

# Incidence and Risk Factors for Pregnancy-Related de Quervain's Tenosynovitis in South Korea: A Population-Based Epidemiologic Study

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**Background:** Although pregnant or lactating women have been recognized to be predisposed to de Quervain's tenosynovitis (DQT), there is a lack of epidemiologic evidence. The purpose of this study was to estimate the nationwide incidence of pregnancy-related DQT (PRDQT) and to analyze risk factors using the Korean National Health Insurance (NHI) database.

**Methods:** A retrospective epidemiologic study of pregnant women in South Korea from 2013 to 2017 was conducted using the NHI claims database. Using corresponding diagnostic codes, we identified women diagnosed with DQT during pregnancy or the postpartum period. We calculated the cumulative incidence and analyzed risk factors such as demographics, pregnancy type, delivery method, gestational complications, and comorbidities using multivariate logistic regression analysis.

**Results:** Between 2013 and 2017, 34,342 patients with PRDQT were identified among 1,601,501 pregnant women, representing a cumulative incidence of approximately 2.1%. Age  $\geq$  30 years, multiple gestation, cesarean delivery, hypertensive disorders in pregnancy, and underlying rheumatoid arthritis were all identified as significant risk factors for the occurrence of PRDQT, whereas diabetic disorders in pregnancy and underlying diabetes mellitus were not.

**Conclusions:** In South Korea, PRDQT was found to affect approximately 2.1 out of 100 pregnant women between 2013 and 2017. The incidence and risk factors identified in this study can be used for clinical consultations and prediction, as well as for development of national health policies.

Keywords: Wrist joint, De quervain disease, Pregnancy complications, Epidemiology

De Quervain's tenosynovitis (DQT) is a stenosing tenosynovitis of the first extensor compartment of the wrist. The first dorsal compartment is the fibro-osseous tunnel just proximal to the radial styloid, through which the abductor pollicis longus (APL) and extensor pollicis brevis (EPB) tendons glide. Attritional force secondary to repetitive friction of tendons causes swelling and thickening of the

Received March 17, 2022; Revised May 10, 2022; Accepted June 6, 2022 Correspondence to: Kee Jeong Bae, MD Department of Orthopedic Surgery, Seoul National University Boramae Medical Center, 20 Boramae-ro 5-gil, Dongjak-gu, Seoul 07061, Korea Tel: +82-2-870-2319, Fax: +82-2-831-0714 E-mail: grant903@gmail.com extensor retinaculum that covers the first dorsal compartment, resulting in the narrowing of the fibro-osseous tunnel. Resisted gliding of tendons though the narrowed tunnel provokes the pain and decreased motion.<sup>1)</sup>

According to previous reports, DQT occurs up to six times more frequently in women than men and is associated with the dominant hand use during middle age.<sup>1-3)</sup> The most common DQT patient is a woman in her 50s or 60s who overuses her wrist or thumb, performing house-hold tasks or occupations that require repetitive motions such as typing and lifting.<sup>1)</sup> Another significantly younger cohort of patients who are diagnosed with DQT consists of pregnant or lactating women, who are classified as having pregnancy-related DQT (PRDQT).<sup>3-7)</sup>

Although it has been recognized that PRDQT is

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Clinics in Orthopedic Surgery • pISSN 2005-291X eISSN 2005-4408

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usually self-limiting following delivery or cessation of breastfeeding, there is a paucity of epidemiologic evidence for PRDQT patients. As only a few case series involving a small number of patients have been reported, there is currently a lack of information necessary to consult pregnant or lactating women complaining of their radial wrist pain or to predict the occurrence of PRDQT.<sup>4-6)</sup> Since the occurrence of PRDQT is empirically considered to be much rarer than DQT, investigations involving a large number of pregnant or lactating women are anticipated to provide more relevant findings. Due to the fact that almost all South Korean nationals are covered by the Korean National Health Insurance (NHI) system, analyzing the NHI database enables epidemiological studies on a large population, providing an accurate estimate of this uncommon condition in South Korea. Therefore, in this study, we used the NHI database to estimate the nationwide incidence of PRDQT in South Korea and to analyze risk factors.

#### **METHODS**

The Institutional Review Board of the Seoul National University Boramae Medical Center approved this study protocol as an exemption from review in compliance with the exemption criteria (No. 07-2022-14). Informed consent was waived because this study used only publicly available de-identified data.

#### **Data Sources**

A 5-year retrospective population-based epidemiologic study from 2013 to 2017 was conducted using the Korean NHI claims database. Almost all healthcare providers in South Korea are required to submit claims data to the NHI, including each patient's diagnosis and medical costs, in order to be reimbursed by the government for medical expenses. As a result, except for procedures such as cosmetic surgery that are not covered by the NHI system, this centralized claims database covers over 50 million South Korean nationals, providing nearly all information about the nationwide volume and burden of any specific disease or condition. Several epidemiological studies on pregnant women in South Korea have already been conducted using this centralized database.<sup>8,9)</sup>

#### **Data Collection**

The NHI claims database contains de-identified information on all insurance claims, including demographics, diagnostic codes based on the 10th revision of the International Classification of Diseases (ICD-10) system, medical services provided, and medical expenses incurred. We identified pregnant women aged 15 to 49 years who gave birth between 2013 and 2017 using the ICD-10 codes for delivery (O80, O81, O82, O83, and O84). We excluded those who gave birth at their ages of < 15 or  $\ge 50$  years, because they are extremely rare (less than 0.001% of total pregnancies) according to Korean National Statistics.<sup>10)</sup> The pregnancy period was defined as the 10 months prior to a delivery date and the postpartum period as the following 1 year. Although the delayed postpartum period is frequently defined in obstetrics as 6 months following delivery, we defined it as 1 year following childbirth, given that the average duration of lactation for South Korean mothers is approximately 13 months according to Korean National Statistics.<sup>10,11)</sup> The following criteria were used to identify patients with PRDQT: (1) those who gave birth between 2013 and 2017, (2) those who had at least one claim under the ICD-10 code corresponding to DQT (M65.4) during the pregnant or postpartum period, and (3) those who did not have any claim for DQT prior to pregnancy (Fig. 1). To exclude women who had been diagnosed with DQT prior to the study period, we excluded those who had any claims data related to DQT from 2011 to pregnancy. A woman who had two or more deliveries during the study period was counted as one mother.

We collected data on demographics, pregnancy type, delivery method, gestational complications, and comorbidities such as rheumatoid arthritis (RA) and diabetes mellitus (DM) using the corresponding ICD-10 codes. According to pregnancy type, all subjects were classified as single gestation (O80, O81, O82, and O83) or multiple gestation (O84). They were classified according to their delivery method as vaginal (O80, O81, O83, O84.0, O84.1, O84.8, and O84.9) or cesarean section (O82 and O84.2) delivery. Data on the presence of two major gestational



**Fig. 1.** Criteria for identifying patients with pregnancy-related de Quervain's tenosynovitis in the Korean National Health Insurance claims database.

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complications were collected: hypertensive disorders of pregnancy (O11, O13, O14, O15, and O16) and diabetic disorders of pregnancy (O24). Hypertensive disorders of pregnancy included disorders such as preeclampsia superimposed on chronic hypertension, pregnancy-induced hypertension, preeclampsia, and eclampsia. Regarding comorbidities, the presence of RA (M05.8, M05.9, M06.0, M06.8, and M06.9) and DM (E10, E11, E12, E13, and E14) were determined, respectively. Table 1 summarizes all ICD-10 codes that were used to collect data.

#### **Data Analysis**

The cumulative incidence of PRDQT was calculated by dividing the number of patients with PRDQT by the total number of women who delivered during the study period. Additionally, the cumulative incidence for each 5-year age group was calculated. To determine the risk factors for PRDQT, all pregnant women were divided into two

(	Condition	ICD-10 code	Description
Delivery		080	Single spontaneous delivery
		081	Single delivery by forceps and vacuum extractor
		082	Single delivery by caesarean section
		083	Other assisted single delivery
		084.0	Multiple delivery, all spontaneous
		084.1	Multiple delivery, all by forceps and vacuum extractor
		084.2	Multiple delivery, all by caesarean section
		084.8	Other multiple delivery
		084.9	Multiple delivery, unspecified
de Quervain's tenosynovitis		M65.4	Radial styloid tenosynovitis (de Quervain)
Gestational complication	Hypertensive disorders of pregnancy	011	Pre-eclampsia superimposed on chronic hypertension
		013	Gestational (pregnancy-induced) hypertension
		014	Pre-eclampsia
		015	Eclampsia
		016	Unspecified maternal hypertension
	Diabetic disorder of pregnancy	024	Diabetes mellitus in pregnancy
Comorbidity	Rheumatoid arthritis	M05.8	Other seropositive rheumatoid arthritis
		M05.9	Seropositive rheumatoid arthritis, unspecified
		M06.0	Seronegative rheumatoid arthritis
		M06.8	Other specified rheumatoid arthritis
		M06.9	Rheumatoid arthritis, unspecified
	Diabetes mellitus	E10	Type 1 diabetes mellitus
		E11	Type 2 diabetes mellitus
		E12	Malnutrition-related diabetes mellitus
		E13	Other specified diabetes mellitus
		E14	Unspecified diabetes mellitus

ICD-10: the 10th revision of the International Classification of Diseases.

groups: those with PRDQT (PRDQT group) and those without PRDQT (non-PRDQT group). The chi-square test was used to compare categorical variables between the two groups. To analyze age as a categorical variable, subjects were divided into two groups: those aged < 30 years and those aged  $\geq$  30 years. A multivariate logistic regression analysis was conducted on variables that had been determined to be significant in univariate analyses. SAS software ver. 9.3 (SAS Institute Inc., Cary, NC, USA) was used to analyze all data, with a *p*-value < 0.05 considered statistically significant.

## RESULTS

Between 2013 and 2017, the total number of women who gave birth was 1,601,501. During this time period, the total number of women with PRDQT was 34,342. As a result, the cumulative incidence of PRDQT can be estimated to be 2.1%. The cumulative incidence was 1.6% for those aged 15–19 years; 1.8% for those aged 20–24 years; 2.1% for those aged 30–34 years; 2.4% for those aged 35–39 years; 2.5% for those aged 40–44 years; and 3.9% for those aged 45–49 years (Table 2).

The univariate analysis revealed that the PRDQT group had a significantly greater number of patients aged  $\geq$  30 years than the non-PRDQT group (70.2% vs. 68.5%, *p* < 0.001). Both multiple gestation rate and cesarean section delivery rate were significantly higher in the PRDQT group (2.7% vs. 1.8%, *p* < 0.001 and 42.3% vs. 37.1%, *p* < 0.001, respectively). Prevalence of hypertensive disorders of pregnancy was significantly greater in the PRDQT

Table 2. Cumulative Incidence of PRDQT in Korea from 2013 to2017					
Age (yr)	Total pregnant women	Patients with PRDQT	Cumulative incidence (%)		
15–19	9,115	146	1.6		
20–24	90,434	1,628	1.8		
25–29	403,578	8,475	2.1		
30–34	772,798	16,229	2.1		
35–39	288,885	6,933	2.4		
40–44	35,710	893	2.5		
45–49	981	38	3.9		
Total	1,601,501	34,342	2.1		

PRDQT: pregnancy-related de Quervain's tenosynovitis.

group (3.5% vs. 2.7%, p < 0.001). However, diabetic disorders of pregnancy were significantly less prevalent in the PRDQT group than in the non-PRDQT group (30.7% vs. 33.4%). Both RA and DM were significantly more prevalent in the PRDQT group (1.9% vs. 0.7%, p < 0.001 and 3.3% vs. 3.0%, p = 0.005, respectively) (Table 3).

The multivariate logistic regression analysis showed that age  $\geq$  30 years, multiple gestation, cesarean section delivery, hypertensive disorders of pregnancy, and underlying RA were significant factors, but underlying DM was not. The odds ratio for underlying RA was the highest of all significant factors, followed by multiple gestation, hypertensive disorders of pregnancy, cesarean section delivery, and age  $\geq$  30 years (Table 4).

#### DISCUSSION

There have been only a few case series reports on the epidemiology of PRDQT. Schumacher et al.<sup>6)</sup> previously reported 6 patients with DQT associated with pregnancy. All 6 patients' symptoms began prior to delivery. Another study by Schned reported that there were 6 pregnant women among 24 women diagnosed with DQT over a 1-year period.<sup>5)</sup> The onset was the second trimester for 1 patient and the third trimester for 5 patients. The author suggested that there might be a true relationship between pregnancy and DQT based on the fact that the mean age of the pregnant women (31.7 years) was lower than that of the entire group (39.5 years). Avci et al. described 18 patients with DQT who were either pregnant or breastfeeding.<sup>4)</sup> The mean age of the patients was 28 years (range, 20-36 years). Five patients developed symptoms during pregnancy, while 13 patients developed symptoms during lactation.

According to our review of the literature, this is the first population-based epidemiologic study focusing exclusively on PRDQT. The purpose of this study was to estimate the nationwide incidence of PRDQT and the associated risk factors using the population-based NHI claims database. This study included over 1.6 million pregnant women across the country and determined that the nationwide cumulative incidence of PRDQT in South Korea was approximately 2.1% from 2013 to 2017. Among identifiable risk factors, age  $\geq$  30 years, multiple gestation, cesarean section delivery, hypertensive disorders of pregnancy, and underlying RA were all associated with the occurrence of PRDQT, but diabetic disorders of pregnancy and underlying DM were not.

Several hypotheses regarding the occurrence of PRDQT have been suggested. According to some authors,

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Table 3. Univariate Analysis of Risk Factors Associated with PRDQT				
Risk factor	Pregnant women with PRDQT (n = 34,342)	Pregnant women without PRDQT (n = 1,567,159)	<i>p</i> -value	
Age (yr)			< 0.001	
< 30	10,249 (29.8)	492,878 (31.5)		
≥ 30	24,093 (70.2)	1,074,281 (68.5)		
Pregnancy type			< 0.001	
Single gestation	33,398 (97.3)	1,538,238 (98.2)		
Multiple gestation	944 (2.7)	28,921 (1.8)		
Delivery method			< 0.001	
Vaginal delivery	19,808 (57.7)	986,231 (62.9)		
C-section delivery	14,534 (42.3)	580,928 (37.1)		
Gestational complication				
Hypertensive disorders of pregnancy	1,207 (3.5)	41,911 (2.7)	< 0.001	
Diabetic disorders of pregnancy	10,549 (30.7)	523,153 (33.4)	< 0.001	
Comorbidity				
Rheumatoid arthritis	658 (1.9)	10,395 (0.7)	< 0.001	
Diabetes mellitus	1,121 (3.3)	46,284 (3.0)	0.005	

Values are presented as number (%).

PRDQT: pregnancy-related de Quervain's tenosynovitis, C-section: cesarean section.

fluid retention during pregnancy and lactation contributed to the edematous state within the first dorsal compartment, increasing susceptibility to stenosing tenosynovitis of APL and EPB tendons.<sup>12,13)</sup> Another hypothesis is that hormonal changes, particularly increased prolactin secretion during pregnancy and lactation, may contribute to the occurrence of DQT.<sup>5,14)</sup> Changes in the secretion of estrogen and progesterone have also been proposed as possible hormonal contributors.<sup>5)</sup> A recent molecular biologic study of intraoperative retinaculum samples from 16 patients with DQT discovered a correlation between estrogen receptor-β expression levels and histologic grades, indicating the presence of a hormonal background.<sup>15)</sup> Another mechanical hypothesis is that postpartum infant care activities such as breastfeeding or milk feeding, lifting, and supporting infants in specific positions require continuous overexertion of the mothers' wrist, which may result in the development of DQT.<sup>5,12,13)</sup>

This study discovered that pregnant women aged  $\geq$  30 years were more prone to DQT than those aged < 30 years. Previous epidemiologic research on DQT in a young and active population using a large military personnel database indicated that age greater than 40 years was a major risk factor for DQT, with an adjusted rate ratio of 3.65 when compared to the age < 20-year group.<sup>16)</sup> Read et al.<sup>17)</sup> described the histologic characteristics of PRDQT in 6 tendon sheath specimens as myxoid degeneration in the absence of acute or chronic inflammation, which were comparable to pregnancy-unrelated DQT. With these clinical and histologic features, one can surmise that PRDQT has a similar pathogenesis to other forms of degenerative tendinopathy. Further studies are necessary to ascertain which elements contribute to the development of degenerative alterations in pregnant women who are relatively young.

In this study, we examined factors associated with pregnancy and delivery, such as pregnancy type, delivery method, and gestational complications. Multiple gestation, cesarean section delivery, and hypertensive disorders in pregnancy were identified as gestational risk factors for PRDQT. Given the association between these factors and increased body fluid during pregnancy, it is quite probable that they did not exist in isolation, but interacted with each other throughout pregnancy.<sup>18)</sup> Even though confounding effects between these factors cannot be ruled out, it can be extrapolated that edema caused by pregnancy-induced body fluid retention may be one of the elements respon-

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Table 4. Multivariate Logistic Regression Analysis for PredictingPregnancy-Related de Quervain's Tenosynovitis					
Risk factor	OR	95% CI	<i>p</i> -value		
Age (yr)					
< 30	1.000				
≥ 30	1.130	1.106-1.154	< 0.001		
Pregnancy type					
Single gestation	1.000				
Multiple gestation	1.379	1.298-1.460	< 0.001		
Delivery method					
Vaginal delivery	1.000				
C-section delivery	1.150	1.121-1.179	< 0.001		
Gestational complication					
Hypertensive disorders in pregnancy	1.233	1.160–1.306	< 0.001		
Comorbidity					
Rheumatoid arthritis	2.461	2.267-2.655	< 0.001		
Diabetes mellitus	0.935	0.878-0.992	0.03		

OR: odds ratio, CI: confidence interval, C-section: cesarean section.

sible for PRDQT development. It is critical to explain the probability of PRDQT occurrence in women with certain gestational risk factors during prenatal evaluation.

Underlying RA was identified as a comorbid risk factor for PRDQT in this study. A previous epidemiologic study about DQT has shown that the RA is an independent risk factor with an adjusted odds ratio of 1.53.<sup>19)</sup> Interestingly, some prior studies have shown that women with RA had lower disease activity during their pregnancy or breastfeeding.<sup>20-22)</sup> According to one observational cohort study, most women with RA improved significantly during pregnancy.<sup>20)</sup> Another laboratory study on the cytokines in pregnant women with RA found that antiinflammatory cytokines increased during pregnancy and decreased after delivery.<sup>21)</sup> Another epidemiologic study based on a community-based registry found that women who breastfed for a longer period of time had a significantly lower risk of developing RA.<sup>22)</sup> These prior findings indicating improvement in RA during pregnancy and breastfeeding seem to be contradictory with the findings of our study, which indicate that pregnant women with underlying RA are more likely to develop PRDQT. Nevertheless, given that a paucity of inflammatory process was detected in previous histologic researches on DQT, it could be hypothesized that underlying RA-related inflammatory responses caused narrowing of the fibro-osseous tunnel, thereby increasing susceptibility to stenosing tenosynovitis of the APL and EPB tendons regardless of the current inflammatory state.<sup>15,23)</sup> Unlike RA, the multivariate analysis established that DM was not a risk factor for PRDQT in the current study. However, prior epidemiologic research on DQT has demonstrated that DM is a risk factor.<sup>19)</sup> These epidemiological distinctions regarding comorbidity between PRDQT and DQT may provide an insight into the different pathogenesis of these two conditions.

There are several limitations of this study. First, the incidence of PRDQT was estimated by calculating the cumulative incidence, not the incidence rate. As the time period of observation was not considered when calculating the cumulative incidence, it cannot be interpreted as indicating the actual risk of a new disease occurrence at any given time, but rather representing an approximate proportion of occurrence. Second, considering the ease of accessibility of medical services in South Korea, this study estimated the incidence based on the assumption that patients with PRDQT would seek medical services for their radial side wrist pain at least once during their pregnant or postpartum period. Nonetheless, given that pregnant or lactating women are generally reluctant to medications or injections, those with milder symptoms might have avoided seeking medical services, which could have caused underestimation of the incidence of PRDQT. Third, because we included only pregnant women who delivered babies, those who terminated their pregnancy prior to delivery via abortion or stillbirth were excluded, which may have influenced the results. Fourth, DQT was clinically diagnosed by a variety of clinicians across the country. Although DQT could be easily diagnosed based on symptoms and simple physical examinations such as the Finkelstein test, less experienced clinicians might have difficulty differentiating DQT from other conditions such as thumb carpometacarpal or scaphotrapeziotrapezoidal joint osteoarthritis or intersection syndrome, all of which can cause radial wrist pain.<sup>1)</sup> Finally, claims data, which were originally intended to be used to obtain government reimbursement, have an inherent limitation. Data that are unnecessary for reimbursement, such as occupation, hand dominance, and breastfeeding status, could not be identified in our data source.

In summary, between 2013 and 2017, the cumulative incidence of PRDQT was approximately 2.1% in South Korea. Age  $\geq$  30 years, multiple gestation, cesarean

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section delivery, hypertensive disorders in pregnancy, and underlying RA were identified as significant risk factors for PRDQT occurrence. The findings of this study could serve as a reference for clinicians seeking to consult patients with PRDQT or to predict its occurrence in pregnant women with certain risk factors, as well as for government health care administrators developing national public health strategies and distributing national health resources. Moreover, further research on the risk factors identified in this study may provide clues to identify etiology and pathophysiology. Future research examining more risk factors in a larger study population recruited over a longer period of time is anticipated to enhance our understanding of not only PRDQT, but also DQT itself.

# **CONFLICT OF INTEREST**

No potential conflict of interest relevant to this article was reported.

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