smoking were grouped as smokers. Patients 96 with a daily alcohol drinking habit were grouped as alcohol drinkers.

97 Radiographic evaluations included pre- and postoperative LCEA, acetabular roof obliquity (ARO) [19],

98 and head lateralization index (HLI) [20]. The anterior center-edge angle (ACEA) [21] was assessed on

99 false-profile radiographs. The amounts of change of transfer of the acetabular fragment were assessed by 100 differences in the pre- and postoperative measurements of the LCEA, ARO and ACEA. At 1 week 101 postoperatively, a pelvic CT scan was routinely performed to examine the osteotomy area 102 (TSX-101A/HA; Toshiba Medical Systems, Tochigi, Japan). CT coronal imaging was used to measure the 103 gap at the pubic ramus osteotomy site, and the narrowest gap on all the slices was taken as the smallest 104 gap (Fig. 1).

Patients showing no bone continuity on AP radiographs at 1 year postoperatively were assigned to the delayed union group (D group), while patients showing bone continuity were assigned to the union group (U group). Bone continuity on AP radiographs was determined by two investigators (A.M. and S.A.). Among the cases judged to have delayed union, cases in which bone union was not obtained even at the time of the final follow-up were regarded as a nonunion. In the case of delayed union, the presence of pubic tenderness was investigated by reading their electronic charts retrospectively.

111

112 Statistical Analysis

113 The Mann–Whitney U test was used to compare the demographic data, clinical scores, and radiographic 114 data between the D and U groups. The chi-squared test was used to assess differences in clinical factors 115 such as sex, lifestyle habits, and comorbidities. A cut-off value for the gap distance at the superior pubic 116 ramus osteotomy site that optimized sensitivity and specificity was identified using a receiver-operating 117 characteristic curve. A multivariate regression analysis was performed to identify factors related to 118 delayed union of the superior pubic ramus. Statistical analyses were performed using SPSS ver. 20.0 119 (IBM Inc., Armonk, New York, USA). Statistical significance was defined as p < 0.05.

120

121 **Results**

122Delayed union was observed in 19 hips (16.8%; D group). The clinical data for the D and U groups are 123shown in Table 1. There were no significant differences between the two groups in mean age at the time 124of surgery, sex, mean follow-up duration, mean time to radiographic follow-up, and mean BMI. The mean 125score for the pain component of the HHS at 1 year postoperatively was significantly greater in the U 126group than the D group (p = 0.005). Pubic tenderness was present in 21.0% (four of 19 hips) in the D 127 group. Hip osteoarthritis had not progressed at 1 year postoperatively in any of the D group patients, 128 whereas it had progressed from Tönnis Grade1 to Grade 2 in one hip and from Tönnis Grade 2 to Grade 3 129 in another hip in the U group. 130 The lifestyle habits and comorbidities in the two groups are shown in Table 2. Smokers were present at a 131significantly higher rate in the D group (p = 0.025). 132 The radiographic evaluations are shown in Table 3. Delayed union was observed in 19 hips (16.8%) at 1 133 year postoperatively. Among these 19 hips with delayed union, five hips obtained bone union at an 134average of 24 months postoperatively (Fig. 2). However, the remaining 14 hips showed nonunion of the 135pubis even at the final follow-up, and mean follow up duration was 48.7 (range 24.0 to 62.9) months (Fig. 136 3). Posterior column fracture rate did not show a significant difference between the two groups (p =137 0.174). Compared with the U group, the D group had a significantly smaller postoperative ARO (p = 138 (0.009), and a significantly greater amount of correction of the ARO (p = 0.007). The gap at the pubic 139 osteotomy site on CT coronal images was significantly larger in the D group (p < 0.001), and the cut-off 140 value for the risk of nonunion was a larger than 5.1 mm, with a sensitivity of 78.9% and a specificity of 141 76.6%.

142 The results of the multivariate regression analysis are shown in Table 4. The multivariate regression

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analysis indicated that smoking (odds ratio (OR) 10.701, 95% confidence interval (CI) 2.06-55.44) and 144a gap at the superior pubic ramus > 5.1 mm (OR 16.5, 95% CI 3.7–73.7) were significantly associated 145with delayed union as independent risk factors.

146

147Discussion

148PAO has been performed as a surgical treatment for symptomatic acetabular dysplasia, and good short-149 and mid-term outcomes have been reported [3-9]. However, some reports have described that the 150incidence of superior pubic ramus nonunion is reportedly 1.0 to 17.0% [7-9,11-14], although these 151reports did not mention the evaluation time. In this study, the incidence of delayed union of the superior 152pubic ramus was 16.8% at 1 year postoperatively. Among these cases of delayed union, five of 19 153 patients obtained bone fusion at an average of 24 months postoperatively. As a result, the nonunion rate 154was 12.4% at the final follow up. This rate was almost equivalent to the previously reported incidence. 155Several studies have reported that pubic ramus nonunion is asymptomatic in most patients and has no 156effect on the clinical outcome [7,13]. However, symptomatic pubic ramus nonunion rates were reported 157to be 1% and 4% by Peters et al. [11] and Clohisy et al. [9], respectively. Furthermore, Siebenrock et al. 158[12] and Clohisy et al. [9] reported that surgery was necessary for nonunion in 1% and 4% of patients, 159respectively. In the present study, the mean score for the pain component of the postoperative HHS was 160 significantly lower in the D group. Since hip osteoarthritis had not progressed at 1 year postoperatively

161 in any of the hips in the D group, the lower score for the pain component in the D group seems to be 162 related to the delayed union itself rather than the hip osteoarthritis. The present results indicate that

163 although bone fusion may be obtained in the long-term in cases of delayed union, delayed union may

164 cause early postoperative pain and lead to a decrease in the activities of daily living. 165 Pubic delayed union might be related to several factors such as severe hip dysplasia, large acetabular 166correction, posterior column fracture, patient age, presence of bone grafting, and early rehabilitation. In 167 our study, there were a significantly greater proportion of smokers in the D group. Regarding the effects 168 of smoking on the musculoskeletal system, nicotine and carbon monoxide are known to reduce oxygen 169supply and peripheral blood flow to tissues, and to reduce bone metabolic activity [22]. Brown et al. [23] 170 reported that smoking reduces oxygen levels in the blood and increases the rate of nonunion. Andersen 171 et al. [24] reported that smoking has negative effects on fusion and patient satisfaction after spinal fusion 172procedures. A smoking cessation program prior to surgery is reportedly effective in decreasing the 173 incidence of postoperative complications caused by smoking [25]. In the present study, the OR for 174delayed union of the superior pubic ramus was 10.7 times higher in smokers, indicating that guidance on 175quitting smoking prior to surgery is necessary to decrease the risk of delayed union.

176 Some studies have reported on the relationship between severity of acetabular dysplasia and pubic 177 ramus nonunion. Clohisy et al. [8] reported a pubic ramus nonunion rate of 12.0% hips with severe 178 dysplasia that underwent Bernese PAO. Peters et al. [11] reported a pubic ramus nonunion rate of 12.0% 179 treated by PAO, and most of these cases involved hips with large acetabular corrections. In our study, the 180 factors associated with the preoperative severity of acetabular dysplasia did not significantly differ 181 between the two groups. However, the amounts of correction of the ARO was significantly greater in the 182D group, indicating that a large amount of acetabular correction may have been involved in delayed 183 union.

To further evaluate the transfer of the acetabular fragment, the gap at the superior pubic ramus osteotomy site was measured on CT images. The mean gap at the osteotomy site was 6.9 mm in the D group, was significantly larger than that in the U group, and the cut-off value for the risk of delayed

187 union was 5.1 mm, the OR was 16.4 times higher. Thus, to prevent delayed union of the superior pubic 188 ramus, the osteotomy site must have a large contact area. Osteotomy of the superior pubic ramus was 189 made at the location just medial to the iliopubic eminence in the present study, and so it will be 190 displaced further with rotation of the acetabular fragment. Performing the pubic osteotomy at the medial 191aspect, where the bone is narrower and the contact area is smaller, may make it more difficult to heal 192 compared with osteotomy performed at the more lateral aspect of the pubis. In our hospital, we have 193 performed superior pubic osteotomy using a flap form since 2013, and have transplanted a bone graft 194 into the superior pubic osteotomy site to promote bone union. 195This study had several limitations. The first limitation was the low follow up rate of 88.3%. As a result, 196 the sample size was insufficient, and there was a large difference in the number of patients in the two 197 groups. The second limitation was that delayed union of the pubis was assessed only based on 198 radiographic evaluation. The actual rate of delayed union of the superior pubic ramus may show higher 199 percentage if CT scans were used for the evaluation. Third, the correlation between the pack-year history 200of smoking and delayed union was not examined. An increasing pack-year history of smoking may have 201resulted in an increased risk of delayed union. Fourth, the radiographic evaluation of transfer of the 202 acetabular fragment was assessed in two dimensions only. It may be difficult to measure the amount of 203acetabular fragment transfer on pelvic radiography; this measurement requires a three-dimensional 204assessment. 205206 Conclusions 207 In summary, surgeons must be aware that delayed union and nonunion of the osteotomy site at the pubic

208 ramus can sometimes occur after CPO. The rate of delayed union of the superior pubic ramus on AP

209	radiographs	at 1	year	after	СРО	was	16.8%.	Smoking	and	a gap	larger	than	5.1	mm	at	the	pubis
210	osteotomy si	te ar	e risk	factor	s for c	lelaye	ed union	after CPO).								

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212 Conflict of Interest: On behalf of all authors, the corresponding author states that there is no conflict of

213 interest.

214	Ethical approval: All procedures performed in studies involving human participants were in
215	accordance with the ethical standards of the institutional and/or national research committee and with
216	the 1964 Helsinki declaration and its later amendments or comparable ethical standards.
217	Informed consent: Informed consent was obtained from all individual participants included in the study.
218	

219 Funding: No external funding was received for this study.

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Figure captions

Fig. 1. Measurement of the smallest gap (yeallow line) at the superior pubic ramus osteotomy site on a coronal computed tomography image.

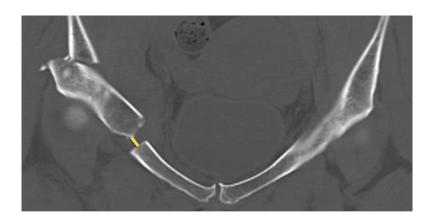


Fig. 2. Radiographs of a 35-year-old female who underwent curved periacetabular osteotomy for hip dysplasia. a: Preoperative anteroposterior radiograph; b: Postoperative anteroposterior radiograph; c: There was no bone continuity of the superior public ramus on anteroposterior radiographs taken 1 year postoperatively; d: Bone union was obtained at 24 months postoperatively.

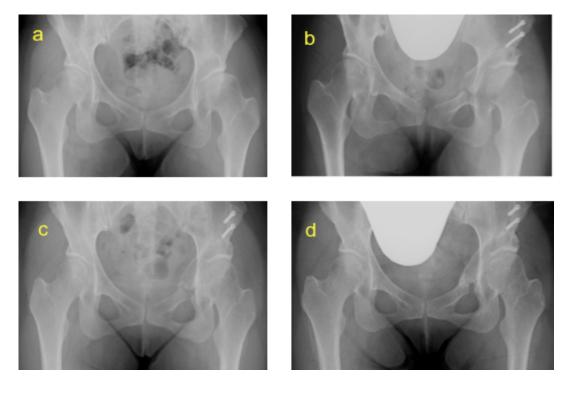
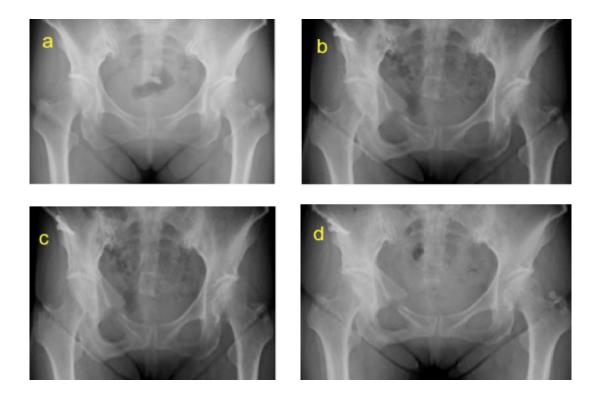


Fig. 3. Radiographs of a 66-year-old female patient who underwent curved periacetabular osteotomy for hip dysplasia. a: Preoperative anteroposterior radiograph; b: Postoperative anteroposterior radiograph; c: There was no bone continuity of the superior pubic ramus on anteroposterior radiographs taken 1 year postoperatively; d: There was nonunion of the superior pubic ramus at final follow-up (5.2 years postoperatively)



Table

Table 1. Demographic Data in the Delayed Union Group and the Union Group

	Delayed union group $(n = 19)$	Union group (n = 94)	P value
Sex, M:F (no. of hips)	0:19	10:84	0.138
Age at surgery (y)	41.0 (20 to 66)	37.7 (14 to 68)	0.184
Follow-up duration (mo)	49.2 (12.1 to 72.0)	51.6 (12.0 to 81.0)	0.673
Time to radiographic follow-up (mo)	12.3 (10.7 to 13.2)	12.2 (10.8 to 13.4)	0.686
Body mass index (kg/m ²)	21.7 (19.1 to 25.9)	22.2 (16.8 to 31.7)	0.416
HHS (points)			
Preoperative	77.0 ± 8.2	75.8 ± 11.1	0.160
Postoperative	91.2 ± 8.3	93.8 ± 5.5	0.251
Pain component of the HHS (points)			
Preoperative	25.3 ± 5.1	15.6 ± 7.3	0.394
Postoperative	36.2 ± 7.3	39.9 ± 6.2	0.005
Tönnis grade†			
Preoperative	10/9/0/0	62/31/1/0	
Postoperative	10/9/0/0	62/30/1/1	

The values are given as the mean \pm standard deviation (range).

HHS: Harris Hip Score. F: female, M: male.

† no. of hips with Tönnis grade 0/ grade1/ grade2/ grade3

	Delayed union group	Union group	P value*
	(n = 19)	(n = 94)	1 vulue
Smokers	7 (36.8%)	12 (12.7%)	0.025
Alcohol drinker	3 (15.7%)	7 (7.4%)	0.370
BMI > 25 kg/m ²	2 (10.5%)	16 (17.0%)	0.338
Liver disease	0 (0%)	3 (3.1%)	0.554
Heart disease	0 (0%)	4 (4.2%)	0.453
Diabetes	0 (0%)	2 (2.1%)	0.676
Depression	0 (0%)	2 (2.1%)	0.676

Table 2 Preoperative Patient-Reported Lifestyle Habits and	Comorbidities in the Delayed Union Group and the Union Group	
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† Calculated using chi-squared tests. Smokers included those who were smokers at the time of surgery and those with a prior history of smoking. Alcohol drinkers included those who drank alcohol daily. BMI: body mass index. Table 3. Radiographic Evaluations in the Delayed Union Group and the Union Group

	Delayed union group $(n = 19)$	Union group $(n = 94)$	P value
	(1 - 19)	(11 - 94)	
Lateral center-edge angle (°)			
Preoperative	9.1 ± 7.4 (-5 to 17)	9.6 ± 5.6 (-6 to 26)	0.362
Postoperative	$32.9 \pm 5.4 (25 \text{ to } 44)$	$32.4 \pm 5.4 (20 \text{ to } 48)$	0.521
Correction	23.9 ± 8.4 (13 to 39)	$22.7 \pm 6.4 (1 \text{ to } 38)$	0.127
Acetabular roof obliquity (°)			
Preoperative	$19.6 \pm 6.0 (10 \text{ to } 29.4)$	$17.3 \pm 5.8 (2 \text{ to } 34)$	0.152
Postoperative	1.4 ± 1.9 (-1.7 to 6.7)	3.1 ± 2.8 (-4 to 12)	0.009
Correction	$18.1 \pm 5.8 \ (9 \text{ to } 29.9)$	14.1 ± 5.3 (-1.3 to 30.3)	0.007
Anterior center-edge angle (°)			
Preoperative	20.5 ± 12.6 (-4.7 to 49.7)	18.5 ± 10.2 (-5.8 to 36.2)	0.651
Postoperative	$36.3 \pm 8.9 (29 \text{ to } 59.1)$	35.1 ± 8.2 (20 to 57.2)	0.499
Correction	$15.5 \pm 6.9 (7.2 \text{ to } 34.6)$	$16.3 \pm 8.3 (0.9 \text{ to } 40)$	0.525
Head lateralization index			
Preoperative	$0.55 \pm 0.05 \ (0.47 \text{ to } 0.67)$	$0.57 \pm 0.06 \ (0.47 \text{ to } 0.73)$	0.196
Postoperative	$0.50 \pm 0.06 \ (0.39 \text{ to } 0.62)$	$0.53 \pm 0.07 \ (0.40 \text{ to } 0.72)$	0.057
Distance of the pubic ramus osteotomy site (mm)	6.9 ± 3.2 (2.5 to 15.9)	$4.0 \pm 2.3 \ (0 \text{ to } 15.6)$	< 0.001
Posterior column fracture (n, %)	3 (15.8 %)	6 (6.4 %)	0.174

The values are given as the mean \pm standard deviation (range).

Table 4. Multivariate Regression Analysis for Risk Factors Predicting Delayed Union of the Superior Pubic Ramus

Risk factors	OR	95% CI	P value
Postoperative ARO	0.689	0.51-0.94	0.017
Correction of the ARO	1.165	1.04-1.31	0.009
Smoking	10.701	2.06-55.44	0.005
Distance at the pubic ramus osteotomy site > 5.1mm	16.466	3.67-73.72	0.001

OR: odds ratio, CI: confidence interval.