# **Supplementary Online Content**

Giustino G, Harari R, Baber U, et al. Long-term safety and efficacy of new-generation drug-eluting stents in women with acute myocardial infarction: from the Women in Innovation and Drug-Eluting Stents (WIN-DES) Collaboration. *JAMA Cardiol*. Published online June 28, 2017. doi:10.1001/jamacardio.2017.1978

eTable 1. Characteristics of Included Randomized Controlled Trials.

eTable 2. Clinical Endpoint Definitions Used Across Randomized Controlled Trials.

eTable 3. Baseline Clinical and Angiographic Characteristics.

#### eReferences

**eFigure.** Kaplan-Meier curves for death, myocardial infarction or target lesion revascularization (1A), death, myocardial infarction or stent thrombosis (1B), definite or probable stent thrombosis (1C) and target lesion revascularization (1D) in women presenting with unstable angina, non-ST-elevation myocardial infarction or ST-elevation myocardial infarction.

This supplementary material has been provided by the authors to give readers additional information about their work.

Study	Year	Patients	Women	Stents used	Key inclusion criteria	Recommended DAPT Duration
RAVEL <sup>1</sup>	2002	238	58 (24)	Cypher, BMS	Stable CAD or UA, single de-novo lesion	2 months
SIRIUS <sup>2</sup>	2003	1058	305 (29%)	Cypher, BMS	Stable CAD or UA, single de-novo lesion	3 months
E-SIRIUS <sup>3</sup>	2003	352	103 (29%)	Cypher, BMS	Stable CAD or UA, single de-novo lesion	2 months
C-SIRIUS⁴	2004	100	31 (31%)	Cypher, BMS	Stable CAD or UA, single de-novo lesion	3 months
TAXUS I⁵	2003	61	7 (11%)	Taxus, BMS	Stable CAD or UA, single lesion	6 months
TAXUS II SR <sup>6</sup>	2003	267	67 (25%)	Taxus, BMS	Stable CAD or UA, single de-novo lesion	6 months
TAXUS IV <sup>7</sup>	2004	1314	367 (28%)	Taxus, BMS	Stable CAD or UA, single de-novo lesion	6 months
TAXUS V <sup>8</sup>	2005	1156	353 (31%)	Taxus, BMS	Stable CAD or UA, single de-novo lesion	6 months
SIRTAX <sup>9</sup>	2005	1012	231 (23%)	Cypher, Taxus	Stable CAD or UA, single de-novo lesion	12 months
ENDEAVOR II <sup>10</sup>	2006	1197	283 (24%)	Endeavor, BMS	Stable CAD or UA, single de-novo lesion	3 months
ENDEAVOR III <sup>11</sup>	2006	436	133 (31%)	Endeavor, Cypher	Stable CAD or UA, single	3 months

eTable 1. Characteristics of Included Randomized Controlled Trials.

					de-novo lesion	
ENDEAVOR IV <sup>12</sup>	2010	1548	500 (32%)	Endeavor, Taxus	Stable CAD or UA, single de-novo lesion	6 months
PROTECT <sup>13</sup>	2012	8709	2061 (24%)	Endeavor, Cypher	Stable CAD or UA, single de-novo lesion	12 months
RESOLUTE AC <sup>14</sup>	2010	2292	529 (23%)	Resolute, Xience	Stable CAD, UA, NSTEMI or STEMI	6 months
TWENTE <sup>15</sup>	2012	1391	382 (27%)	Resolute, Xience	Stable CAD, UA or NSTEMI	12 months
SPIRIT II <sup>16</sup>	2006	300	80 (27%)	Xience, Taxus	Stable CAD, UA or 2 de- novo lesions	6 months
SPIRIT III <sup>17</sup>	2008	1002	314 (31)	Xience, Taxus	Stable CAD, UA or 2 de- novo lesions	6 months
SPIRIT IV <sup>18</sup>	2010	3687	1189 (32)	Xience, Taxus	Stable CAD, UA or 3 de- novo lesions	12 months
COMPARE I <sup>19</sup>	2010	1800	526 (29%)	Xience, Taxus	Stable CAD, UA, NSTEMI or STEMI	12 months
BASKET-PROVE <sup>20</sup>	2010	2314	565 (24%)	Xience, Cypher, BMS	Stable CAD, UA or acute MI, target vessel diameter ≥ 3.0 mm	12 months
EXCELLENT <sup>21</sup>	2011	1443	512 (35%)	Xience, Promus, Cypher	Stable CAD, UA, NSTEMI	6 or 12 months*
RESET <sup>22</sup>	2012	3197	742 (23%)	Xience, Cypher	Stable CAD, UA, NSTEMI or STEMI	3 or 12 months*
PRODIGY <sup>23</sup>	2012	2013	473 (23%)	Xience, Promus,	Stable CAD, UA, NSTEMI	6 or 24 months*

				Endeavor, Taxus, BMS	or STEMI	
LEADERS <sup>24</sup>	2008 1707 430		430 (25%)	Riomatrix Cyphor	Stable CAD, UA, NSTEMI	12 months
LEADENS	2000	1707	430 (2370)	Biomatrix, Cypher	or STEMI	12 11011115
COMPARE II <sup>25</sup>	2013 2707 293	293 (26%)	Nobori, Xience, Promus	Stable CAD, UA, NSTEMI	12 months	
	2013	2013 2101 29	293 (2070)	Noboli, Alence, I Tollida	or STEMI	12 11011115
ISAR-TEST 4 <sup>26</sup> 2	2009 2603 623 (24%)	Yukon, Xience, Cypher	Stable CAD, UA, NSTEMI	6 months		
		rakon, Alence, Cypher	or STEMI	0 months		

CAD: Coronary Artery Disease; BMS: Bare Metal Stent; NSTEMI: Non-ST segment Elevation Myocardial Infarction; STEMI: ST segment Elevation Myocardial Infarction; UA: Unstable Angina. Cypher and Cordis, Johnson & Johnson, Miami Lakes, FL, USA; Taxus, Boston Scientific, Natick, MA, USA; Xience, Abbott Vascular, Santa Clara, CA, USA; Promus, Boston Scientific; Endeavor, Medtronic, Santa Rosa, CA, USA; Resolute, Medtronic; Biomatrix, Biosensors, Newport Beach, CA, USA; Nobori, Terumo, Tokyo, Japan; Yukon, Translumina, Hechingen, Germany. \*Patients were randomized to different DAPT durations.

Trial name	Myocardial infarction definition	Myocardial	Target lesion revascularization	Stent thrombosis
		infarction definition	definition	definition
		(old or new)		
RAVEL	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent.	
SIRIUS	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent.	
E-SIRIUS	Development of Q waves in $\geq 2$		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent.	
C-SIRIUS	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent.	

eTable 2. Clinical Endpoint Definitions Used Across Randomized Controlled Trials.

TAXUS I	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with CK and CK-MB		a stenosis of the luminal diameter	
	levels elevated above normal	Old	anywhere within the stent or within	
			the 5-mm borders proximal or	
			distal to the stent.	
TAXUS II SR	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of		a stenosis of the luminal diameter	
	Q waves, increase in the CK level	Old	anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	
			distal to the stent.	
TAXUS IV	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of		a stenosis of the luminal diameter	
	Q waves, increase in the CK level	Old	anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	
			distal to the stent.	
TAXUS V	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of		a stenosis of the luminal diameter	
	Q waves, increase in the CK level	Old	anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	
			distal to the stent	
SIRTAX	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of	Old	a stenosis of the luminal diameter	
	Q waves, increase in the CK level	Olu	anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	

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	or troponin I		distal to the stent	
ENDEAVOR II	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of		a stenosis of the luminal diameter	
	Q waves, increase in the CK level	Old	anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	
			distal to the stent	
ENDEAVOR III	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent	
ENDEAVOR IV	Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads with elevated cardiac		a stenosis of the luminal diameter	
	enzymes or, in the absence of Q	Old	anywhere within the stent or within	
	waves, increase in the CK level		the 5-mm borders proximal or	
	≥2*ULN and increased level of CK-MB		distal to the stent	
				ARC criteria
PROTECT	II Universal Definition (Thygesen K et		Revascularization for ischemia for	ARC criteria
	al. Circulation 2007): Periprocedural		a stenosis of the luminal diameter	
	MI: cardiac biomarkers increase		anywhere within the stent or within	
	≥3*ULN Spontaneous: Typical rise and	New	the 5-mm borders proximal or	
	fall of cardiac biomarkers (preferably		distal to the stent	
	troponin) with at least 1 value >URL			
	and at least 1 of the following:			

symptoms, ST-T changes at ECG, pathological Q waves, or imaging evidence of ischemia **RESOLUTE AC** Extended historical definition (Vranckx et al. Eurointervention 2010). In summary: development of Q waves in ≥2 contiguous leads and elevated cardiac enzymes or, in the absence of Q waves, increase in the CK level ≥2\*ULN and increased level of CK-MB or troponin. In patients with acute MI at baseline: if cardiac biomarkers still raising new chest pain of ischemia equivalent and rise in cardiac biomarkers >50% previous level; if cardiac biomarkers have returned to normal, CK level ≥2\*ULN. TWENTE Extended historical definition (Vranckx et al. Eurointervention 2010). In summary: development of Q waves in ≥2 contiguous leads and elevated cardiac enzymes or, in the absence of Q waves, increase in the CK level ≥2\*ULN and increased level of CK-MB

Revascularization for ischemia for ARC criteria a stenosis of the luminal diameter anywhere within the stent or within the 5-mm borders proximal or distal to the stent

New

New

Revascularization for ischemia for<br/>a stenosis of the luminal diameter<br/>anywhere within the stent or within<br/>the 5-mm borders proximal or<br/>distal to the stentARC criteria

SPIRIT II	or troponin. In patients with acute MI at baseline: if cardiac biomarkers still raising new chest pain of ischemia equivalent and rise in cardiac biomarkers >50% previous level; if cardiac biomarkers have returned to normal, CK level ≥2*ULN. Development of Q waves in ≥2		Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of Q waves, a typical rise and fall of CK- MB (if non-procedural/spontaneous MI, CK-MB >2 times upper limit of normal; if post PCI, CK-MB >3 times upper limit of normal; if post CABG, CK-MB >5 times upper limit of normal)	New	a stenosis of the luminal diameter anywhere within the stent or within the 5-mm borders proximal or distal to the stent	
SPIRIT III	Development of Q waves in ≥2 contiguous leads with elevated cardiac enzymes or, in the absence of Q waves, increase in the CK level ≥2*ULN and increased level of CK-MB	New	Revascularization for ischemia for a stenosis of the luminal diameter anywhere within the stent or within the 5-mm borders proximal or distal to the stent	ARC criteria
SPIRIT IV	Development of Q waves in ≥2 contiguous leads with elevated cardiac enzymes or, in the absence of Q waves, increase in the CK level	New	Revascularization for ischemia for a stenosis of the luminal diameter anywhere within the stent or within the 5-mm borders proximal or	ARC criteria

	≥2*ULN and increased level of CK-MB		distal to the stent	
COMPARE	Periprocedural MI (in patients without		Revascularization for ischemia for	ARC criteria
	acute MI at baseline): any elevation in		a stenosis of the luminal diameter	
	concentrations of CK ≥2*ULN and		anywhere within the stent or within	
	increase in CK-MB or troponin.		the 5-mm borders proximal or	
	Spontaneous MI: typical rise and fall of	Nous	distal to the stent	
	troponin or CK-MB with at least one of	New		
	the following: ischemic symptoms,			
	development of pathological Q waves,			
	ischemic ECG changes, or			
	pathological findings of an acute MI			
BASKET-	Typical rise and fall of cardiac		Target vessel Revascularization	ARC criteria
PROVE	biomarkers (preferably troponin) with at		was used	
	least 1 value >URL and at least 1 of	Nous		
	the following: symptoms, ST-T	New		
	changes at ECG, pathological Q			
	waves, or recent angioplasty.			
EXCELLENT	Academic Research Consortium		Revascularization for ischemia for	ARC criteria
	criteria (Cutlip DE et al. Circulation		a stenosis of the luminal diameter	
	2007) In summary:		anywhere within the stent or within	
	Periprocedural MI: troponin >3*URL or	New	the 5-mm borders proximal or	
	CK-MB>3*URL if baseline cardiac		distal to the stent	
	biomarkers <url. or<="" stable="" td=""><td></td><td></td><td></td></url.>			
	decreasing values on 2 samples			

	followed by 20% increase if baseline			
	cardiac biomarkers >URL.			
	Spontaneous MI: troponin >URL or			
	CK-MB >URL			
RESET	Periprocedural MI: CK-MB ≥3*ULN or		Revascularisation for ischemia for	ARC criteria
	CK ≥3*ULN in the absence of CKMB		a stenosis of the luminal diameter	
	measurement.		anywhere within the stent or within	
	Spontaneous MI: Academic Research	New	the 5-mm borders proximal or	
	Consortium criteria (Cutlip DE et al.		distal to the stent	
	Circulation 2007), troponin >URL or			
	CK-MB >URL			
		New		ARC criteria
PRODIGY	II Universal Definition (Thygesen K et		Target vessel Revascularisation	ARC criteria
	al. Circulation 2007): Periprocedural		was used	
	MI: cardiac biomarkers increase			
	≥3*ULN Spontaneous: Typical rise and			
	fall of cardiac biomarkers (preferably	New		
	troponin) with at least 1 value >URL	New		
	and at least 1 of the following:			
	symptoms, ST-T changes at ECG,			
	pathological Q waves, or imaging			
	evidence of ischemia			
LEADERS	Development of Q waves in ≥2	New	Revascularization for ischemia for	ARC criteria
	contiguous leads or, in the absence of		a stenosis of the luminal diameter	

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	Q waves, increase in the CK level		anywhere within the stent or within	
	≥2*ULN and increased level of CK-MB		the 5-mm borders proximal or	
	or troponin I		distal to the stent	
COMPARE-2	Periprocedural MI (in patients without		Revascularization for ischemia for	ARC criteria
	acute MI at baseline):any elevation in		a stenosis of the luminal diameter	
	concentrations of CK ≥2*ULN and		anywhere within the stent or within	
	increase in CK-MB or troponin.		the 5-mm borders proximal or	
	Spontaneous MI: typical rise and fall of	Nierre	distal to the stent	
	troponin or CK-MB with at least one of	New		
	the following: ischemic symptoms,			
	development of pathological Q waves,			
	ischemic ECG changes, or			
	pathological findings of an acute MI			
ISAR-TEST 4	Periprocedural MI: CK-MB (or CK)		Revascularization for ischemia for	ARC criteria
	≥3*ULN and at least 50% over the		a stenosis of the luminal diameter	
	most recent pre-PCI levels, or the		anywhere within the stent or within	
	development of new ECG changes		the 5-mm borders proximal or	
	consistent with MI and CK-MB (CK)		distal to the stent	
	elevation >ULN at 2 measurements for	New		
	patients with stable angina pectoris or			
	NSTE-ACS and falling or normal CK-			
	MB (CK) levels. Recurrent chest pain			
	lasting .30 min with either new ECG			
	changes consistent with second MI or			

next CK-MB (CK) level at least 8–12 h after PCI elevated at least 50% above the previous level was considered procedure-related MI for patients presenting with elevated CK-MB (CK) level prior to PCI. Spontaneous MI: any CK-MB increase with or without the development of Q-waves on ECG.

ARC: Academic Research Consortium; CK: Creatine-Kinase; ECG = Electrocardiogram; MI: Myocardial Infarction; URL: Upper Reference Limit.

eTable 3. Baseline Clinical and Angiographic Characteristics.

	Overall	UA	NSTEMI	STEMI	P-value	
	(N=4373)	(N=2197)	(N=2197) (N=1397)		F-value	
Age (Years)	66.81 ± 11.28	67.01 ± 10.89	66.49 ± 11.27	66.82 ± 12.34	0.42	
BMI (kg/m²)	27.47 ± 5.45	27.81 ± 5.66	27.44 ± 5.28	26.47 ± 5.01	<0.0002	
Risk factors						
Diabetes mellitus	1229 (28.1%)	712 (32.4%)	377 (27.0%)	140 (18.0%)	<0.0002	
IDDM	386 (31.4%)	211 (29.6%)	139 (36.9%)	36 (25.7%)	0.02	
Hypertension	3067 (70.1%)	1695 (77.2%)	942 (67.4%)	430 (55.2%)	< 0.000	
Hypercholesterolemia	2638 (60.3%)	1519 (69.1%)	802 (57.4%)	317 (40.7%)	<0.000	
Serum creatinine (mg/dl)	0.91 ± 0.59	$0.90 \pm 0.53$	0.91 ± 0.63	$0.90 \pm 0.66$	0.92	
Smoking	1385 (31.7%)	565 (25.8%)	515 (36.9%)	305 (39.2%)	<0.000	
Family history of CAD	1517 (35.4%)	745 (35.0%)	510 (36.9%)	262 (34.2%)	0.37	
Clinical history						
Previous MI	807 (18.5%)	427 (19.5%)	311 (22.3%)	69 (8.9%)	< 0.000	
Previous PCI	693 (15.9%)	514 (23.4%)	127 (9.1%)	52 (6.7%)	< 0.000	

Previous CABG	196 (4.5%)	143 (6.5%)	43 (3.1%)	10 (1.3%)	<0.0001
Stent generation					0.32
Early-generation DES	1608 (36.8%)	832 (37.9%)	498 (35.6%)	278 (35.7%)	
New-generation DES	2765 (63.2%)	1365 (62.1%)	899 (64.4%)	501 (64.3%)	
Angiographic Characteristics					
Multivessel disease	1148 (30.5%)	591 (30.6%)	387 (30.4%)	170 (29.9%)	0.94
Number of lesions treated	1.33 ± 0.66	1.27 ± 0.59	1.45 ± 0.76	1.29 ± 0.62	<0.0001
Number of DES implanted	1.58 ± 0.97	1.49 ± 0.88	1.72 ± 1.09	1.58 ± 0.94	<0.0001
Mean stent diameter (mm)	$3.00 \pm 0.39$	$3.00 \pm 0.40$	2.96 ± 0.39	3.10 ± 0.38	<0.0001
Total stent length (mm)	30.96 ± 20.48	28.98 ± 18.24	33.90 ± 23.56	31.96 ± 20.47	<0.0001
LVEF (%)	52.45 ± 17.81	56.14 ± 16.84	47.58 ± 20.47	47.72 ± 11.58	<0.0001
Type B2/C lesion	2176 (65.2%)	1167 (62.7%)	674 (64.1%)	335 (78.8%)	<0.0001
Moderate/severe calcifications	661 (24.0%)	370 (23.0%)	256 (28.4%)	35 (14.4%)	<0.0001
At least 1 bifurcation lesion	231 (14.9%)	157 (18.2%)	50 (12.2%)	24 (8.6%)	<0.0001

Results reported as n (%) or mean ± standard deviation. BMI: Body Mass Index; DES: Drug-Eluting Stent; UA: Unstable Angina; NSTEMI: Non-ST segment Elevation Myocardial Infarction; STEMI: ST segment Elevation Myocardial Infarction;

CAD: Coronary Artery Disease; IDDM: Insulin-Dependent Diabetes Mellitus; MI: Myocardial Infarction; PCI: Percutaneous Coronary Intervention; CABG: Coronary Artery By-pass Graft.

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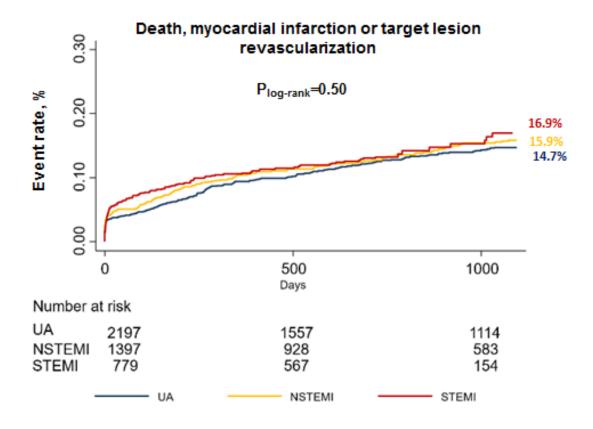
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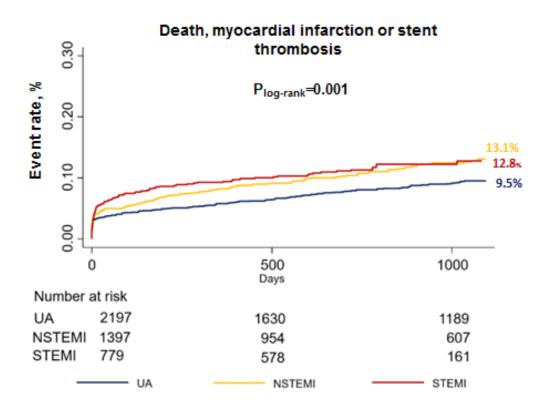
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**eFigure 1.** Kaplan-Meier curves for death, myocardial infarction or target lesion revascularization (2A), death, myocardial infarction or stent thrombosis (2B), definite or probable stent thrombosis (2C) and target lesion revascularization (2D) in women presenting with unstable angina, non-ST-elevation myocardial infarction or ST-elevation myocardial infarction. NSTEMI: Non-ST Elevation Myocardial Infarction; STEMI: ST Elevation Myocardial Infarction; UA: Unstable Angina.

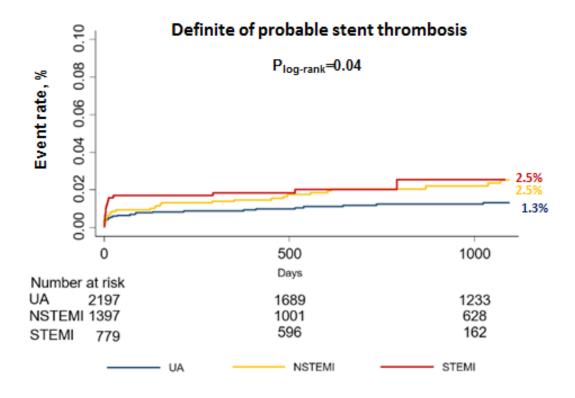
## eFigure 1A.



eFigure 1B.



## eFigure 1C.



## eFigure 1D.

