ORIGINAL RESEARCH

Initiation And Persistence With Antiplatelet Agents Among The Patients With Acute Coronary Syndromes: A Retrospective, Observational Database Study In China

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Objectives: To investigate the initiation and persistence of antiplatelet agents and the associated factors for patients with acute coronary syndromes (ACS) in Tianjin, China.

Methods: Data were obtained from Tianjin Urban Employee Basic Medical Insurance database (2011-2015). Adult patients who were discharged alive after the first ACS-related hospitalization (index hospitalization) between January 2012 and December 2014 were included. Patients who initiated with antiplatelet therapy, including aspirin monotherapy, clopidogrel monotherapy, or dual antiplatelet with aspirin and clopidogrel at discharge or within the initial 30-day follow-up after discharge were further identified. Patients with no gaps of ≥30 days in antiplatelet therapy were deemed persistent. The logistic model and Cox model were used to explore the associated factors of initiation and persistence with antiplatelet agents, respectively.

Results: In total, 21,450 patients (64.6±10.7 years; 46.0% female) were included. Only 70.3% (N=15,071) of them initiated with antiplatelet agents within the initial 30-day followup; 85.0% (N=12,809) of the initial users discontinued their antiplatelet therapy, and the average time to discontinuation was 117.4±119.7 days. The patients who had prior antiplatelet agents utilization (Odds ratio [95% CI]=1.93 [1.78-2.09]; hazard ratio [95% CI]=0.78 [0.74-0.81]), received percutaneous coronary intervention (PCI) during the baseline period (OR=1.47 [1.26-1.73]; HR=0.91 [0.84-0.97]) or index hospitalization (OR=22.40 [18.63-26.92]; HR=0.51 [0.49-0.53]) were more likely to initiate and persist with antiplatelet agents, while the female (OR=0.75 [0.70-0.81]; HR=1.22 [1.88-1.27]) patients were less likely to initiate and persist with antiplatelet agents.

Conclusion: The initiation and persistence with antiplatelet agents are poor among the ACS patients in Tianjin. Females are associated with poorer initiation and persistence, while prior antiplatelet agents use and receiving PCI during baseline period or index hospitalization are associated with better initial use and better persistence.

Keywords: acute coronary syndromes, antiplatelet agents, initiation, persistence, China

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Introduction

Acute coronary syndrome (ACS), including unstable angina (UA), non-ST segment elevation myocardial infarction (NSTE-MI), and ST segment elevation myocardial infarction (STE-MI), is an umbrella term for situations where the blood supplied to the heart muscle is suddenly blocked. 1-3 Rupture of coronary atherosclerotic plaque, vasospasm, and consequent platelet adhesion, aggregation and secondary thrombosis were the major causes of ACS. With the specific pathological base, the ACS patients face long-term high risk of recurrence and other potential adverse cardiovascular events, even death during secondary prevention. This disease seriously affects the quality and length of life. AS one of the most widespread causes of morbidity and mortality, ACS not only threatens human health seriously but also presents a significant burden for healthcare providers globally. 10–12

Preventing platelet activation and subsequent aggregation are the key targets in the management of ACS.¹³ Guidelines of the American College of Cardiology/ American Heart Association (ACC/AHA), European Society of Cardiology (ESC), and China Heart Society (CHS) all recommend antiplatelet therapy use for ACS patients in the acute setting. 14 For all ACS patients without contraindications, aspirin is recommended for long-term use and a P2Y₁₂ inhibitor, eg, clopidogrel, prasugrel, or ticagrelor, is recommended for up to 12 months. 15 As the first P2Y₁₂ inhibitor which entered the Chinese market, Clopidogrel is the most widely used P2Y₁₂ inhibitor which is recommended by Chinese diagnostic and therapeutic guideline for ACS, 9,16 and is also the only P2Y₁₂ inhibitor reimbursed by the basic medical insurance system in China during the study period.

Many studies suggested that optimal antiplatelet therapy according to the guidelines will decrease the risk of adverse cardiovascular events, such as stroke, myocardial infarction (MI), and all-cause mortality. ^{4,17,18} However, many patients do not initiate with antiplatelet agents after ACS or cannot persist to the regimen prescribed to them. 19,20 The prior studies suggested that 10-20% of ACS patients were left untreated within the initial period after discharge. 21,22 It was also reported that, after 12-month follow-up, about half of the patients did not persist to use antiplatelet agents. 18,23 It is evident that there is room for improvement in the prevention management of ACS. Despite some related studies being reported abroad, 24-26 corresponding observational studies using real-world data are lacking among the Chinese population. This study aimed to evaluate the initiation and persistence of antiplatelet agents and explore their associated factors among patients with ACS in Tianjin, China.

Methods

Data Source

The data in this retrospective study were extracted from Tianjin Urban Employee Basic Medical Insurance (UEBMI)

from January 1, 2011 to December 31, 2015. As the most extensive basic medical insurance system in China, the UEBMI system covers all employed workers and retirees from both public and private sectors in Tianjin. By 2015, the UEBMI covered almost 5.2 million unique members in Tianjin, representing 50.8% of the registered residents.²⁷ The analytical sample in this study was a 30% random sample of all enrollees. The dataset contained the unique patient-level demographic characteristics, pharmacy claims (quantity, strength, date of prescription), medical claims of inpatient and outpatient services, and related medical and medication costs of employees and retirees in Tianjin. The Tolerability and Ethics Committee at the School of Pharmaceutical Science and Technology, Tianjin University, waived the requirement of ethics approval for the current study, as this was a retrospectively observational study using the de-identified medical claims data.

Study Population

Adult patients who were discharged alive with primary discharge diagnosis of ACS (ICD-10 code I20.0, I21, I22 supplemented by Chinese description) from January 1, 2012 through December 31, 2014 were identified. The first qualified hospitalization during the identification period was defined as the index hospitalization. Patients with malignant diseases, who had no continuous insurance enrollment for 12 months before (baseline period) and 12 months after (follow-up period) the index hospitalization, or who experienced major adverse cardiovascular events (MACEs) which included MI, stroke, and death within the initial 30 days after discharge were further excluded. The included patients were followed up from the discharge of index hospitalization until the first MACE or 12 months after this discharge, whichever came first.

Outcome Measures

Baseline Characteristics

Baseline information was reported, including demographic characteristics (age, gender, and employment status), clinical characteristics (Charlson Comorbidity Index [CCI)], medical history during baseline), prior medication utilization (antiplatelet agents, angiotensin-converting enzyme inhibitors/ angiotensin receptor blockers [ACEIs/ARBs], β -blockers, statins and calcium channel blockers [CCB]), prior procedure received (percutaneous coronary intervention, PCI) and health care resource utilization (HRU) and direct medical costs. The procedure received (PCI or not) and the length

of stay (LOS) during the index hospitalization were also described.

Initiation

Initial use of antiplatelet treatment was examined by the presence of the first claimed antiplatelet prescriptions at discharge or within the initial 30 days after the index discharge. Specifically, patients who had at least one prescription of aspirin or clopidogrel during the timeframe mentioned above were defined as the initial users.

Persistence

Persistence was analyzed among the initial users. Patients who had no gap of ≥30 days between the timepoint of exhausting the supplied antiplatelet agents and the following refill during the follow-up period were defined as persistent. Persistence was measured by the rate of discontinuation and persistence duration, ie, time (days) to discontinuation, which was calculated as the number of days between the first day of the initial prescription and the last day of the last prescription before the discontinuation. Overlaps of the prescriptions were not accumulated.

Clopidogrel was the only drug of P2Y₁₂ that can be reimbursed by UEBMI during the study period. Based on the recommend guidelines and the reimbursement policies, the antiplatelet agents in this study mainly consist of aspirin and clopidogrel.

Statistical Analysis

The continuous variables were reported as mean and standard deviation (SD), and the categorical variables were described using frequencies and percentages. Differences between two subcohorts were assessed using Chi-squared tests for categorical variables, and t-tests for continuous variables. Logistic regression model was used to explore the potential factors associated with initiation of antiplatelet agents, and Cox's proportion hazard model was used to explore the associated factors for persistence. Demographics (age, gender, and employment status), clinical characteristics (CCI, medical history, prior medication utilization, and PCI received), baseline HRU and direct medical costs (any hospitalization, any outpatient visit, total direct medical cost, and LOS of index hospitalization) were included as potential associated factors in the regression models. A two-side P-value of <0.05 was considered statistically significant.

All statistics analyses were conducted using STATA version 12.1.

Results

Patient Characteristics

Figure 1 shows the patients flow chart with the number of patients included and excluded in the study. A total of 25,397 adult patients who had a primary discharge diagnosis of ACS were identified between 2012 and 2014, among which 21,450 eligible patients were included as the study population.

In Table 1, the mean (SD) age of the study population was 64.6 (10.7) years, 46.0% were female and 80.5% were retired. The mean (SD) CCI was 1.7 (1.6). The most common co-morbidities during the baseline period among ACS patients were hypertension (75.0%), hyperlipidemia (46.6%), and diabetes (37.5%). Among these patients, 54.7% (N=11,728) had prior use of an antiplatelet prescription during the baseline period, 31.1% patients experienced \geq 1 all-cause hospitalizations during the baseline, and the average annual number of outpatient visits was 30.5 (31.7) among the patients. The mean direct medical cost of the patients was 12,772 (18,893) CNY. The LOS was 10.5 (5.5) days on average and 36.5% of patients received PCI during the index hospitalization.

Initiation With Antiplatelet Treatment And Associated Factors

About 70.3% (N=15,071) of the total patients had the initial antiplatelet prescription within 30 days after discharge, while 29.7% (N=6,379) of patients did not have any antiplatelet agents. The percentages of initiation with aspirin or clopidogrel were similar, which were 47.4% (N=10,160) and 47.7% (N=10,226), respectively, among ACS patients during the initial 30 days (Table 2).

The initial (N=15,071) and non-initial (N=6,379) patient subcohorts differed on most of the baseline characteristics. The result suggested that age, gender, medical history, prior use, baseline MRU and index characteristics were all associated with initiation (Table 3). Compared with patients aged ≤54, patients aged older were more likely to initiate antiplatelet treatment (age 55–64: adjusted Odd Ratio [95% CI]=1.46 [1.31–1.63]; age 65–74: adjusted OR=1.82 [1.61–2.07]; age ≥75: adjusted OR=1.70 [1.49–1.93], respectively). The patients who had higher CCI (adjusted OR=1.04 [1.00–1.07]) had prior MI (adjusted OR=1.18 [1.07–1.30]) or had baseline antiplatelet prescription (adjusted OR=1.93 [1.78–2.09]) were more likely to initiate with antiplatelet treatment. Also, receiving PCI during the baseline period (adjusted

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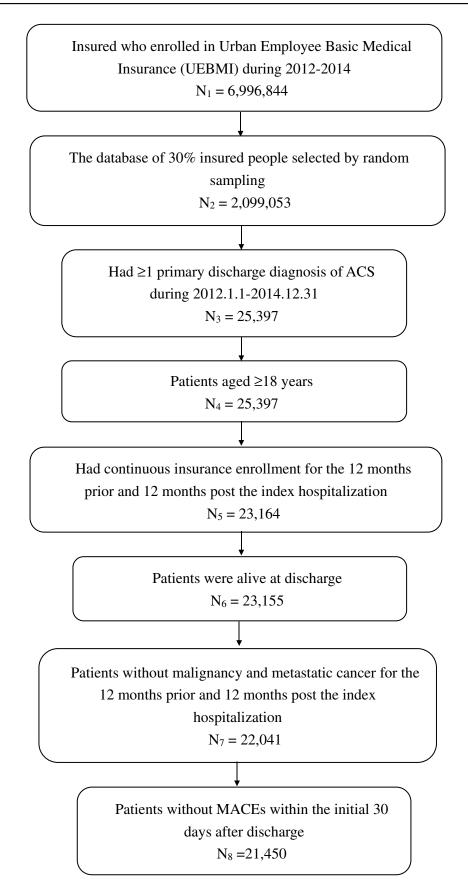


Figure I Flow chart of sample selection.

Table I Baseline Characteristics

Demographic Characteristics	N=21,4	50
Age, Mean (SD)	64.6	10.7
Age group, n (%)		
≤54	3,584	16.7%
55–64	7,670	35.8%
65–74	5,731	26.7%
≥75	4,465	20.8%
Female, n (%)	9,861	46.0%
Retired, n (%)	17,275	80.5%
Charlson Comorbidity Index, Mean (SD)	1.7	1.6
Medical history, n (%)		
Hypertension	16,090	75.0%
Hyperlipidemia	10,006	46.6%
Diabetes mellitus	8,038	37.5%
Prior stroke	5,586	26.0%
Prior angina	5,388	25.1%
Prior myocardial infraction	3,270	15.2%
Prior percutaneous coronary intervention (PCI)	1,418	6.6%
Prior use, ^a n (%)		
Antiplatelet	11,728	54.7%
β-blockers	8,801	41.0%
Calcium channel blockers	12,010	56.0%
Antiplatelet agents, angiotensin-	11,446	53.4%
converting enzyme inhibitors/angiotensin		
receptor blockers		
Statins	9,347	43.6%
Resource utilization and cost		
Any hospitalizations, n (%)	6,668	31.1%
Number of outpatient visits among all patients,	30.5	31.7
Mean (SD)		
Total direct medical cost (CNY), Mean (SD)	12,772	18,893
Index hospitalization		
Any PCI, n (%)	4,771	22.2%
Mean length of stay (days), Mean (SD)	10.5	5.5

Note: ^aHad at least one prescription during the baseline

 Table 2 Initiation with Antiplatelet Therapy Within the 30 Days

 After Discharge

•		
	N=21,4	50
Initiation with antiplatelet therapy, n (%)	15,071	70.3%
Aspirin	10,160	47.4%
Clopidogrel	10,226	47.7%
Dual therapy	5,336	24.9%
Initiation without antiplatelet therapy, n (%)	6379	29.7%

OR=1.47 [1.26-1.73]) or index hospitalization (adjusted OR=22.40 [18.63-26.92]) was associated with an increased possibility of using antiplatelet agents within the initial 30 days after discharge. While compared with

males, the female patients were less likely to initiate antiplatelet agents (adjusted OR=0.75 [0.70-0.81]).

Persistence With Antiplatelet Agents And Associated Factors

As presented by the Kaplan–Meier survival curve (Figure 2), the percentage of non-persistent patients increased with time after discharge. During the follow-up period, 85.0% of patients (N=12,809) had discontinued the antiplatelet therapy and the average time to discontinuation was 117.4 (119.7) days. Almost half of initial users had discontinuation within 3 months after discharge of the index hospitalization.

The associated factors of persistence were analyzed by using Cox regressions (Table 4). Compared with patients aged ≤54, the likelihood of being persistent was lower for patients aged ≥75 years (adjusted Hazard Ratio [95% CI]=1.08 [1.00-1.17]). And, compared with males, female patients (adjusted HR=1.22 [1.18-1.27]) were more likely to discontinue the antiplatelet treatment. Complications of prior hypertension (adjusted HR=1.10 [1.03-1.17]) and hyperlipidemia (adjusted HR=1.06 [1.01-1.11]) were both risk factors of discontinuation with antiplatelet agents. Baseline utilization of β-blockers (adjusted HR=1.05 [1.01-1.10]) or CCB (adjusted HR=1.08 [1.03-1.13]) was related with higher risk of discontinuation, while baseline antiplatelet agents utilization (adjusted HR=0.78 [0.74–0.81]) or stain utilization (adjusted HR=0.91 [0.87-0.96]) was related with lower risk of discontinuation. In addition, receiving PCI during baseline period (adjusted HR=0.91 [0.84-0.97]) or index hospitalization (adjusted HR=0.51 [0.49-0.53]) was associated with significantly lower possibility of discontinuation.

Discussion

To the best of our knowledge, this is the first observational study which presents the initiation and persistence of antiplatelet treatment in ACS population using real-word data in China. We found that both the initial use and subsequent persistence were poor among ACS patients in Tianjin, China; and the patients who were female were less likely to initiate or persist the treatment pattern, while patients who had no prior antiplatelet agents use or had not received PCI during baseline period or index hospitalization were more likely to initiate and persist that. The findings provide useful information on the precise management of ACS in China.

Table 3 Unadjusted And Adjusted Characteristics For Patients Initiated With And Without Antiplatelet Agents (N=21,450)

Patient Characteristics	Initiation With Antiplatelet (N=15,071)	With et)	Initiation Without Antiplatelet (N=6,379)	ithout (N=6,379)	P-value	Adjust	Adjusted Analysis	lysis	
	z	%	z	%		OR	95% CI	_	P-value
Age group (vs ≤54)									
<554	2,267	15.0%	1,317	20.6%	<0.001	l (reference)	rence)		
55–64	5,449	36.2%	2,221	34.8%	0.062	1.46	1.31	1.63	<0.001
65–74	4,253	28.2%	1,478	23.2%	100:0>	1.82	19:1	2.07	<0.001
≥75	3,102	20.6%	1,363	21.4%	610.0	1.70	1.49	1.93	<0.001
Female (vs male)	6,356	42.2%	3,505	54.9%	100:0>	0.75	0.70	0.81	<0.001
Retired (vs Working)	12,130	80.5%	5,145	%2'08	0.774	26:0	0.87	60.1	0.646
Charlson Comorbidity Index	1.8	0:0	9.1	0.0	100:0>	1.04	1.00	1.07	0.032
Medical history (Yes vs No)									
Hypertension	11,475	76.1%	4,615	72.3%	<0.001	66'0	0.88	1.10	0.804
Hyperlipidemia	7,266	48.2%	2,740	43.0%	<0.001	1.00	0.92	1.08	0.949
Diabetes mellitus	5,867	38.9%	2,171	34.0%	100 [.] 0>	96:0	0.89	1.05	0.363
Prior stroke	3,958	26.3%	1,628	25.5%	0.258	86:0	0.90	1.07	0.700
Prior angina	3,779	25.1%	1,609	25.2%	0.818	0.94	0.87	1.03	0.180
Prior myocardial infraction	2,420	%1.91	850	13.3%	<0.001	1.18	1.07	1.30	0.001
Prior percutaneous coronary intervention (PCI)	1,182	7.8%	236	3.7%	<0.001	1.47	1.26	1.73	<0.001
Prior use (Yes vs No)									
Antiplatelet	8,970	29.5%	2,758	43.2%	<0.001	1.93	1.78	2.09	<0.001
β -blockers	6,493	43.1%	2,308	36.2%	<0.001	1.13	1.05	1.22	0.00
Calcium channel blockers	8,579	%6.99	3,431	53.8%	<0.001	0.95	0.87	1.03	0.214
Antiplatelet agents, angiotensin-converting enzyme inhibitors/angiotensin receptor blockers	8,188	54.3%	3,258	%1'15	100:0>	86'0	06:0	90:1	0.599
									,

P-value <0.00 <0.001 <0.001 <0.00 0.903 0.447 26.92 0.98 90: 99.0 1.02 <u>=</u> Adjusted Analysis ប 18.63 82% 0.88 0.55 8. 0.97 22.40 0.60 8 8 0.97 0.97 P-value <0.00 <0.00 <0.00 <0.00 <0.00 <0.00 Antiplatelet (N=6,379) 6,012 39.4% 35.5% Initiation Without 26.8 %: '' -. % 11,358.0 2,263 2,514 25.5 133 z 19,958.4 45.3% 29.2% 30.8% 33.4 Initiation With 0.0 Antiplatelet (N=15,071)13,370. 6,833 4,648 4,405 32.6 0.7 z Total baseline all-cause direct medical cost Any baseline inpatient visits (yes vs no) Length of stay during the index event Number of baseline outpatient visits Patient Characteristics Index PCI (Yes vs No) Statins

proportion of patients continuing antiplatelet treatment 0.00 0.25 0.50 0.75 1.00 85.0% (N=12,809) of initial users had discontinuation and the average time to discontinuation was 117.4±119.7 days 400 200 300 time since first prescription (days)

Figure 2 Persistence with antiplatelet agents during the follow-up period.

Compared with the results of previous studies from different countries, the proportion of initiation of antiplatelet agents was relatively lower (70%) within the 30 days after discharge and the proportion of discontinuation was much higher (85%) during the 12 months after discharge in the present study. In the previous studies, it was reported that more than 90% of ACS patients possessed prescription of antiplatelet agents at discharge. 18,22,28 As for persistence, the previous retrospective studies using a large observational database from different countries in Europe indicated that 50-73% of patients had discontinuation of antiplatelet agents with gaps >30 days during 1year follow-up in patients after an ACS. 18,29 There are many possible reasons for the poorer initiation and persistence in this study. First, physicians may be unfamiliar with treatment recommendations and/or unaware of the importance of preventive treatment. According to the previous studies, "not indicated" by the physician was recorded as the most common reason for not prescribing medications, which means that the medication was not prescribed because of a perceived lack of indication rather than the presence of a contraindication or intolerance. ^{21,30} Second, heavy economic burden of ACS is a potential barrier. Despite the universal health insurance coverage in China, patients still need to pay a proportion of total medical costs (ie, out-of-pocket cost). It reported that the mean annual ACS-related direct medical costs was 8,531 CNY among the Chinese ACS patients, while the per capita disposable income was only 18,311 CNY.³¹

Compared with males, female patients were less likely to initiate with antiplatelet treatment but more likely to discontinue. The results were consistent with the previous

Table 3 (Continued).

 Table 4
 Unadjusted and Adjusted Characteristics for Patients Who Were Persistent and Non-Persistent Antiplatelet Agents (N=15,071)

Patient Characteristics	Persistent (N=2.262)	(N=2.262)	Non-Persister	Non-Persistent (N=12,809)	P-value	Adius	Adjusted Analysis	alvsis	
		(======================================		(222(=: 21) 21					
	z	%	Z	%		HR	95% CI		P-value
Age group (vs ≤54)									
<554	345	15.3%	1,922	15.0%	0.762	I (refe	l (reference)		
55–64	885	39.1%	4,564	35.6%	0.001	96.0	06:0	1.02	0.179
65–74	626	27.7%	3,627	28.3%	0.532	0.98	16:0	1.05	0.548
≥75	406	17.9%	2,696	21.0%	0.001	1.08	00:	1.17	0.044
Female (vs male)	782	34.6%	5,574	43.5%	<0.001	1.22	81.	1.27	<0.001
Retired (vs Working)	1,782	78.8%	10,348	80.8%	0.026	1.05	66:0	1.12	0.129
Charlson Comorbidity Index	2.6	0.0	1.7	0.0	<0.001	0.99	0.97	00.1	0.145
Medical history (Yes vs No)									
Hypertension	1,794	79.3%	9,681	75.6%	<0.001	1.10	1.03	1.17	0.003
Hyperlipidemia	1,229	54.3%	6,037	47.1%	<0.001	1.06	1.01	1.1	0.010
Diabetes mellitus	1,276	56.4%	4,591	35.8%	<0.001	96.0	16:0	00.1	0.068
Prior stroke	793	35.1%	3,165	24.7%	<0.001	1.02	0.97	1.07	0.523
Prior angina	920	28.7%	3,129	24.4%	<0.001	1.01	96.0	1.05	0.731
Prior myocardial infraction	396	17.5%	2,024	88'51	0.042	1.05	1.00	1:1	0.064
Prior percutaneous coronary intervention (PCI)	239	%9:01	943	7.4%	<0.001	16.0	0.84	0.97	0.007
Prior use (Yes vs No)									
Antiplatelet	1,675	74.0%	7,295	27.0%	<0.001	0.78	0.74	0.81	<0.001
β -blockers	1,009	44.6%	5,484	42.8%	0.112	1.05	1.01	1.10	0.011
Calcium channel blockers	1,391	61.5%	7,188	26.1%	<0.001	1.08	1.03	1.13	0.001
Antiplatelet agents, angiotensin-converting enzyme inhibitors/angiotensin receptor blockers	1,377	60.9%	6,811	53.2%	<0.001	0.98	0.94	1.03	0.391
Statins	1,254	55.4%	5,579	43.6%	<0.001	16.0	0.87	96.0	<0.001

Table 4 (Continued).

Patient Characteristics	Persistent	(N=2,262)	Non-Persister	Persistent (N=2,262) Non-Persistent (N=12,809) P-value Adjusted Analysis	P-value	Adjus	ted Ar	ıalysis	
	z	%	z	%		HR	15% CI		P-value
Any baseline inpatient visits (yes vs no)	818	36.2%	3,587	28.0%	<0.001 0.95 0.90 1.00 0.035	0.95	06.0	00:1	0.035
Number of baseline outpatient visits	54.9	52.2	28.6	27.0	<0.001	0.99	66'0 66'0	66.0	100:0>
Total baseline all-cause direct medical cost	22,445.4	22,445.4 25,437.9 11,767.5	11,767.5	18,367.0	<0.001 1.04 1.03 1.05 <0.001	1.04	1.03	1.05	<0.001
Index PCI (Yes vs No)	626	43.3%	3,669	28.6%	<0.001 0.51 0.49 0.53 <0.001	0.51	0.49	0.53	<0.001
Length of stay during the index event	6.01	0.1	10.1	0.0	100.0> 00.1 69.0 69.0 100.0>	66'0	0.99	1.00	<0.001

studies in Finland and Denmark.^{18,32} The possible reason for this finding is that females have a higher risk of bleeding and refractory ischemia than males.^{32,33} Despite the guidelines recommending clopidogrel treatment to all patients regardless of gender, treatment with antiplatelets should be considered for every individual fulfilling indications.^{32,34} The results suggested that physicians should pay more attention to the antiplatelet management of females.

The study found that patients who had any prescription of antiplatelet agents during the baseline period were more likely to initiate and persist the follow-up antiplatelet treatment. A Finnish study also reported that the patients who used antiplatelet agents during the baseline period were 3.7-times more likely of initiating antiplatelet after ACS and half the risk of discontinuing than those who had no records of antiplatelet utilization. The patients who used antiplatelet agents prior to index hospitalization indicated that they did not have any contraindication, allergy history, or adverse drug reaction. They may have already formed a good habit for antiplatelet agents utilization, which is a possible reason for higher compliance with the guidelines.

The results from this study suggested receiving PCI during the index hospitalization was positively related to both initiation and persistence, which was consistent with previous studies that reported patients who received PCI during index hospitalization had a nearly 20-fold possibility of initiation with antiplatelet agents and faced lower risk of non-persistence than those who did not receive PCI. 18,23 The results may be related to the guidelines in China. The most updated international guidelines for ACS during the study period recommend that all the patients with ACS receiving PCI or not should use the antiplatelet treatment of aspirin and clopidogrel from the discharge of the first ACS up to 12 months. 1,15 Despite the Chinese guidelines recommending that all patients with STE-MI should use the aspirin and clopidogrel whether with or without PCI, the clopidogrel only be recommend for the NSTE-MI/UA patients with PCI.^{9,16} In clinical practice, physicians usually highly recommend the DAPT for the ACS patients with PCI in China. This may explain why the patients with PCI tend to have better initiation and persistence in the present study.

The present study does have some limitations. First, aspirin could be obtained OTC and ticagrelor was not reimbursed by UEBMI (during the study period) and this may have caused the underestimation. However, because

of the high price without reimbursement by insurance, the market share of other P2Y₁₂ inhibitors was very limited during the study period in China, which resulted in minimal influence on the study. Secondarily, the claim database has no detailed information of socioeconomics, living habits, and disease severity, thus the analyses exploring the factors associated were not able to control these unmeasured factors, which may be associated with initiation and persistence. Furthermore, due to the limitation of the database, estimates of both initiation and persistence in this study were based on patterns of drugs dispensed, but not necessarily consumed. Additionally, the generalizability is limited due to data from an insured population under employment and those who had retired in Tianjin. Although there are some limitations, this study provides real-world evidence that helps discover problems and promote management of secondary prevention for ACS.

Conclusion

The initiation of antiplatelet therapy is unsatisfactory among the ACS patients in Tianjin, China, and the persistence to antiplatelet is poor among the initial users. It represents suboptimal treatment practices as well as opportunities for improvement. The study found that patients who are female, have no prior antiplatelet agent utilization, and do not receive PCI during the baseline period or index hospitalization are the population at risk of non-initiation and non-persistence with antiplatelet treatment, and who need more attention from the physicians.

Disclosure

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