

Stent Selection for Primary Angioplasty and Outcomes in the Era of Potent Antiplatelets. Data from the Multicenter Randomized Prague-18 trial.

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Introduction

The current generation of **drug-eluting stents (DES)** has been shown to be superior to **bare-metal stents (BMS)** in reducing the risk of recurrent myocardial infarction (MI), stent thrombosis, and target lesion revascularization.

However, the use of **bioresorbable vascular scaffolds (BVS)** has been hypothesized to overcome the limitations of DES

Task Force on Myocardial Revascularization of the European Society of Cardiology (2018) **recommended that BVS should not be used outside well-controlled clinical studies**

Aims

- **why different types of stents were used in AMI patients** who underwent primary angioplasty
- **how it influenced the prognoses** of the study population.
- The **efficacy and safety outcomes** of the different stent types were also compared in patients treated with **prasugrel vs. ticagrelor**

Prasugrel versus Ticagrelor in Patients with Acute Myocardial Infarction Treated with Primary Percutaneous Coronary Intervention: Multicenter Randomized PRAGUE-18 Study

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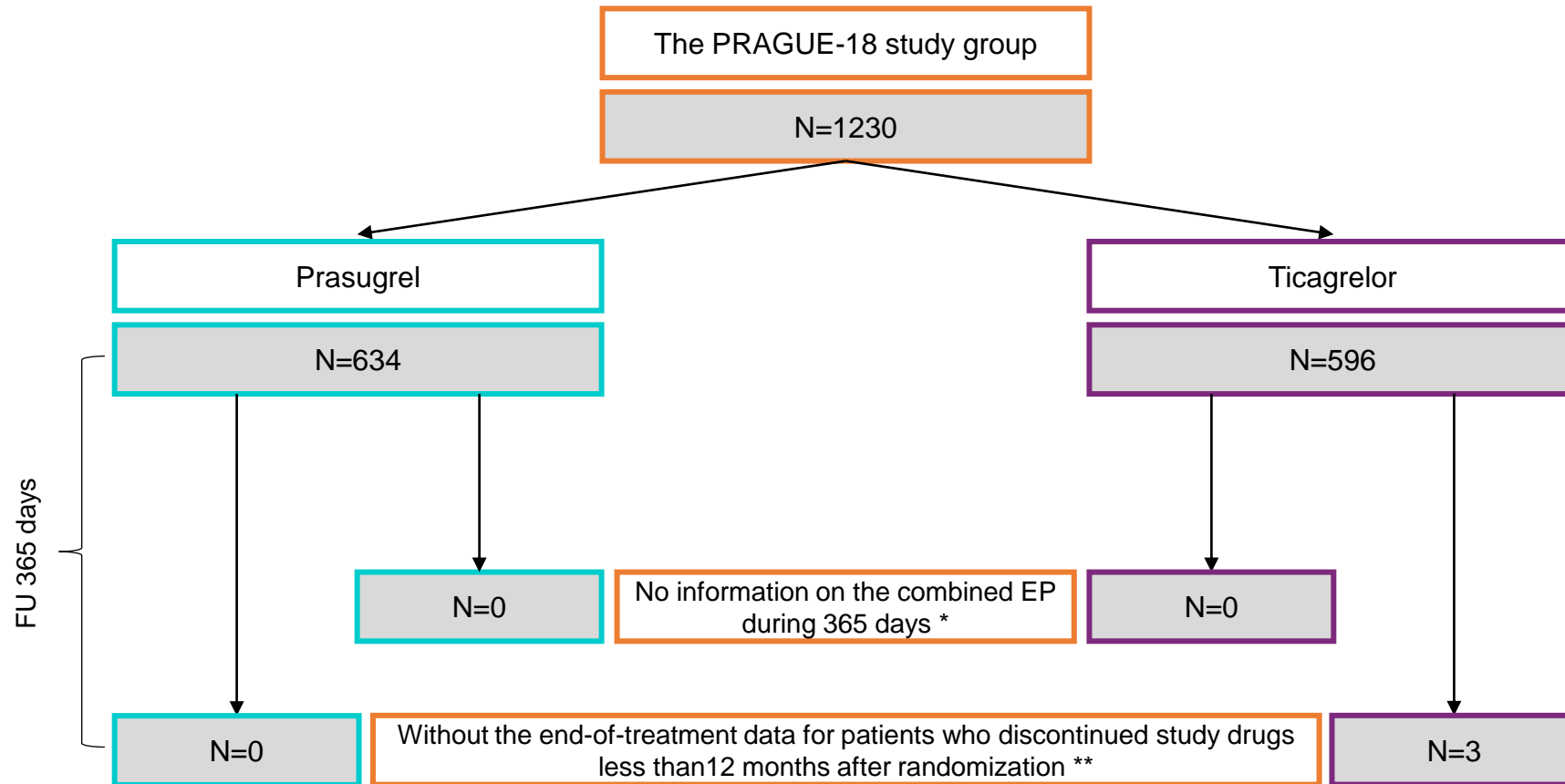
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ORIGINAL INVESTIGATIONS

1-Year Outcomes of Patients Undergoing Primary Angioplasty for Myocardial Infarction Treated With Prasugrel Versus Ticagrelor

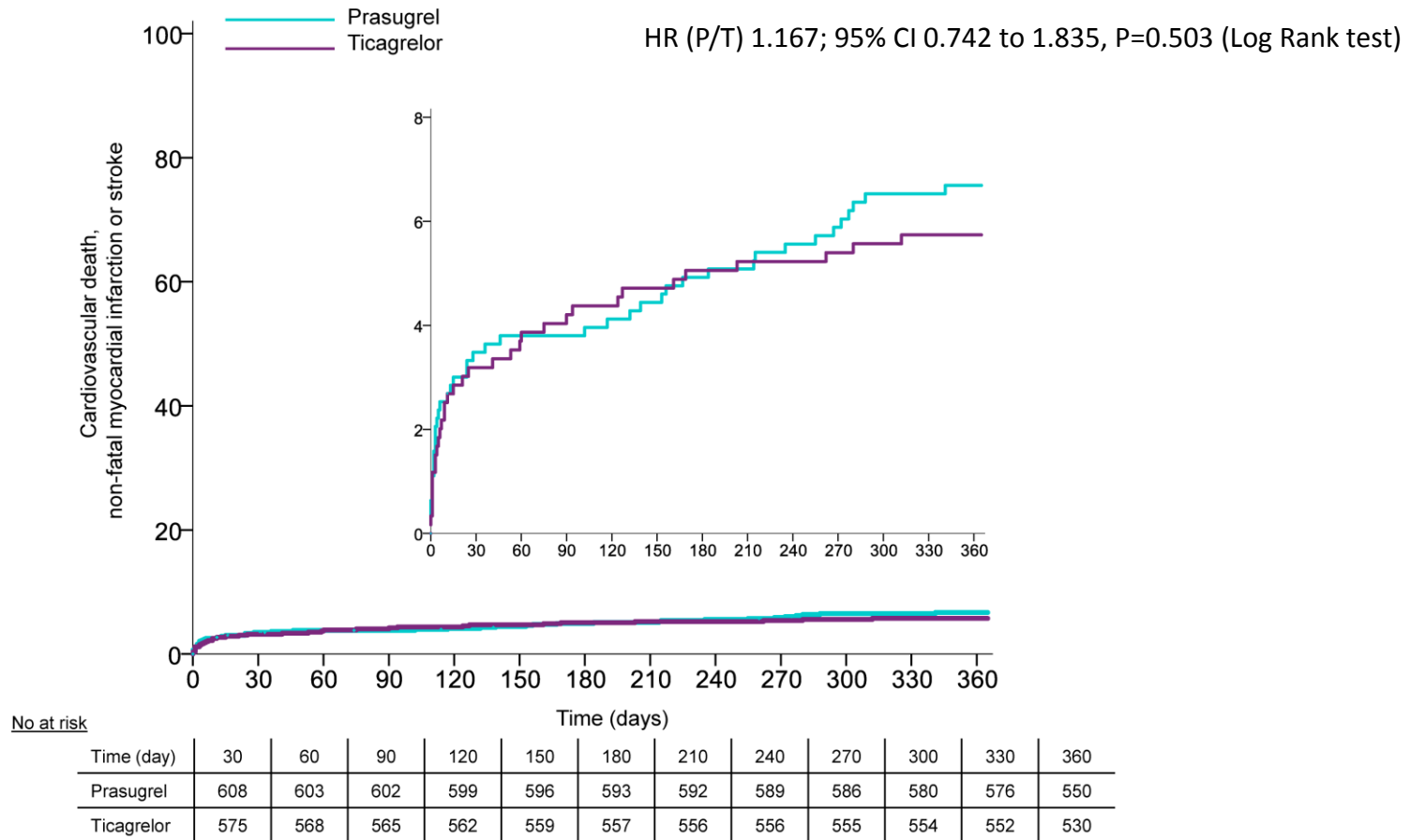


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PRAGUE-18 Study Group



* The combined efficacy endpoint (EP) = Cardiovascular death, Non-fatal myocardial infarction, Stroke: Missing information in 19 patients were supplemented from national registries of the Institute of Health information and Statistics of the Czech Republic.
 ** For missing end-of-treatment data in 3 patients, a visit data were added for which treatment discontinuations were reported.

KEY EFFICACY ENDPOINT: CV Death/Non-fatal MI/Stroke



END POINTS

	Prasugrel	Ticagrelor	P-value
CV Death, Non-fatal MI or Stroke	42 (6.6%)	34 (5.7%)	0.503
Death from cardiovascular causes	21 (3.3%)	18 (3.0%)	0.769
Non-fatal myocardial infarction	19 (3.0%)	15 (2.5%)	0.611
Stroke	7 (1.1%)	4 (0.7%)	0.423
Definite stent thrombosis	7 (1.1%)	9 (1.5%)	0.535
Death from any cause	30 (4.7%)	25 (4.2%)	0.654
Bleeding	69 (10.9%)	66 (11.1%)	0.930
TIMI major	6 (0.9%)	4 (0.7%)	0.754
BARC \geq 3	15 (2.4%)	9 (1.5%)	0.308

		HR (95% CI)	P-value
Risk of ischemic endpoint *	Economically motivated switch (N=481)	0.433 (0.210–0.894)	0.024
	Switch from other reasons (N=178)	3.420 (1.823–6.415)	<0.001
Risk of bleeding	Economically motivated switch (N=481)	0.416 (0.246–0.701)	0.001

* Cardiovascular death, non-fatal myocardial infarction or stroke.

The hazard ratio was based on the Cox proportional hazard model with time dependent covariates

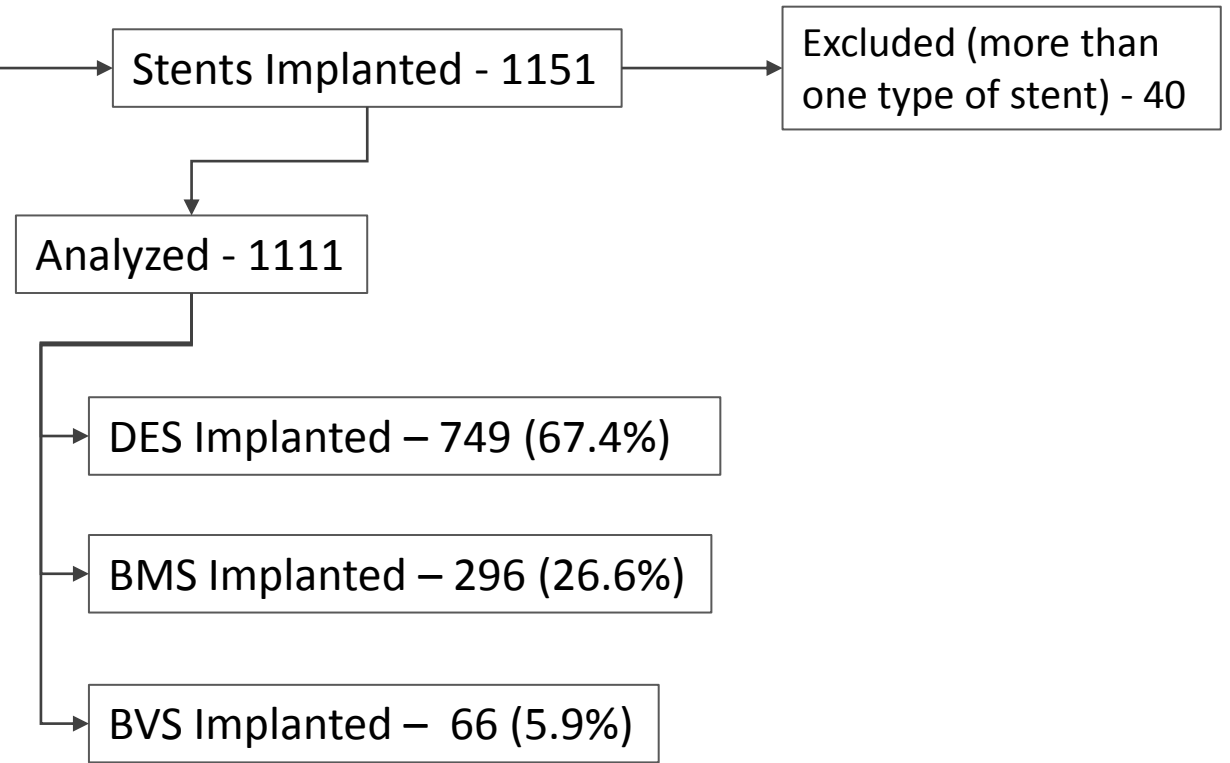
CONCLUSIONS

- 1) Prasugrel and Ticagrelor are similarly effective and safe during the first year after MI
- 1) Economically motivated, early post-discharge switch to clopidogrel, when approved by treating physicians, was not associated with increased risk of ischemic events

Methods

PRAGUE-18 study:

- 14 sites in the Czech Republic
- Between April 2013 – May 2016
- 1230 patients enrolled



Baseline Characteristics

		Stent			p-Value
		DES (n = 749)	BMS (n = 296)	BVS (n = 66)	
Basic characteristics					
Gender—male		574 (76.6%)	223 (75.3%)	45 (68.2%)	0.292
Age		61.7 (42.9; 78.1)	62.7 (46.7; 81.5)	56.9 (40.8; 71.9)	<0.001
BMI		27.8 (22.3; 36.1)	28.3 (22.7; 36.3)	26.4 (21.2; 35.9)	0.022
Laboratory results					
Urea		5.2 (3.1; 9.0)	5.4 (3.4; 9.7)	4.9 (2.7; 8.4)	0.011
Creatinine		82.0 (55.0; 124.0)	85.0 (54.0; 136.0)	73.0 (47.0; 106.0)	<0.001
Risk factors and comorbidities					
Obesity		155 (20.7%)	53 (17.9%)	6 (9.1%)	0.05
Smoking		485 (64.8%)	179 (60.5%)	52 (78.8%)	0.016
Killip class	1	667 (89.1%)	253 (85.5%)	64 (97.0%)	0.041
	2	50 (6.7%)	19 (6.4%)	2 (3.0%)	
	3	11 (1.5%)	4 (1.4%)	0 (0.0%)	
	4	21 (2.8%)	20 (6.8%)	0 (0.0%)	
Coronarography and primary PCI					
Left main stenosis ≥50%	Yes	17 (50.0%)	16 (47.1%)	1 (2.9%)	0.036
Left main stenosis as culprit lesion	Yes	4 (36.4%)	7 (63.6%)	0 (0.0%)	0.035
LAD	Yes	332 (74.3%)	86 (19.2%)	29 (6.5%)	<0.001
RCA	Yes	288 (62.9%)	142 (31.0%)	28 (6.1%)	0.018

pPCI -primary PCI, LAD—left anterior descending artery, LCx—left circumflex artery, OM—obtuse marginal artery, RCA—right coronary artery. Fisher's exact test and Kruskal-Wallis test.

Endpoint Occurrence in Relation to Stent Type

Primary net-clinical endpoint (i.e., death, nonfatal MI, stroke, major bleeding, and revascularization)
(DES vs BMS; DES vs BVS)

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
7 days								
Primary endpoint	19 (2.5%)	19 (6.3%)	2 (3.0%)	0.011	2.70 (1.42–5.15)	0.002	1.25 (0.29–5.39)	0.763
30 days								
CV death	12 (1.6%)	9 (3.0%)	1 (1.5%)	0.303	1.92 (0.80–4.55)	0.139	0.94 (0.12–7.23)	0.953
Re-MI	9 (1.2%)	3 (1.0%)	1 (1.5%)	0.791	0.85 (0.23–3.14)	0.808	1.26 (0.16–10.01)	0.822
Stroke	2 (0.3%)	1 (0.3%)	0 (0.0%)	0.999	1.27 (0.11–14.10)	0.841	–	–
CV death/Re-MI/Stroke	19 (2.5%)	13 (4.4%)	2 (3.0%)	0.281	1.75 (0.86–3.55)	0.119	1.20 (0.27–5.15)	0.807
Death	14 (1.9%)	12 (4.1%)	1 (1.5%)	0.101	2.20 (1.02–4.76)	0.045	0.81 (0.11–6.13)	0.835
Stent thrombosis	6 (0.8%)	2 (0.7%)	1 (1.5%)	0.587	0.84 (0.17–4.19)	0.838	1.89 (0.22–15.75)	0.553
Bleeding	40 (5.3%)	24 (8.1%)	3 (4.5%)	0.218	1.57 (0.94–2.61)	0.079	0.85 (0.26–2.77)	0.799
TIMI—severe	3 (0.4%)	4 (1.4%)	0 (0.0%)	0.232	3.43 (0.76–15.33)	0.106	–	–
BARC—severe	7 (0.9%)	6 (2.0%)	0 (0.0%)	0.346	2.21 (0.74–6.58)	0.154	–	–

Cox proportional risk model

Endpoint Occurrence in Relation to Stent Type

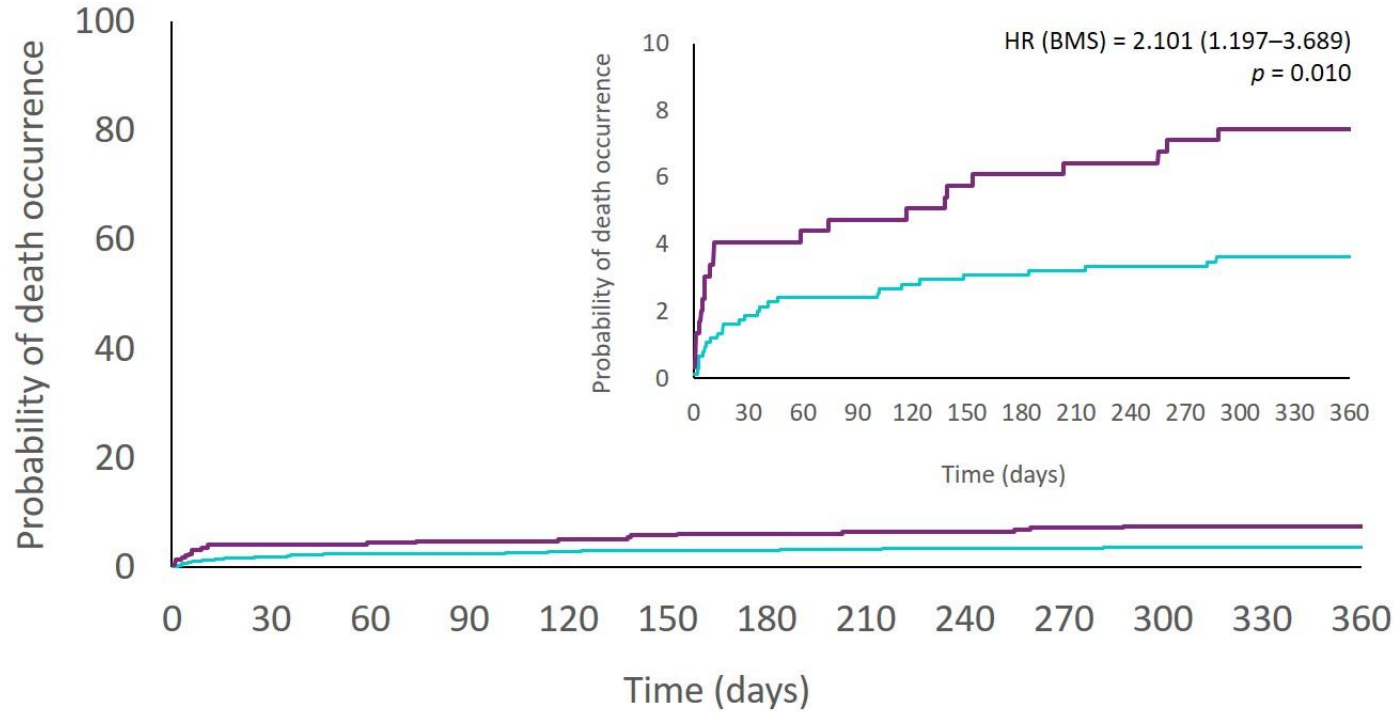
Primary net-clinical endpoint (i.e., death, nonfatal MI, stroke, major bleeding, and revascularization)
(DES vs BMS; DES vs BVS)

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
365 days								
CV death	20 (2.7%)	15 (5.1%)	1 (1.5%)	0.119	1.93 (0.98–3.76)	0.054	0.56 (0.07–4.18)	0.573
Re-MI	20 (2.7%)	8 (2.7%)	1 (1.5%)	0.999	1.03 (0.45–2.34)	0.935	0.56 (0.07–4.19)	0.575
Stroke	6 (0.8%)	3 (1.0%)	1 (1.5%)	0.523	1.29 (0.32–5.18)	0.713	1.85 (0.22–15.42)	0.566
CV death/Re-MI/Stroke	39 (5.2%)	25 (8.4%)	3 (4.5%)	0.150	1.66 (1.01–2.74)	0.047	0.86 (0.26–2.80)	0.810
Death	27 (3.6%)	22 (7.4%)	1 (1.5%)	0.018	2.10 (1.19–3.69)	0.010	0.41 (0.05–3.05)	0.388
Stent thrombosis	10 (1.3%)	3 (1.0%)	1 (1.5%)	0.812	0.77 (0.21–2.79)	0.690	1.13 (0.14–8.82)	0.907
Bleeding	78 (10.4%)	32 (10.8%)	10 (15.2%)	0.461	1.08 (0.71–1.62)	0.715	1.45 (0.75–2.80)	0.268
TIMI—severe	4 (0.5%)	4 (1.4%)	2 (3.0%)	0.051	2.58 (0.64–10.32)	0.180	5.63 (1.03–30.73)	0.046
BARC—severe	12 (1.6%)	6 (2.0%)	2 (3.0%)	0.453	1.29 (0.48–3.44)	0.609	1.87 (0.41–8.36)	0.412

Cox proportional risk model

Secondary Endpoint Occurrence in Relation to Stent Type

Secondary clinical endpoint (death rate)



		30	180	365
DES	Number of patients at risk	734	721	634
	Probability of death (95% CI)	1.9 (0.9; 2.8)	3.1 (1.8; 4.3)	3.6 (2.3; 5.0)
BMS	Number of patients at risk	284	277	245
	Probability of death (95% CI)	4.1 (1.8; 6.3)	6.1 (3.4; 8.8)	7.4 (4.5; 10.4)

Kaplan-Meier curves of cumulative incidence of death during 365 days in DES and BMS group.

Endpoint Occurrence in Relation to Stent Type in Patients Treated with Prasugrel vs. Ticagrelor

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
Patients Randomized to Prasugrel								
7 days								
Primary Endpoint	10 (2.6%)	9 (6.3%)	2 (4.7%)	0.104	2.74 (1.09–6.92)	0.032	1.98 (0.42–9.19)	0.380
30 days								
CV death	6 (1.6%)	5 (3.5%)	0 (0.0%)	0.280	2.30 (0.70–7.55)	0.167	–	–
Re-MI	5 (1.3%)	1 (0.7%)	1 (2.3%)	0.649	0.54 (0.06–4.68)	0.583	1.81(0.21–15.55)	0.586
Stroke	2 (0.5%)	1 (0.7%)	0 (0.0%)	0.999	1.38 (0.12–15.22)	0.792	–	–
CV death/Re-MI/Stroke	11 (2.8%)	7 (4.9%)	1 (2.3%)	0.427	1.75 (0.67–4.51)	0.246	0.82 (0.10–6.39)	0.854
Death	7 (1.8%)	6 (4.2%)	0 (0.0%)	0.203	2.37 (0.79–7.07)	0.120	–	–
In stent thrombosis	2 (0.5%)	1 (0.7%)	1 (2.3%)	0.314	1.36 (0.12–15.08)	0.798	4.53(0.41–50.05)	0.217
Bleeding	23 (5.9%)	10 (7.0%)	3 (7.0%)	0.810	1.22 (0.58–2.56)	0.597	1.20 (0.36–4.00)	0.763
TIMI—severe	2 (0.5%)	2 (1.4%)	0 (0.0%)	0.483	2.77 (0.39–19.73)	0.307	–	–
BARC—severe	5 (1.3%)	2 (1.4%)	0 (0.0%)	0.999	1.11 (0.21–5.73)	0.898	–	–

Fisher's exact test and Cox proportional risk model

Endpoint Occurrence in Relation to Stent Type in Patients Treated with **Prasugrel vs. Ticagrelor**

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
Patients Randomized to Prasugrel								
365 days (biased by high switch rate to clopidogrel)								
CV death	11 (2.8%)	9 (6.3%)	0 (0.0%)	0.081	2.28 (0.94–5.51)	0.066	–	–
Re-MI	12 (3.1%)	3 (2.1%)	1 (2.3%)	0.913	0.69 (0.19–2.46)	0.575	0.74 (0.09–5.70)	0.774
Stroke	4 (1.0%)	2 (1.4%)	1 (2.3%)	0.425	1.40 (0.25–7.67)	0.694	2.19 (0.24–19.59)	0.483
CV death/Re-MI/Stroke	23 (5.9%)	13 (9.2%)	2 (4.7%)	0.398	1.58 (0.80–3.12)	0.186	0.77 (0.18–3.28)	0.728
Death	15 (3.9%)	13 (9.2%)	0 (0.0%)	0.018	2.42 (1.15–5.09)	0.019	–	–
In stent thrombosis	4 (1.0%)	2 (1.4%)	1 (2.3%)	0.425	1.39 (0.25–7.63)	0.699	2.23 (0.25–20.02)	0.471
Bleeding	40 (10.3%)	12 (8.5%)	9 (20.9%)	0.075	0.84 (0.44–1.61)	0.611	2.069 (1.00–4.26)	0.049
TIMI—severe	2 (0.5%)	2 (1.4%)	2 (4.7%)	0.035	2.80 (0.39–19.88)	0.303	8.90 (1.25–63.18)	0.029
BARC—severe	7 (1.8%)	2 (1.4%)	2 (4.7%)	0.325	0.79 (0.16–3.83)	0.777	2.52 (0.52–12.15)	0.248

Fisher’s exact test and Cox proportional risk model

Endpoint Occurrence in Relation to Stent Type in Patients Treated with **Prasugrel vs. Ticagrelor**

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
Patients Randomized to Ticagrelor								
7 days								
Primary Endpoint	9 (2.5%)	10 (6.6%)	0 (0.0%)	0.080	2.65 (1.07–6.52)	0.034	–	–
30 days								
CV death	6 (1.7%)	4 (2.6%)	1 (4.3%)	0.343	1.58 (0.44–5.60)	0.478	2.61 (0.31–21.68)	0.374
Re-MI	4 (1.1%)	2 (1.3%)	0 (0.0%)	0.999	1.19 (0.21–6.50)	0.839	–	–
Stroke	0 (0.0%)	0 (0.0%)	0 (0.0%)	–	–	–	–	–
CV death/Re-MI/Stroke	8 (2.2%)	6 (3.9%)	1 (4.3%)	0.345	1.78 (0.62–5.15)	0.282	1.95 (0.24–15.66)	0.526
Death	7 (1.9%)	6 (3.9%)	1 (4.3%)	0.265	2.04 (0.68–6.07)	0.199	2.23 (0.27–18.19)	0.451
In stent thrombosis	4 (1.1%)	1 (0.6%)	0 (0.0%)	0.999	0.58 (0.06–5.26)	0.636	–	–
Bleeding	17 (4.7%)	14 (9.1%)	0 (0.0%)	0.090	2.01 (0.99–4.09)	0.052	–	–
TIMI—severe	1 (0.3%)	2 (1.3%)	0 (0.0%)	0.310	4.76 (0.43–52.56)	0.202	–	–
BARC—severe	2 (0.6%)	4 (2.6%)	0 (0.0%)	0.144	4.77 (0.87–26.08)	0.071	–	–

Fisher's exact test and Cox proportional risk model

Endpoint Occurrence in Relation to Stent Type in Patients Treated with **Prasugrel vs. Ticagrelor**

	Stent			<i>p</i> -Value	BMS *		BVS *	
	DES	BMS	BVS		HR (95% CI)	<i>p</i>	HR (95% CI)	<i>p</i>
Patients Randomized to Ticagrelor								
365 days (biased by high switch rate to clopidogrel)								
CV death	9 (2.5%)	6 (3.9%)	1 (4.3%)	0.420	1.58 (0.56–4.44)	0.384	1.74 (0.22–13.79)	0.596
Re-MI	8 (2.2%)	5 (3.2%)	0 (0.0%)	0.742	1.50 (0.49–4.59)	0.475	–	–
Stroke	2 (0.6%)	1 (0.6%)	0 (0.0%)	0.999	1.19 (0.10–13.16)	0.885	–	–
CV death/Re-MI/Stroke	16 (4.4%)	12 (7.8%)	1 (4.3%)	0.294	1.80 (0.85–3.80)	0.124	0.98 (0.13–7.38)	0.984
Death	12 (3.3%)	9 (5.8%)	1 (4.3%)	0.315	1.78 (0.75–4.24)	0.188	1.31 (0.17–10.09)	0.794
In stent thrombosis	6 (1.7%)	1 (0.6%)	0 (0.0%)	0.765	0.39 (0.04–3.27)	0.388	–	–
Bleeding	38 (10.5%)	20 (13.0%)	1 (4.3%)	0.496	1.29 (0.75–2.22)	0.351	0.39 (0.05–2.89)	0.363
TIMI—severe	2 (0.6%)	2 (1.3%)	0 (0.0%)	0.653	2.38 (0.33–16.91)	0.385	–	–
BARC—severe	5 (1.4%)	4 (2.6%)	0 (0.0%)	0.638	1.91 (0.51–7.12)	0.333	–	–

Fisher's exact test and Cox proportional risk model

Conclusions

- Patients with the highest initial risk profile were preferably treated with BMS over BVS.
- BMS were associated with a significantly higher rate of cardiovascular events whether treated with prasugrel or ticagrelor.

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Article

Stent Selection for Primary Angioplasty and Outcomes in the Era of Potent Antiplatelets. Data from the Multicenter Randomized Prague-18 Trial

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