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# Impact of routine invasive strategy on outcomes in patients with non-ST-segment elevation myocardial infarction during 2005–2014: A report from the Polish Registry of Acute Coronary Syndromes (PL-ACS)

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### **Abstract**

**Background:** Non-ST-segment elevation myocardial infarction (NSTEMI) has become the most frequently encountered type of myocardial infarction. The patient clinical profile and management has evolved over the past decade. As there is still a scarcity of data on the latest trends in NSTEMI, changes herein were observed and assessed in the treatment and outcomes in Poland between 2005 and 2014. **Methods:** A total of 197,192 patients with NSTEMI who enrolled in the Polish Registry of Acute Coronary Syndromes (PL-ACS) between 2005 and 2014 were analyzed. In-hospital and 12-month mortality were assessed.

**Results:** Coronary angiography use increased from 35.8% in 2005–2007 to 90.7% in 2012–2014 (p < 0.05), whereas percutaneous coronary intervention increased from 25.7% in 2005–2007 to 63.6% in 2012–2014 (p < 0.05). There was a 50% reduction in in-hospital mortality (from 5.6% in 2005–2007 to 2.8% in 2012–2014; p < 0.05) and a 30% reduction in 1-year mortality (from 19.4% in 2005–2007 to 13.7% in 2012–2014; p < 0.05). A multivariate analysis confirmed an immense impact of invasive strategy on patient prognosis during in-hospital observation with an odds ratio (OR) of 0.31 (95% confidence interval [CI] 0.29–0.33; p < 0.05) as well as during the 12-month observation with an OR of 0.51 (95% CI 0.49–0.52; p < 0.05).

**Conclusions:** Over the past 10 years, an important advance in the management of NSTEMI has taken place in Poland. Routine invasive strategy resulted in a significant decrease in mortality rates in all groups of NSTEMI patients. (Cardiol J XXXX; XX, X: xx–xx)

Key words: non-ST-elevation myocardial infarction, invasive strategy, percutaneous coronary intervention, outcomes, temporal trends

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# Introduction

Non-ST-segment elevation myocardial infarction (NSTEMI) represents the majority of all MI cases. In Western Europe, NSTEMI accounts for over 60% of all MI cases. Dynamic changes in the clinical profile and treatment strategy have been observed in recent years. Contemporary analyses for a wide national population are scarce.

The analysis of clinical characteristics, treatment strategies and outcomes in almost 200,000 NSTEMI cases registered in the Polish Registry of Acute Coronary Syndromes (PL-ACS) between 2005 and 2014 are presented.

# **Methods**

The study population was drawn from 463 hospitals in Poland that provide care to patients with MIs. The population consists of patients admitted with a diagnosis of NSTEMI according to the European Society of Cardiology guidelines [1, 2]. The study covers a 10-year period from 2005 to 2014. Contribution to the study was voluntary; nevertheless, half of all estimated NSTEMI cases in Poland during the study time period were included. The study complies with the Declaration of Helsinki and was approved by the PL-ACS Registry committee.

Data were collected from PL-ACS Registry questionnaires that include variables on demographic factors (gender, age), risk factors (smoking, hypertension, hypercholesterolemia, diabetes mellitus and obesity), previous coronary incidences and related procedures (MI, percutaneous coronary intervention [PCI], coronary artery bypass grafting), clinical presentation on admission (Killip class, heart rate, systolic blood pressure), electrocardiographic abnormalities, left ventricular ejection fraction, coronary angiography (CA), coronary intervention details, and in-hospital and post-discharge treatment. The mortality rate was evaluated for the in-hospital observation period as well as for 30-day, 6-month and 12-month followup periods.

# Statistical analysis

The gender groups were analyzed separately and subsequently compared to each other. To investigate the impact of age on outcomes, the analysis was conducted in age groups (< 55, 55–64, 65–74, ≥ 75 years) as well as in consecutive decades of life. Changes over time were investigated using the following two models: a comparison between subgroups in marginal 3-year intervals (2005–2007

and 2012–2014) and an evaluation of temporal trends over a 10-year period.

Categorical data are presented as numbers and percentages while continuous data are presented as the median or arithmetic mean  $\pm$  standard deviation (SD). Differences in categorical variables were tested using the  $\chi^2$  test with Pearson modification, whereas in continuous variables differences were tested with the Student t-test. A two-sided p value  $\leq 0.05$  was considered significant. Trend importance was verified by the Cochran-Armitage test for categorical data and the Jonckheere-Terpstra test for continuous data. A logistic regression was used to identify variables that independently contributed to mortality.

### Results

A total of 197,192 patients (including 77,550 women, 39.3%) hospitalized in Poland due to NSTEMI between 2005 and 2014 were included in the analysis. The contribution of younger patients (under 55) decreased compared to older patients. The average age of males increased from 65 to 66 while in women it was stable at 72. Nevertheless, men predominated in the group under 70, whereas women did in the group over 70.

The frequency of diabetes, arterial hypertension, obesity (in men only), and smoking (in women only) also increased. A history of prior coronary artery interventions (especially PCI) was more common in the later years were observed in this study (Table 1).

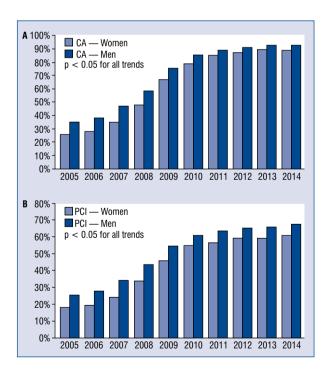
Over the last decade, NSTEMI treatment strategy has changed significantly. The frequency of coronary angiography increased from 35.8% in 2005-2007 up to 90.7% in 2012-2014; p < 0.05. From 2012 to 2014, the gender disparity in CA implementation was still visible with 88.4% in women vs. 92.1% in men; p < 0.05. PCI use increased from 25.7% in 2005–2007 to 63.6% in 2012–2014; p < 0.05. From 2012 to 2014 we achieved 59.6% in women and 66.1% in men; p < 0.05. The most intensive growth of an invasive procedure use took place between 2007 and 2011. In later years, only a mild further increase was observed. In 2014, only 10.9% of women and 7.1% of men (p < 0.05) were treated conservatively. Temporal trends in the invasive treatment of NSTEMI patients are presented in Figure 1.

The age group analysis revealed that the percentage of invasive treatment increased most in the oldest patients (over 75 years) and achieved a level of 80% in CA and 50% in PCI. Although differences

Table 1. Clinical characteristics of non-ST-elevation myocar	ial infarction patients
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Clinical characteristics	characteristics 2005–2007		2012–2014			2005–2007 vs. 2012–2014		
_	Women	Men	Р	Women	Men	Р	Women	Men
Age < 55 years	1738 (7.5%)	6162 (18.6%)	< 0.05	1647 (6.4%)	5468 (13.3%)	< 0.05	< 0.05	< 0.05
Age ≥ 7 years	11208 (48.3%)	8746 (26.4%)	< 0.05	12098 (47.4%)	11571 (28.1%)	< 0.05	< 0.05	< 0.05
Hypertension	17908 (77.2%)	22792 (68.8%)	< 0.05	20568 (80.5%)	31219 (75.9%)	< 0.05	< 0.05	< 0.05
Diabetes	8180 (35.3%)	9623 (23.7%)	< 0.05	7865 (37.3%)	11999 (29.2%)	< 0.05	< 0.05	< 0.05
Hyperlipidemia	10182 (43.9%)	11264 (43.6%)	0.43	1446 (44.1%)	18067 (43.9%)	0.67	0.67	0.33
Current smoking	2403 (10.4%)	10595 (32.0%)	< 0.05	3340 (13.1%)	10989 (26.7%)	< 0.05	< 0.05	< 0.05
Obesity	5879 (25.4%)	5143 (15.5%)	< 0.05	6391 (25.0%)	7807 (19.0%)	< 0.05	0.40	< 0.05
Prior MI	5899 (25.4%)	10097 (30.5%)	< 0.05	5681 (22.2%)	10728 (26.1%)	< 0.05	< 0.05	< 0.05
Prior PCI	736 (3.2%)	1680 (5.1%)	< 0.05	4301 (16.8%)	8534 (20.8%)	< 0.05	< 0.05	< 0.05
Prior CABG	1321 (5.7%)	2764 (8.3%)	< 0.05	1092 (4.3%)	2755 (6.7%)	< 0.05	< 0.05	< 0.05

CABG — coronary artery bypass grafting; MI — myocardial infarction; PCI — percutaneous coronary intervention

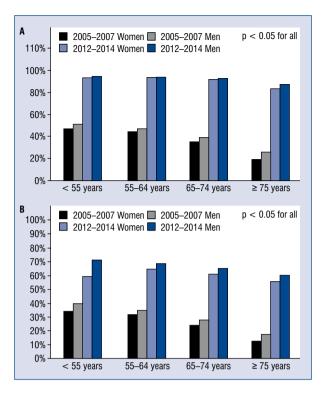


**Figure 1.** Temporal trends in utilization of coronary angiography (CA) (**A**) and percutaneous coronary intervention (PCI) (**B**).

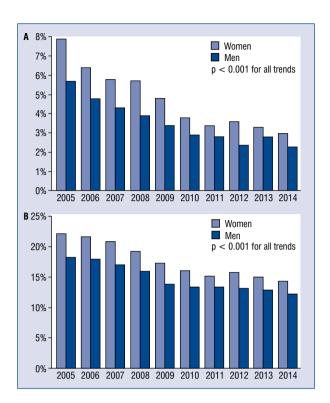
among the age groups diminished during the study, a slight disproportion still existed. In all age groups, invasive strategies were more widespread in men than in woman. Nevertheless, this disproportion has decreased in recent years (Fig. 2).

There have been significant changes in pharmacotherapy over the last decade, especially in the use of antiplatelet agents. The utilization of P2Y12-receptor blockers substantially increased from 51.0% in women and 59.1% in men from 2005 to 2007 to 92.9% in women and 93.2% in men (p < 0.05). Ticlopidine, which was commonly used, was almost completely substituted by clopidogrel. In addition, during the last years a continuous shift from clopidogrel to ticagrelol or prasugrel was observed.

The management outcomes of NSTEMI patients have improved considerably over the last decade. In the present analysis, the risk of reinfarction was reduced from 4.5% in 2005–2007 to 0.3% in 2012–2014 (p < 0.05), and the risk of stroke was reduced from 0.5% in 2005–2007 to 0.2% in 2012–2014 (p < 0.05). On the other hand, a side effect of the intensive invasive treatment application was observed, especially in the frequency of



**Figure 2.** Percentage of coronary angiography (CA) (**A**) and percutaneous coronary intervention (PCI) (**B**) in age groups.



**Figure 3.** Temporal trends of in-hospital mortality (A) and 12-month mortality (B).

Table 2. Mortality rates by gender.

Mortality rates	ates 2005–2007		2012–2014			2005–2007 vs. 2012–2014		
	Women	Men	Р	Women	Men	Р	Women	Men
In-hospital mortality	1541 (6.6%)	1633 (4.9%)	< 0.05	851 (3.3%)	1026 (2.5%)	< 0.05	< 0.05	< 0.05
30-day mortality	2267 (9.8%)	2571 (7.8%)	< 0.05	1686 (6.6%)	2101 (5.1%)	< 0.05	< 0.05	< 0.05
6-month mortality	3988 (17.2%)	4586 (13.8%)	< 0.05	2997 (11.7%)	3994 (9.7%)	< 0.05	< 0.05	< 0.05
12-month mortality	5005 (21.6%)	5897 (17.8%)	< 0.05	3865 (15.1%)	5258 (12.8%)	< 0.05	< 0.05	< 0.05

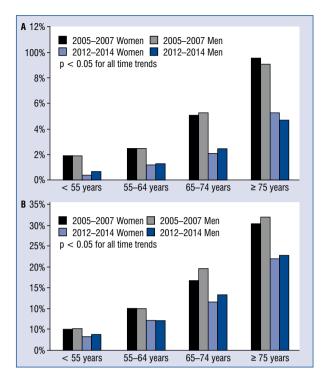
major bleeding (0.7% in 2005-2007 vs. 1.2% in 2012-2014; p < 0.05).

In-hospital mortality decreased by 50% (from 5.6% in 2005–2007 to 2.8% in 2012–2014; p < 0.05) and 1-year mortality by up to 30% (from 19.4% in 2005–2007 to 13.7% in 2012–2014; p < 0.05) (Fig. 3). These positive tendencies are apparent in all age groups and genders (Table 2). Mortality rates in the four age groups (under 55, 55–64, 65–74, 75 and over) are presented in Figure 4.

A multivariable analysis showed that the impact of the invasive strategy on mortality decrease was immense, with a 3-fold improvement in outcomes in short-term observation and a 2-fold improvement in long-term observation (Tables 3 and 4).

# Discussion

The clinical characteristics and management of NSTEMI patients together with treatment



**Figure 4.** In-hospital mortality (**A**) and 12-month mortality (**B**) in age groups.

outcomes have significantly changed all over the world in recent years [3–5]. In Poland, as in many other countries, NSTEMI has become the most common type of MI.

According to many previous reports on NSTE-MI, there are common trends in the baseline clinical profile of patients. The prevalence of major atherosclerosis risk factors like arterial hypertension, diabetes, obesity and chronic kidney disease is still increasing [4, 5]. Similar trends were observed in the present study. Contrary to findings in other countries, the mean age of NSTEMI patients in Poland increased, especially in men. Presumably this is a result of a noticeable smoking decrease and better pharmacological risk factor control, i.e., hypercholesterolemia and arterial hypertension. Over the last 10 years, some important changes in medical therapy have taken place. The vast majority of patients received P2Y12-receptor blockers. Interestingly, from 2005 to 2007 many patients were still administered ticlopidine, which was gradually substituted by clopidogrel and later by new drugs such as ticagrelol, according to the European Society of Cardiology guidelines [1, 2].

Table 3. Multivariate analysis (in-hospital mortality).

	Р	OR	OR (95% CI)
Invasive treatment	< 0.0001	0.31	0.31 (0.29–0.33)
Hypercholesterolemia	< 0.0001	0.73	0.73 (0.69–0.77)
Hypertension	< 0.0001	0.73	0.73 (0.69–0.78)
Previous PCI	< 0.0001	0.80	0.80 (0.73-0.88)
Previous CABG	0.0006	0.80	0.80 (0.71–0.91)
Current smokers	0.6776	1.02	1.02 (0.94–1.10)
Female (vs. male)	0.4485	1.02	1.02 (0.97–1.08)
Previous MI	0.0255	1.07	1.07 (1.01–1.14)
Diabetes	0.0021	1.09	1.09 (1.03–1.15)
Time to admission > 12 h	0.0030	1.09	1.09 (1.03–1.16)
LVEF 35-50%	0.0240	1.10	1.10 (1.01–1.20)
ST-T abnormalities in ECG	0.0007	1.16	1.16 (1.07–1.27)
Obesity	< 0.0001	1.18	1.18 (1.10–1.26)
No sinus rhythm in ECG	< 0.0001	1.19	1.19 (1.12–1.27)
Age (on each decade)	< 0.0001	1.63	1.63 (1.59–1.68)
LVEF < 35%	< 0.0001	2.31	2.31 (2.11–2.53)
Prehospital cardiac arrest	< 0.0001	2.37	2.37 (2.09–2.69)
Killip 3 class	< 0.0001	3.67	3.67 (3.41–3.94)
IABP	< 0.0001	3.89	3.89 (3.23-4.69)
Killip 4 class	< 0.0001	13.17	13.2 (12.0–14.4)

CABG — coronary artery bypass grafting; CI — confidence interval; ECG — electrocardiogram; IABP — intraaortic balloon pump; LVEF — left ventricular ejection fraction; MI — myocardial infarction; OR — odds ratio; PCI — percutaneous coronary intervention

Table 4. Multivariate analysis (12-month mortality).

	P	OR	OR (95% CI)
Invasive treatment	< 0.0001	0.51	0.51 (0.49–0.52)
Hypercholesterolemia	< 0.0001	0.81	0.81 (0.79–0.83)
Previous CABG	< 0.0001	0.84	0.84 (0.80-0.88)
Hypertension	< 0.0001	0.85	0.85 (0.83-0.88)
Previous PCI	< 0.0001	0.90	0.90 (0.87-0.94)
Female (vs. male)	< 0.0001	0.94	0.94 (0.92-0.97)
Obesity	0.37	0.99	0.99 (0.96–1.02)
Time to admission > 12 h	0.02	1.02	1.03 (1.00–1.06)
Current smokers	0.0005	1.07	1.06 (1.03–1.10)
Previous MI	< 0.0001	1.12	1.12 (1.09–1.15)
ST-T abnormalities in ECG	< 0.0001	1.14	1.15 (1.11–1.19)
No sinus rhythm in ECG	< 0.0001	1.15	1.14 (1.11–1.18)
Diabetes	< 0.0001	1.29	1.29 (1.26–1.32)
LVEF 35-50%	< 0.0001	1.52	1.52 (1.47–1.57)
Age (on each decade)	< 0.0001	1.57	1.57 (1.55–1.59)
Prehospital cardiac arrest	< 0.0001	1.74	1.74 (1.63–1.85)
Killip 3 class	< 0.0001	1.98	1.98 (1.91–2.06)
IABP	< 0.0001	2.17	2.17 (1.99–2.38)
LVEF < 35%	< 0.0001	2.67	2.67 (2.57–2.78)
Killip 4 class	< 0.0001	4.48	4.48 (4.26–4.71)

CABG — coronary artery bypass grafting; CI — confidence interval; ECG — electrocardiogram; IABP — intraaortic balloon pump; LVEF — left ventricular ejection fraction; MI — myocardial infarction; OR — odds ratio; PCI — percutaneous coronary intervention

The European Society of Cardiology-recommended implementation of a routine invasive strategy has been the most important change in NSTEMI treatment over the last 10 years [1, 2]. A rapid increase in the number of 24/7 catheterization centers in Poland (up to 150) enabled the successful introduction of this recommendation. The percentage of CAs as well as PCIs in NSTEMI patients in Poland reached the same level as countries in Western Europe such as France [4] and Denmark [6] as well as the United States [5]. Importantly, previously reported underutilization of an invasive strategy in women as well as in older patients was not as pronounced in Poland [7–11]. The significant advances in treatment have resulted in a spectacular decrease in mortality rates at an even higher rate than in previous analyses [3–5, 12, 13].

In the multivariable analysis, the invasive strategy was the most important factor contributing to a better prognosis for in-hospital and 12-month observations. The advantages of invasive treatment were apparent in all patients regardless of age or gender. The final results are comparable to data from other countries that have successfully introduced contemporary guidelines [14].

# Limitations of the study

This study has several limitations. PL-ACS is a voluntary, observational study, and not all hospitals treating NSTEMI in Poland participated in data collection. The present analysis was retrospective, and some potentially important parameters might not be included. Finally, this is a single country study; therefore, some trends should be interpreted with caution.

# **Conclusions**

In Poland, outcomes of NSTEMI patients have improved substantially over the last 10 years due to the implementation of routine invasive treatment. The invasive approach was beneficial to all age groups and both genders.

Conflict of interest: None declared

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