The prevalence of first-onset temporomandibular disorder in low back pain and associated risk factors

A nationwide population-based cohort study with a 15-year follow-up

Kuei-Chen Lee, MS^{a,b}, Yung-Tsan Wu, MD^b, Wu-Chien Chien, PhD^{c,d,e,*}, Chi-Hsiang Chung, PhD^{c,d}, Liang-Cheng Chen, MD^b, Yi-Shing Shieh, PhD^{a,f,*}

Abstract

The coexistence of low back pain (LBP) and temporomandibular disorder (TMD) has often been noted clinically. However, studies of the association between these two conditions involving a large population with longitudinal evidences are lacking. Therefore, the study aimed to investigate the association between LBP and TMD in a nationwide-matched cohort population with a 15-year follow-up.

Data of 65,121 patients newly diagnosed with LBP were analyzed, along with those of 195,363 (1:3) sex- and age-matched controls. Multivariate Cox regression analysis was used to determine TMD risk between the LBP and non-LBP groups. Kaplan-Meier method was used for determining the cumulative risk of first-onset TMD between groups, with a 15-year follow-up.

The LBP group was more likely to develop first-onset TMD (adjusted hazards ratio (HR) = 1.561, P < .001), after adjusting for demographic variables and comorbidities. The risk factors for TMD were LBP, young age, higher insured premium, and osteoporosis. In the subgroup analysis, the LBP group had a higher risk of TMD than the non-LBP group in all stratifications.

LBP is the risk factor contributing to the development of first-onset TMD. Therefore, clinicians should be reminded to manage LBP disorders concurrently when treating TMD.

Abbreviations: $CCL_R = Charlson$ comorbidity index revised, CI = Confidence interval, HR = Hazard ratio, ICD-9-CM = International Classification of Diseases, Ninth Revision, Clinical Modification, <math>IP = Insured premium, LBP = Low back pain, LPH = Lumbar pelvis hip, NHIRD = National Health Insurance Research Database, RDC = Research Diagnostic Criteria, TMD = Temporomandibular disorder, TMJ = Temporomandibular joint.

Keywords: first-onset temporomandibular disorder, low back pain, National Health Insurance Research Database

Editor: Dennis Enix.

This work was supported by the Tri-Service General Hospital Research Foundation (Grant Reference Numbers TSGH-C105-012, TSGH-C106-010, TSGH-C107-183, TSGH-C108-003, and TSGH-C108-005-006-006-S03).

The authors have no conflicts of interest to disclose.

^a Graduate Institute of Medical Sciences, ^b Department of Physical Medicine and Rehabilitation, Tri-Service General Hospital, ^c School of Public Health, ^d Department of Medical Research, Tri-Service General Hospital, ^e Graduate Institute of Life Science, ^f Department of Dentistry, Tri-Service General Hospital, National Defense Medical Center, Taipei, Taiwan.

^{*} Correspondence: Yi-Shing Shieh, Department of Dentistry, Tri-Service General Hospital, National Defense Medical Center, 114 No.161, Sec. 6, Minquan E. Rd., Neihu Dist., Taipei, Taiwan (e-mail: ndmcyss@mail.ndmctsgh.edu.tw), Wu-Chien Chien, Department of Medical Research, Tri-Service General Hospital, National Defense Medical Center, 7115R, No. 325, Section 2, Cheng-Kung Road, Neihu District, Taipei, Taiwan (e-mail: chienwu@ndmctsgh.edu.tw).

Copyright © 2020 the Author(s). Published by Wolters Kluwer Health, Inc. This is an open access article distributed under the terms of the Creative Commons Attribution-Non Commercial License 4.0 (CCBY-NC), where it is permissible to download, share, remix, transform, and buildup the work provided it is properly cited. The work cannot be used commercially without permission from the journal.

How to cite this article: Lee KC, Wu YT, Chien WC, Chung CH, Chen LC, Shieh YS. The prevalence of first-onset temporomandibular disorder in low back pain and associated risk factors: A nationwide population-based cohort study with a 15-year follow-up. Medicine 2020;99:3(e18686).

Received: 26 August 2019 / Received in final form: 22 November 2019 / Accepted: 7 December 2019

http://dx.doi.org/10.1097/MD.00000000018686

1. Introduction

Temporomandibular disorders (TMD) are complex and multifactorial diseases. The common signs and symptoms of TMD are orofacial pain, clicking or crepitus of the temporomandibular joints (TMJ), limited mandible movement, and deviation from the midline during movement.^[1,2] Besides local TMJ pain, TMD is often associated with simultaneous pain in the head, neck, and back areas. The evidence from clinical and epidemiologic studies shows that TMD coexists with numerous systemic disorders including fibromyalgia, low back pain (LBP), spinal pain, chronic fatigue syndrome, irritable bowel syndrome, sleep disorders, tension-type and migraine headaches, and allergic conditions.^[3,4,5] TMD seems to be of multifactorial etiology, including parafunctional habits, bruxism, deleterious body posture, occlusal features, trauma, overload, and stress^[6,7,8]; hence, TMD is not easy to cure. Owing to the complexity of the symptoms and etiology, many patients with TMD cannot be appropriately treated according to the cause of the disease.

Medicine

In fact, LBP and TMD often coexist in clinical practice. If we treat patients focusing on the TMJ only, the symptoms will easily recur. This reminds us that we should determine the cause of the condition in advance or manage comorbidities simultaneously. Previous studies have documented a significant association between LBP and TMD.^[4,5,9,10] In 2000–2005 US National Health Interview Survey, 64% of 189,977 patients with TMD

reported LBP.^[4] Wiesinger et al.^[9] reported significant associations between back pain and musculoskeletal disorders in the jaw or face and indicated the comorbidity between these two conditions. However, these investigations were only crosssectional studies; studies involving a large population with longitudinal evidences are limited. Moreover, data on the possible mechanism linking LBP to TMD are lacking.

LBP is the most common musculoskeletal condition affecting the adult population, and >70% of the population has experienced acute episode of LBP at some point in their lives.^[11] The common causes of LBP include muscle strain, herniated intervertebral disc, degenerative joint, sacrolilitis, spinal stenosis, sciatica, and other unspecified back disorders.^[12,13,14] If LBP and TMD occur concurrently, functional activities would be mostly affected. Therefore, early prevention of TMD occurrence as comorbidity in patients with LBP is important, to avoid the profound functional limitation. To achieve that, this study aimed to investigate the association between LBP and first-onset TMD and to determine the possible mechanism linking these two diseases, using the Taiwan's National Health Insurance Research Database (NHIRD) with a 15-year follow up.

2. Methods

2.1. Data sources and ethics

Data retrieved from the 2000-2015 Longitudinal Health Insurance Database, a subset of Taiwan's NHIRD, were used in this study to investigate the association between LBP and TMD over a 15-year period. The NHIRD comprises comprehensive data on disease diagnoses coded by the International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM)^[15]; treatments and procedures; service dates; patients' demographic characteristics; and drug prescriptions. The NHIRD is strictly used for research purposes, and all personally identifiable information is encrypted before being released to researchers to maintain patients' confidentiality. The National Health Insurance program was launched in Taiwan in 1995 and, as at June 2009, included contracts with 97% of healthcare providers, with approximately 23 million beneficiaries or >99% of the entire population. Several studies have shown the accuracy and validity of diagnoses in the NHIRD.^[16,17] This study received approval from the Tri-Service General Hospital (IRB No.2-103-05-043). Since the patient identifiers were encrypted to protect patient confidentiality before the data were used for research purposes, the requirement for patient written or verbal consent was waived.

2.2. Study design and sample population

This retrospective matched-cohort study analyzed patients with LBP-related diagnosis from January 1, 2000 to December 31, 2004 according to the following ICD-9-CM codes: 720.2, 721.3, 721.42, 722.10, 722.52, 722.73, 722.9, 724.02, 724.2, 724.3, 724.5, 724.9, and 847.2 (Table 1).^[18,19] Each enrolled LBP patient was required to have made at least six outpatient visits or one inpatient visit. All diagnoses of LBP were made by a board-certified rehabilitation, orthopedics, neurology, neurosurgery, and orthopedic doctor. For each patient with LBP, three patients without LBP, who were matched for sex, age, medical visits, and index year, were allocated to the comparison group. A total of 260,484 patients (study group, n=65,121; comparison group, n=195,363) were enrolled in this study. Both groups were followed-up from the index date until the onset of TMD,

Table 1

Diagnosis groups with corresponding ICD-9-CM codes.

Variables	ICD-9-CM
LBP related diagnosis	
Sacroiliitis	720.2
Lumbosacral spondylosis without myelopathy	721.3
Spondylosis in lumbar region	721.42
Lumbar intervertebral disc without myelopathy	722.10
Lumbar or lumbosacral intervertebral disc	722.52
Intervertebral disc disorders in lumbar region	722.73
Other and unspecified disc disorder in lumbar region	722.9
Spinal stenosis in lumbar region	724.02
Lumbago	724.2
Sciatica	724.3
Backache, unspecified	724.5
Other unspecified back disorders	724.9
Sprains and strains in lumbar region	847.2
Events: TMD	524.6
Excluding	
Malignant neoplasm of the lip, oral cavity, and pharynx	140-149
Injuries on the head, face, or neck	800–804, 830,
	941, 959.0
Comorbidities	
HTN	401–405
Stroke	430–438
COPD	490-496
CKD	585-586
Migraine	346
Osteoporosis	733.0
Hyperlipidemia	272.0-272.4
Nonpsychotic mental disorders	300–314
Psychosis	295–299
CAD	410-414
Insomnia	780.52
KA	714

CAD = coronary artery disease, CHF = congestive heart failure, CKD = chronic kidney disease, COPD = chronic obstructive pulmonary disease, HTN = hypertension, ICD-9-CM = International Classification of Diseases, Ninth Revision, Clinical Modification, LBP = low back pain, NMD = nonpsychotic mental disorders, RA = rheumatoid arthritis, TMD = temporomandibular disorder.

withdrawal from the NHI program, or the end of 2015. The diagnosis of TMD was based on the ICD-9-CM code 524.6, which was adapted from previous studies,^[6,20,21] and has been used widely in population studies. All diagnoses of TMD were included at least 1 year after the LBP diagnosis and confirmed by board-certified rehabilitation physicians or dentists according to the physical examination or imaging findings, including three groups of the Research Diagnostic Criteria for Temporomandibular Disorders (RDC/TMD).^[22]

The flowchart of the study sample selection process is presented in Figure 1.

2.3. Exclusion criteria and comorbidities

Patients diagnosed with

- LBP and TMD before 2000 (or LBP prior to the first visit for TMD);
- (2) those with diseases and injuries that may involve the temporomandibular joint (details provided in Table 1);
- (3) those aged <20 years or >70 years; and
- (4) those with unknown sex were all excluded from the study.

Associated systemic diseases and risk factors for TMD were analyzed to compare the effect of these confounding factors



between the two groups.^[6,23,21] Comorbidities were identified using the ICD-9-CM diagnosis codes, and detailed information was shown in the Table 1.

2.4. Statistical analysis

All analyses were performed using SPSS version 22 (IBM Corp., Armonk, NY). The chi-square test and *t* test were used to statistically examine the differences between the two cohorts. Multivariate Cox regression analysis was performed to determine the risk of TMD, with the results of the analysis presented as hazards ratio (HR) with 95% confidence interval (CI). The difference in the risk of TMD between the study and comparison groups was estimated using Kaplan-Meier analysis with the log-rank test. A two-tailed *P* value <.05 was considered statistically significant.

3. Results

3.1. Sample characteristics

The characteristics of the study participants are presented in Table 2. There were no significant differences between the study and comparison groups in terms of sex, age, and comorbidity such as nonpsychotic mental disorders. There were significant differences in the insured premium (IP), level of care, and most of the comorbidities, such as hypertension, diabetes mellitus, congestive heart failure, stroke, chronic obstructive pulmonary disease, chronic kidney disease, migraine, osteoporosis, hyperlipidemia, nonpsychotic mental disorders, psychoses, coronary artery disease, insomnia, rheumatoid arthritis, and Charlson comorbidity index revised (CCI_R) between the study and control groups.

3.2. Kaplan-Meier analysis of the cumulative risk for TMD

The LBP group showed a significantly higher cumulative risk of TMD than the non-LBP group (log-rank P < .001). The cumulative risk of TMD became significantly different between the 2 groups since the tenth year of tracking (Fig. 2). We divided the follow-up duration into 3 periods and found that the difference became significant since the 6 to 10-year period and became more significant as the LBP became more chronic (Table 3).

3.3. Factors for the incidence of TMD in the Cox regression analysis

The Cox regression analysis showed that the adjusted HR for the development of TMD was 1.561 times greater in the LBP group

Table 2

Characteristics of the study and comparison group at baseline.

	Study	group	Compariso	on group	Р
LBP	W	ith	With	out	
Variables	n	%	n	%	
Total	65,121	25.00	195,363	75.00	
Sex					.999
Male	33,104	50.83	99,312	50.83	
Female	32,017	49.17	96,051	49.17	
Age (years)	52.60 ± 12.66	52.50 ± 12.86	0.214		
Age groups (years)					.999
20–29	3,895	5.98	11,685	5.98	
30–39	8,467	13.00	25,401	13.00	
40–49	13,019	19.99	39,057	19.99	
50–59	16,219	24.91	48,657	24.91	
60–70	23,521	36.12	70,563	36.12	
IP (NTD)					<.001
<18,000	58,045	89.13	175,069	89.61	
18,000-34,999	4625	7.10	13,805	7.07	
≥35,000	2451	3.76	6,489	3.32	
Level of care					<.001
Hospital center	12,822	19.69	62,860	32.18	
Regional hospital	18,504	28.41	57,077	29.22	
Local hospital	33,795	51.90	75,426	38.61	
Comorbidity	,		,		
HTN	6.765	10.39	24.061	12.32	<.001
CHF	562	0.86	2.831	1.45	<.001
Stroke	2,170	3.33	12,157	6.22	<.001
COPD	3,312	5.09	9,542	4.88	.040
CKD	431	0.66	3,808	1.95	<.001
Migraine	86	0.13	144	0.07	<.001
Osteoporosis	2037	3.13	839	0.43	<.001
Hyperlipidemia	2733	4.20	6.565	3.36	<.001
NMD	863	1.33	2.649	1.36	.569
Psychoses	289	0.44	3.526	1.80	<.001
CAD	3.254	5.00	13.037	6.67	<.001
Insomnia	360	0.55	387	0.20	<.001
RA	283	0.43	661	0.34	<.001
CCI_R	0.16:	±0.78	0.45±	1.65	<.001

CAD = coronary artery disease, CCI_R = Charlson comorbidity index revised, CHF = congestive heart failure, CKD = chronic kidney disease, HTN = hypertension, IP = insured premium, LBP = low back pain, NMD = nonpsychotic mental disorders, NTD = new Taiwan dollars, RA = rheumatoid arthritis, TMD = temporomandibular disorder.



Figure 2. Kaplan-Meier for cumulative risk of first-onset TMD in patients aged 20 to 70 years stratified by low back pain with log-rank test. LBP=low back pain, TMD=temporomandibular disorder.

than in the non-LBP group (95% CI: 1.220–1.996, P < .001), after adjusting for demographic variables and comorbidities.

There was a higher risk of TMD in younger patients than in older patients (60–70 years) [adjusted HR for 20–29 years = 5.615, P < .001; for 30–39 years = 2.406, P = .001]. Moreover, those with higher IP showed a higher incidence of TMD than those with lower IP [adjusted HR for 18,000–34,999 NT= 12.741 (P < .001), for $\ge 35,000$ NT = 33.684 (P < .001)]. Moreover, osteoporosis was also a risk factor contributing to TMD occurrence (adjusted HR = 1.860, P = .003) (Table 4).

3.4. Risk of TMD between the study and comparison groups stratified by variables and comorbidities

The LBP group had a higher risk of TMD occurrence than the non-LBP group in all stratifications (Table 5).

4. Discussion

Based on the results, the overall adjusted HR of TMD occurrence in the LBP group was 1.561 (95% CI: 1.220–1.996, P < .001) after adjusting for demographic variables and comorbidities. In other words, the patients with LBP had a 1.561-fold increased risk of developing first-onset TMD than the patients without LBP. The subgroup analysis stratified by demographic variables

hisk of tempor	isk of temporomanubular disorder among unerent tracking year.											
LBP Tracking year	With				Without		With vs Without					
	Events	PYs	ER	Events	PYs	ER	Ratio	Adjusted HR	Р			
Overall	107	736,810.80	14.52	161	2,456,015.93	6.56	2.215	1.561	<.001			
0–5	19	63,676.29	29.84	42	280,818.14	14.96	1.995	1.384	.243			
6–10	37	204,098.16	18.13	58	730,541.15	7.94	2.283	1.701	.013			
≧11	51	469,036.35	10.87	61	1,444,656.64	4.22	2.575	1.823	.002			

 Table 3

 Risk of temporomandibular disorder among different tracking year.

Adjusted HR = hazard ratio adjusted for demographic variables and comorbidities, ER = event rate per 10⁵ person-years, LBP = low back pain, PYs = person-years.

and comorbidities also revealed a higher risk of TMD in the LBP group than in the non-LBP group in all stratifications. It indicated LBP as a strong risk factor contributing to the occurrence of firstonset TMD. To the best of our knowledge, this is the first report that investigated the association between LBP and the risk of developing first-onset TMD through a nationwide populationbased cohort study, with 15 years of follow-up. The study results support previous cross-sectional observations of comorbidity

Table 4

Risk	Factors	for	the	incidence	of	TMD	in	the	Сох	regression
analy	vsis.									

Variables	Adjusted HR	95% CI	95% CI	Р
LBP				
Without	Reference			
With	1.561	1.220	1.996	<.001
Sex				
Male	0.768	0.601	1.002	.053
Female	Reference			
Age groups (years)				
20–29	5.615	2.014	15.653	<.001
30-39	2.406	1.540	3.758	.001
40-49	1.017	0.727	1.576	.923
50-59	1.001	0.646	1.422	.997
60-70	Reference			
IP (NTD)				
<18,000	Reference			
18,000–34,999	12.741	9.176	17.690	<.001
≥35,000	33.684	20.730	54.731	<.001
Level of care				
Hospital center	1.264	0.935	1.741	.742
Regional hospital	1.093	0.873	1.397	.161
Local hospital	Reference			
Comorbidity				
HTN	1.006	0.657	1.194	.426
CHF	1.065	0.602	1.885	.828
Stroke	1.136	0.867	1.597	.105
COPD	1.753	0.944	2.278	.294
CKD	1.241	0.633	2.243	.529
Migraine	0.000	_	_	.975
Osteoporosis	1.860	1.212	3.483	.003
Hyperlipidemia	1.414	0.813	2.459	.220
NMD	0.000	_	_	.922
Psychoses	1.254	0.789	2.315	.687
CAD	1.240	0.833	1.844	.289
Insomnia	1.021	0.253	4.123	.976
RA	1.021	0.253	4.123	.976
CCI_R	0.989	0.938	1.043	.681

CAD = coronary artery disease, CCI_R = Charlson comorbidity index revised, CHF = congestive heart failure, CKD = chronic kidney disease, COPD = chronic obstructive pulmonary disease, HTN = hypertension, IP = insured premium, LBP = low back pain, NMD = nonpsychotic mental disorders, NTD = new Taiwan dollars, RA = rheumatoid arthritis, TMD = temporomandibular disorder.

relationship between LBP and TMD.^[4,5,9,10] However, their results only demonstrate the coexistence of LBP and TMD, but not that LBP may contribute to the development of first-onset TMD, as reported in the present study with 15-year follow-up. In the cumulative risk of TMD, we found that the difference between LBP and non-LBP groups became significant since the 6–10-year follow-up (Table 3), indicating that when LBP became more chronic, the occurrence of TMD would be more obvious.

Patients with LBP usually reveal problems in the lumbar spine and in the pelvic and hip joints.^[24] A recent electromyography study demonstrated that cervical postural muscles (deep neck flexors) were more difficult to recruit in non-neutral spinal lumbopelvic alignment than in a neutral posture.^[25] Therefore, problems in the lumbar-pelvis-hip (LPH) complex will affect the normal coordination of the neck muscle. Fink et al^[26] proposed the functional relationships between the craniomandibular system, cervical spine, and the sacroiliac joint. Another study also documented that LBP might be associated with an increased incidence of neck pain.^[27] Moreover, TMD was already confirmed to be related to cervical pain or upper quarter dvsfuntion.^[28,29] El-Hamalawy^[30] reported that the forward head movement correction can improve myogenic temporomandibular joint dysfunction. Seagreen reported that tensed muscles around the head and neck would induce TMD symptoms.^[31] Therefore, problems in the LPH complex can affect the head and neck, subsequently causing temporomandibular joint dysfunction. This may explain why patients with LBP would develop first-onset TMD, and the trend became obvious with chronic LBP of >6 years. However, the real mechanism of LBP triggering TMD still needs further investigation.

Regarding the factors affecting the incidence of first-onset TMD, patients with younger age, higher IP, and osteoporosis have an increased risk of first-onset TMD. There was a higher risk of TMD in the 20-29- and 30-39-year-old groups than in the other age groups. The age prevalence was similar to those reported in other studies.^[2,32] Gonçalves et al^[2] reported that the age groups with the largest proportion of first-onset TMD were the 31-40-year-old and 21-30-year old age groups (21.3% and 20.2%, respectively). In Aggarwal et al's study, patients aged 18 to 35 years were significantly more likely to report new oral facial pain at the 2-year follow-up than those aged 64 to 75 years.^[32] Stress factors and low adaptive capacity might explain why young people were most affected.^[33] The finding could be supported by other data from our results in the present study, indicating that those with middle (18,000–34,999 NT) and high (≥35,000 NT) IP showed a high risk of TMD occurrence. In the Taiwan NHI Program, a higher IP may indicate a higher socioeconomic status, which may be associated with higher stress. Stress is usually related to the presence of TMD pain that may be due to increased activity of the masticatory muscles, which is a compensatory reaction to stress.^[8,34]

Table 5	ondikular -	lioordor between	n otudu av	in and series	norioon ereen -t-	atified by		oomorbidition	_	
Risk of temporon	nandibular d	Study group	n study grou	ip and com	Comparison group str	ariables and	Risk of TMD			
		With			Without			With vs Without	<u></u>	
LDF	Evente	BVo	ED	Evente	BVo	ED	Potio		D	
Suduneu	Events	F 15	En	Events	FIS	En	nauu	Aujusteu nn	r	
Total	107	736,810.80	14.52	161	2,456,015.93	6.56	2.215	1.561	<.001	
Sex		050 707 70	10.00	70	1 000 101 00	0.45	0.000		0.01	
Male	44	356,797.72	12.33	76	1,236,421.83	6.15	2.006	1.411	<.001	
Female	63	380,013.08	16.58	85	1,219,594.09	6.97	2.379	1.676	<.001	
Age groups (years)										
20–29	4	4075.09	98.16	5	18,142.15	27.56	3.562	2.510	<.001	
30–39	13	32,433.27	40.08	16	125,455.42	12.75	3.143	2.216	<.001	
40-49	12	74,447.28	16.12	14	250,556.82	5.59	2.885	2.033	<.001	
50-59	15	129,932.67	11.54	21	439,522.59	4.78	2.416	1.703	<.001	
60-70	63	495,922.49	12.70	105	1,622,338.95	6.47	1.963	1.382	<.001	
IP (NTD)										
<18,000	71	721,962.87	9.83	119	2,412,265.61	4.93	1.994	1.405	<.001	
18,000-34,999	26	12,981.07	200.29	30	36,523.47	82.14	2.438	1.719	<.001	
≥35,000	10	1,866.86	535.66	12	7226.84	166.05	3.226	2.229	<.001	
Level of care										
Hospital center	39	220.333.69	17.70	66	858.371.31	7.69	2.302	1.672	<.001	
Regional hospital	44	340.514.98	12.92	62	1.065.924.54	5.82	2.222	1.598	<.001	
Local hospital	24	175,962,13	13.64	33	531,720,08	6.21	2.198	1.544	< .001	
Comorbidity	<u> </u>	110,002110	10101	00	001,120100	0.2.1	21100		(1001	
HTN										
Without	76	5/13 303 30	13.00	118	1 810 365 80	6 52	2 1/6	1 500	~ 001	
With	21	102 507 50	16.02	12	645 650 12	0.52	2.140	1.509	< 001	
WIUI CUE	51	193,307.30	10.02	40	040,000.12	0.00	2.400	1.099	<.001	
UTF Without	00	704 010 00	14.00	140	0.040 504 60	6.07	0.000	1 550	< 001	
WILLIOUL	99	704,210.90	14.00	149	2,340,324.00	0.37	2.200	1.070	<.001	
WILLI	8	32,599.90	24.54	12	115,491.25	10.39	2.362	1.078	<.001	
Stroke	100	070 040 00	15.07	150	0 000 700 00	0.00	0.100	1 5 40	. 001	
Without	102	6/6,640.66	15.07	153	2,220,789.62	6.89	2.188	1.543	<.001	
With	5	60,170.14	8.31	8	235,226.31	3.40	2.443	1.782	<.001	
COPD										
Without	99	680,425.86	14.55	154	2,276,096.64	6.77	2.150	1.511	<.001	
With	8	56,384.93	14.19	7	179,919.29	3.89	3.647	2.596	<.001	
CKD										
Without	102	716,544.62	14.23	154	2,352,743.10	6.55	2.175	1.532	<.001	
With	5	20,266.18	24.67	7	103,272.83	6.78	3.640	2.565	<.001	
Migraine										
Without	107	735,684.31	14.54	161	2,453,873.74	6.56	2.217	1.561	<.001	
With	0	1126.49	0.00	0	2142.18	0.00	_	-	-	
Osteoporosis										
Without	102	727,155.27	14.03	158	2,440,591.80	6.47	2.167	1.522	<.001	
With	5	9,655.52	51.78	3	15,424.12	19.45	2.662	1.879	<.001	
Hyperlipidemia		-,			- 1					
Without	99	709.234.44	13.96	154	2.361.100.58	6.52	2.140	1.482	<.001	
With	8	27.576.35	29.01	7	94,915,34	7.37	3.934	2,789	<.001	
NMD	-				,					
Without	107	722 336 46	14.81	161	2 411 933 67	6 68	2 2 1 9	1 561	< 001	
With	0	1/ /7/ 33	0.00	0	4/ 082 26	0.00	2.215	-	<.001	
Peychococ	0	17,77,00	0.00	0	44,002.20	0.00				
Without	102	701 960 46	1/12	150	2 280 670 28	6 68	2 1 1 6	1 465	< 001	
With	102	1/ 0/02.40	14.15	109	2,300,070.30	0.00	10 601	7.403	< 001	
WILLI	Э	14,940.34	33.40	2	70,340.04	2.00	12.001	7.010	<.001	
CAD	00	001 001 71	14.07	1.40	0 100 004 05	0.50	0.157	1 500	. 001	
WITHOUT	93	001,081./1	14.07	143	2,192,804.65	6.52	2.15/	1.520	<.001	
With	14	75,729.08	18.49	18	263,211.27	6.84	2.703	1.909	<.001	
Insomnia		700				o				
Without	105	730,155.26	14.38	159	2,438,425.15	6.52	2.205	1.550	<.001	
With	2	6655.54	30.05	2	17,590.77	11.37	2.643	1.897	<.001	
RA										
Without	107	732,068.67	14.62	161	2,444,015.39	6.59	2.219	1.561	<.001	
With	0	4742.12	0.00	0	12.000.54	0.00	_	_	_	

Adjusted HR = hazard ratio adjusted for demographic variables and comorbidities, CAD = coronary artery disease, CHF = congestive heart failure, CKD = chronic kidney disease, ER = event rate per 10^5 person-years, HTN = hypertension, IP = insured premium, LBP = low back pain, NMD = nonpsychotic mental disorders, NTD = new Taiwan dollars, PYs = person-years, RA = rheumatoid arthritis.

Contrarily, a Brazilian study reported that poor economic class was significantly associated with the diagnosis of TMD.^[35] However, another US study showed that the socioeconomic measures were not significantly associated with TMD incidence in their participants.^[36]

Our data showed that osteoporosis is a risk factor contributing to TMD development in patients with LBP. Previous studies proposed that subchondral bone loss or osteoporotic changes in the temporomandibular joint may account for the high risk of TMD occurrence in LBP patients when combined with osteoporosis.^[37-40] With regard to the association between osteoporosis and the occurrence of TMD, osteoporosis may induce bone loss in both the condyle and temporal components of the TMJ and impede the functional harmony of the masticatory system, increasing the occurrence of TMD.^[39,40] In addition, our results indicate that migraine is not a risk factor for TMD development. Although some previous studies^[41-43] reported that migraine is a common symptom in TMD individuals, they only proposed a comorbidity relationship between migraine and TMD according to crosssectional evidence, using TMD questionnaires or physical examination. However, our cohort study demonstrated that there was no cause-effect relationship between migraine and TMD with 15 years of follow-up. This may be because the prevalence of migraine was significantly higher in individuals with myofascial TMD than in those with non-myofascial TMD.^[44,45] However, myofascial TMD only involves group I of RDC/TMD, and our study enrolled three groups of RDC/TMD. Therefore, the causeeffect relationship between myofascial TMD and migraine needs to be further investigated with a longitudinal follow-up.

The event rate of TMD in the present study is lower than that in other studies,^[3,10] which may be due to the following reasons. First, this study had stricter inclusion criteria such as TMD patients with identification ICD-9 code of 524.6 confirmed by doctors during outpatient or inpatient visits. Many previous studies only included patients who responded to a questionnaire or telephone interview.^[3,4,10,36] The real incidence of TMD in the Taiwan population was much higher than that found in medical records, because many patients with TMD did not seek medical treatment. Moreover, the difference in ethnicity should also be taken into consideration.^[4,10]

4.1. Limitations and strengths

There are several limitations in this study. First, there are many etiologies of LBP, but our study only analyzed the most common related diagnosis. Otherwise, data on pain severity, laboratory parameters, and radiologic images were not available in the NHIRD; thus, we could not classify the severity of LBP among the enrolled participants. Second, the actual prevalence of TMD might have been underestimated in the NHIRD, because some symptoms associated with TMD are often observed in other diseases, resulting in confusion or misdiagnosis.

Moreover, the major strength of our study is that we used data based on clinical diagnosis, which enabled a reliable cohort assessment. Furthermore, our longitudinal cohort study could overcome the major limitation of cross-sectional and case-control study designs by only providing information on disease incidence.

5. Conclusion

LBP is a risk factor contributing to the development of first-onset TMD. Therefore, for appropriate TMD management, clinicians

should be reminded to manage both LBP disorders and TMD concurrently, especially among individuals presenting with associated risk factors, such as age 20 to 39 years, high IP, and osteoporosis.

Author contributions

Data curation: Wu-Chien Chien, Chi-Hsiang Chung, and Kuei-Chen Lee.

- Funding acquisition: Wu-Chien Chien, Yi-Shing Shieh, and Kuei-Chen Lee.
- Methodology: Wu-Chien Chien, Kuei-Chen Lee, and Yi-Shing Shieh.

Project administration: Yung-Tsan Wu and Liang-Cheng Chen. **Software:** Chi-Hsiang Chung.

- Supervision: Yung-Tsan Wu and Liang-Cheng Chen.
- Writing original draft: Kuei-Chen Lee and Yung-Tsan Wu.
- Writing review & editing: Yung-Tsan Wu, Wu-Chien Chien, and Yi-Shing Shieh.

References

- [1] Inglehart MR, Patel HM, Widmalm SE, et al. Self-reported TMJD symptoms, oral health and quality of life of children in kindergarten through grade 5: does gender, race, and socio-economic background matter? J Am Dent Assoc 2016;147:131–41.
- [2] Gonçalves DA, Dal Fabbro AL, Campos JA, et al. Symptoms of temporomandibular disorders in the population: an epidemiological study. J Orofac Pain 2010;24:270–8.
- [3] Sanders AE, Slade GD, Bair E, et al. General health status and incidence of first-onset temporomandibular disorder: the OPPERA prospective cohort study. J Pain 2013;14:T51–62.
- [4] Plesh O, Adams SH, Gansky SA. Temporomandibular joint and muscle disorder (TMJMD) – type pain and co-morbid pains in a national US sample. J Orofac Pain 2011;25:190–8.
- [5] Wiesinger B, Malker H, Englund E, et al. Does a dose-response relation exist between spinal pain and temporomandibular disorders? BMC Musculoskelet Disord 2009;10:28.
- [6] Liao CH, Chang CS, Chang SN, et al. The risk of temporomandibular disorder in patients with depression: a population-based cohort study. Community Dent Oral Epidemiol 2011;39:525–31.
- [7] Furquim BD, Flamengui LM, Conti PC. TMD and chronic pain: A current view. Dental Press J Orthod 2015;20:127–33.
- [8] Reissmann DR, John MT, Schierz O, et al. Stress-Related adaptive versus maladaptive coping and temporomandibular disorder pain. J Orofac Pain 2012;26:181–90.
- [9] Wiesinger B, Malker H, Englund E, et al. Back pain in relation to musculoskeletal disorders in the jaw-face: a matched case–control study. Pain 2007;131:311–9.
- [10] Sipilä K, Suominen AL, Alanen P, et al. Association of clinical findings of temporomandibular disorders (TMD) with self-reported musculoskeletal pains. Eur J Pain 2011;15:1061–7.
- [11] Allegri M, Montella S, Salici F, et al. Mechanisms of low back pain: a guide for diagnosis and therapy. F1000Res 2016;5:F1000Faculty Rev-1530.
- [12] De Palma MJ, Ketchum JM, Saullo T. What is the source of chronic low back pain and does age play a role? Pain Med 2011;12:224–33.
- [13] Jhun HJ, Park JY. Estimated number of Korean adults with back pain and population-based associated factors of back pain: data from the Fourth Korea National Health and Nutrition Examination Survey. J Korean Neurosurg Soc 2009;46:443–50.
- [14] Hestbaek L, Leboeuf-Yde C, Manniche C. Is low back pain part of a general health pattern or is it a separate and distinctive entity? A critical literature review of comorbidity with low back pain. J Manip Physiol Ther 2003;26:243–52.
- [15] Chinese Hospital AssociationICD-9-CM English-Chinese Dictionary. Taipei: Chinese Hospital Association Press; 2000.
- [16] Ho Chan WS. Taiwan's healthcare report 2010. EPMA J 2010;1:563-85.
- [17] Cheng CL, Kao YH, Lin SJ, et al. Validation of the National Health Insurance Research Database with ischemic stroke ases in Taiwan. Pharmacoepidemiol Drug Saf 2011;20:236–42.

- [18] Chia YY, Lo Y, Chen YB, et al. Risk of chronic low back pain among parturients who undergo cesarean delivery with neuraxial anesthesia: a nationwide population-based retrospective cohort study. Medicine (Baltimore) 2016;95:e3468.
- [19] Chung YC, Hung CT, Li SF, et al. Risk of musculoskeletal disorder among Taiwanese nurses cohort: a nationwide population-based study. BMC Musculoskelet Disord 2013;14:144.
- [20] Lin CY, Chung CH, Chu HY, et al. Prevalence of temporomandibular disorders in rheumatoid arthritis and associated risk factors: a nationwide study in Taiwan. J Oral Facial Pain Headache 2017;31:e29–36.
- [21] Lee CF, Lin MC, Lin HT, et al. Increased risk of tinnitus in patients with temporomandibular disorder: A retrospective population-based cohort study. Eur Arch Otorhinolaryngol 2016;273:203–8.
- [22] Dworkin SF, LeResche L. Research diagnostic criteria for temporomandibular disorders: review, criteria, examinations and specifications, critique. J Craniomandib Disord 1992;6:301–55.
- [23] Huang GJ, LeResche L, Critchlow CW, et al. Risk factors for diagnostic subgroups of painful temporomandibular disorders (TMD). J Dent Res 2002;81:284–8.
- [24] Shum GL, Crosbie J, Lee RY. Effect of low back pain on the kinematics and joint coordination of the lumbar spine and hip during sit-to-stand and stand-to-sit. Spine 2005;30:1998–2004.
- [25] Falla D, O'Leary S, Fagan A, et al. Recruitment of the deep cervical flexor muscles during a postural-correction exercise performed in sitting. Man Ther 2007;12:139–43.
- [26] Fink M, Wähling K, Stiesch-Scholz M, et al. The functional relationship between the craniomandibular system, cervical spine, and the sacroiliac joint: a preliminary investigation. Cranio 2003;21:202–8.
- [27] Lee HY, Wang JD, Chang HL, et al. The association between asymmetric hip mobility and neck pain in young adults. J Manipulative Physiol Ther 2013;36:364–8.
- [28] Walczyńska-Dragon K, Baron S. The biomechanical and functional relationship between temporomandibular dysfunction and cervical spine pain. Acta Bioeng Biomech 2011;13:93–8.
- [29] Chang TH, Duh DY, Lee MS, et al. Therapeutic effect of combining muscle relaxant and posture correction in patients with myofascial pain dysfunction syndrome of temporomandibular disorder. J Med Sci 2013;33:183–9.
- [30] El-Hamalawy FA. Forward head correction exercises for management of myogenic temporomandibular joint dysfunction. J Am Sci 2011;7:71–7.
- [31] Seagreen ME. An investigation into the relationship of myofascial trigger points in the head and neck region in association with temporomandibular

joint dysfunction. [masters in technology: chiropractic dissertation]. Durban, South Africa: Durban University of Technology; 2009. in press.

- [32] Aggarwal VR, Macfarlane GJ, Farragher TM, et al. Risk factors for onset of chronic oro-facial pain–results of the North Cheshire oro-facial pain prospective population study. Pain 2010;149:354–9.
- [33] Mazzetto MO, Rodrigues CA, Magri LV, et al. Severity of TMD related to age, sex and electromyographic analysis. Braz Dent J 2014;25:54–8.
- [34] Ohrbach R, Blascovich J, Gale EN, et al. Psychophysiological assessment of stress in chronic pain: Comparisons of stressful stimuli and of response systems. J Dent Res 1998;77:1840–50.
- [35] Magalhães BG, de-Sousa ST, de Mello VV, et al. Risk factors for temporomandibular disorder: Binary logistic regression analysis. Med Oral Patol Oral Cir Bucal 2014;19:e232–6.
- [36] Slade GD, Bair E, Greenspan JD, et al. Signs and symptoms of first-onset TMD and sociodemographic predictors of its development: the OPPERA prospective Cohort Study. J Pain 2013;14:T20–32.
- [37] Dervis E. Oral implications of osteoporosis. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2005;100:349–56.
- [38] Chen K, Zhang N, Ding L, et al. Early intra-articular injection of alendronate reduces cartilage changes and subchondral bone loss in rat temporomandibular joints after ovariectomy. Int J Oral Maxillofac Surg 2014;43:996–1004.
- [39] Gruber HE, Gregg J. Subchondral bone resorption in temporomandibular joint disorders. Cells Tissues Organs 2003;174:17–25.
- [40] Klemetti E, Vainio P, Kroger H. Craniomandibular disorders and skeletal mineral status. Cranio 1995;13:89–92.
- [41] Stuginski-Barbosa J, Macedo HR, Bigal ME, et al. Signs of temporomandibular disorders in migraine patients: a prospective, controlled study. Clin J Pain 2010;26:418–21.
- [42] Goncalves DA, Camparis CM, Speciali JG, et al. Treatment of comorbid migraine and temporomandibular disorders: a factorial, double-blind, randomized, placebo-controlled study. J Orofac Pain 2013;27:325–35.
- [43] Lopes SL, Costa AL, Gamba Tde O, et al. Lateral pterygoid muscle volume and migraine in patients with temporomandibular disorders. Imaging Sci Dent 2015;45:1–5.
- [44] Dahan H, Shir Y, Nicolau B, et al. Self-reported migraine and chronic fatigue syndrome are more prevalent in people with myofascial vs nonmyofascial temporomandibular disorders. J Oral Facial Pain Headache 2016;30:7–13.
- [45] Goncalves DA, Camparis CM, Speciali JG, et al. Temporomandibular disorders are differentially associated with headache diagnoses: a controlled study. Clin J Pain 2011;27:611–5.