Bioresorbable Vascular Scaffolds for the Treatment of Chronic Total Occlusions; An International Multicenter Registry

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FACC, FAHA, FESC, FSCAI

Why BVS ???

About BVS

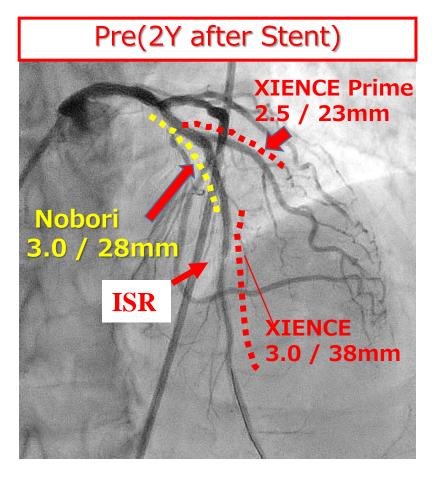


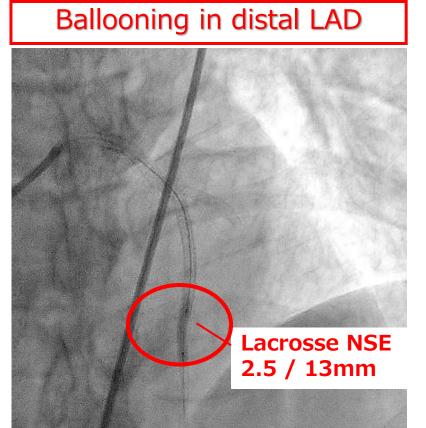
Long procedure time, Expensive, Need IVUS or OCT always..., A lot of requirement And higher incidence of thrombotic event

And all advantages is future advantage...

Case: Angina: 2Year after DES implantation

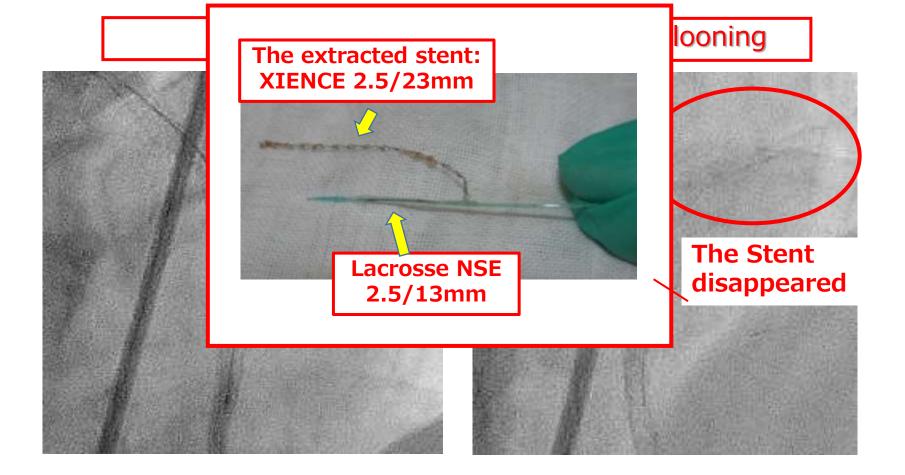
Patient had a late In-Stent restenosis in distal LAD





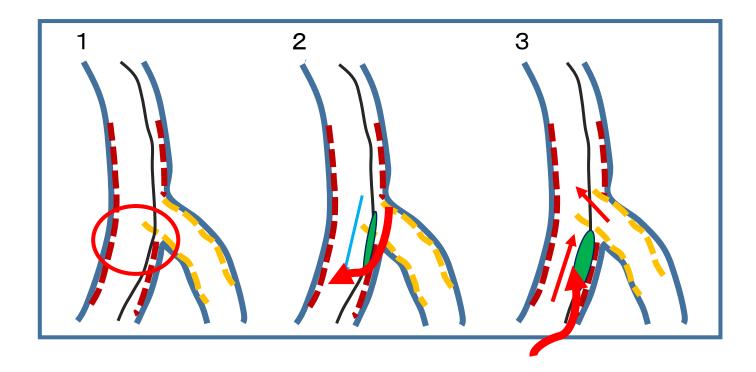
Stent Fell Off

Stent was Fell off by the deflated balloon

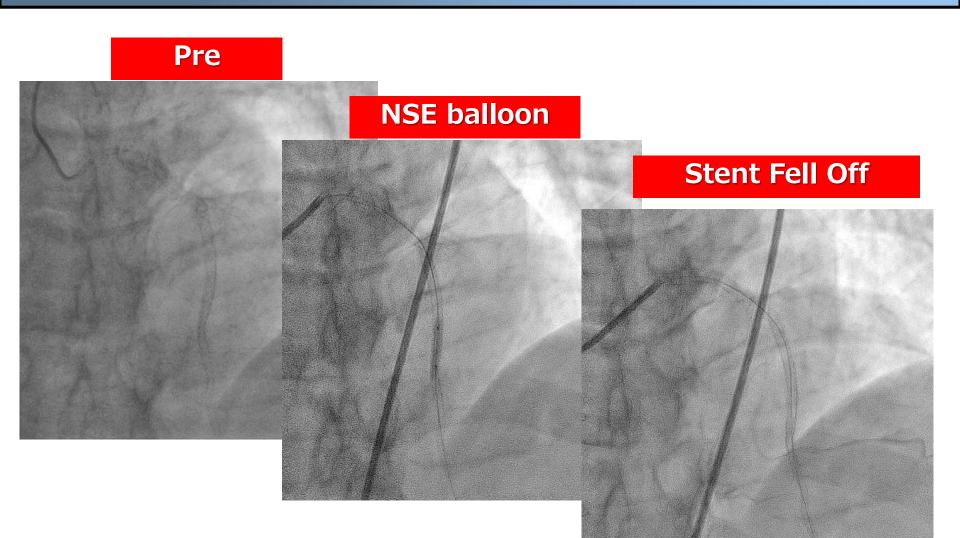


Mechanism of Falling Off

This rare complication was due to the entrapment of the deflated balloon in the protruded struts of the side branch stent to the main branch (neocarina). likely secondary to crossing the guidewire in the middle of these struts.



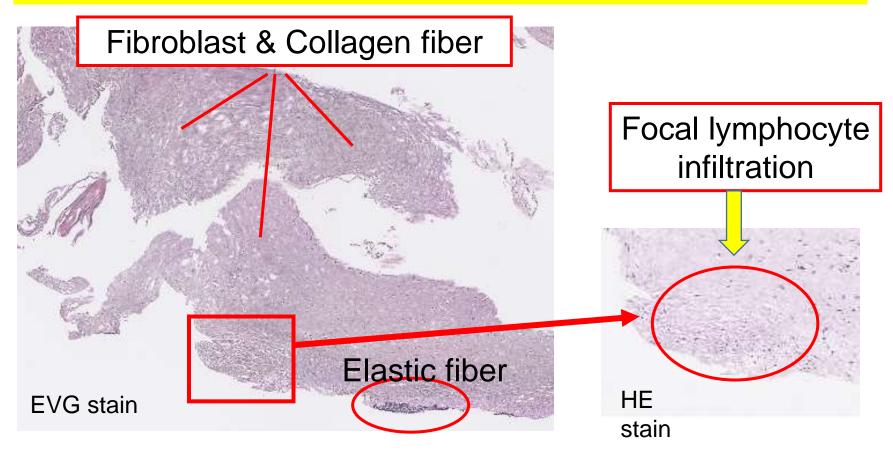
Stent Fell Off



Watch !! Diagonal

Microscopic Findings

Coronary artery specimen adhering to stent



The specimen consists mainly of fibrous thickening intima with a little part of media, and focal chronic inflammatory infiltration.

A potential procedural complication 2 years after percutaneous coronary intervention to treat left anterior descending artery lesions with the T-stenting and the small protrusion technique

Karube K, Naganuma T, Nakamura S et al JACC Cardiovasc Interv. 2015 March

Histological assessment of the tissue adhered to the extracted stent demonstrated fibrous thickening intima with a part of media and focal inflammation.

While difficult to prove, the connection of the vessel wall and stent could eventually be loose probably due to suppression of neo-intiaml proliferation by everolimus and/or chronic inflammation.

After DES implant...(even 2nd gen. Stent) We still have "Delayed Healing"...

DES is not PERFECT !!

But newly emerged BVS also has many challenges

Overcoming the limitations of metallic stents

The 2Y after BVS implant. from meta-analysis including 7 RCT

Absorb China	2 (0.8)	237	0 (0.0)	231			
Absorb II	5 (1.5)	325	0 (0-0)	163		1	
Absorb III	24 (1.9)	1272	5 (0.8)	660		<u> </u> ■	- 2.49 (0.95-6.50)
Absorb Japan	8 (3.1)	257	2 (1.5)	130			— 2.02 (0.44–9.39)
AIDA	31 (3.4)	924	8 (0.9)	921		- j 🖂 🖷	3.86 (1.79–8.36)
EVERBIO II	1(1.3)	77	0 (0.0)	80		1 24	
TROFIII	2 (2.1)	95	1(1.0)	96	5 <u>0</u>		2.02 (0.19-21.92)
Overall DL (l²=0%, p=0.8219)	73 (2.3)	3187	16 (0.7)	2281		٠	2.99 (1.73-5.15)
Overall M-H				1 <u>2</u>	Lis is	• •	3.35 (1.96-5.72)
Target ve				N <u>-</u>	L(† 3		
Overall M-H Target ve Absorb China	5 (2.1)	237	2 (0-8)	237	E? a		
Overall M-H Target ve Absorb China Absorb II	5 (2·1) 22 (6·7)	237 328	4 (2.4)	164	L)		
Overall M-H Target ve Absorb China Absorb II Absorb III	5 (2·1) 22 (6·7) 95 (7·3)	237 328 1296	4 (2·4) 32 (4·8)	164 673	L9 8		
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BVS was associated with increased rates of target vessel MI and definite/probable ScT at 2 years.

Absorb Japan Trial -2year result-

Two-year clinical, angiographic, and serial optical coherence tomographic follow-up after implantation of an everolimuseluting bioresorbable scaffold and an everolimus-eluting metallic stent: insights from the randomised ABSORB Japan trial



Yoshinobu Onuma¹, MD; Yohei Sotomi², MD; Hiroki Shiomi³, MD; Yukio Ozaki⁴, MD; Atsuro Namiki⁵, MD; Satoshi Yasuda⁶, MD; Takafumi Ueno⁷, MD; Kenji Ando⁸, MD; Jungo Furuya⁹, MD; Keiichi Igarashi⁹, MD; Ken Kozuma¹⁰, MD; Kengo Tanabe¹¹, MD;

The rate of TLF was numerically higher in the BVS arm than

in the CoCr-EES arm, although this difference was not

statistically significant.

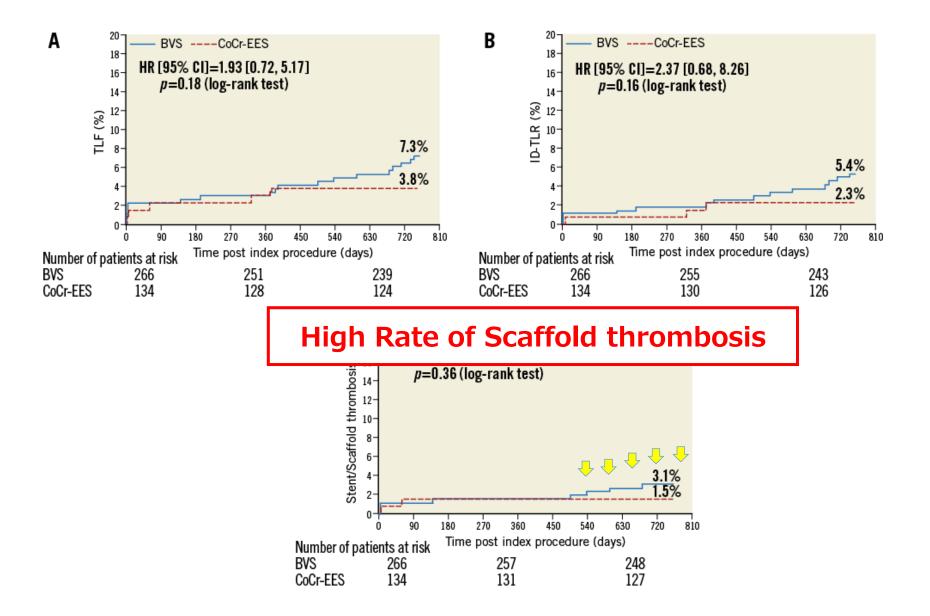
VLST was observed only in the BVS arm at a rate of 1.6%

between one and two years. Further studies are mandatory

to investigate the risk of BVS relative to metallic stents for VLST,

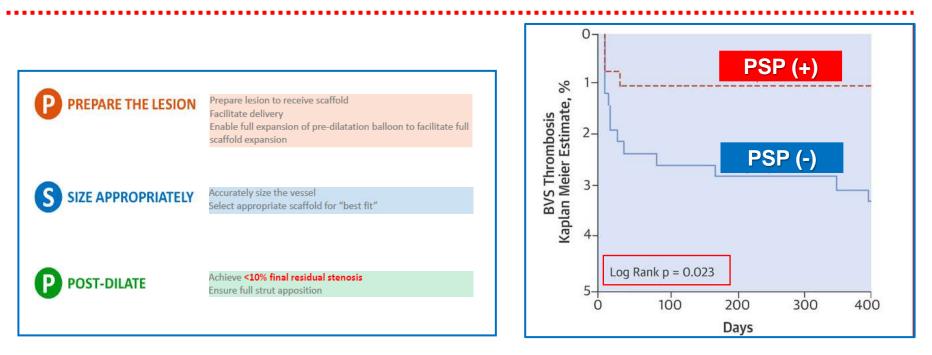
and the underlying mechanisms of BVS VLST.

Absorb Japan Trial -2year result-



What should be concerned?

1) The specific implantation technique is required



Pooled analysis of all patients with 2-year follow-up enrolled ABSORB trials (ABSORB II, III, Japan, and China)

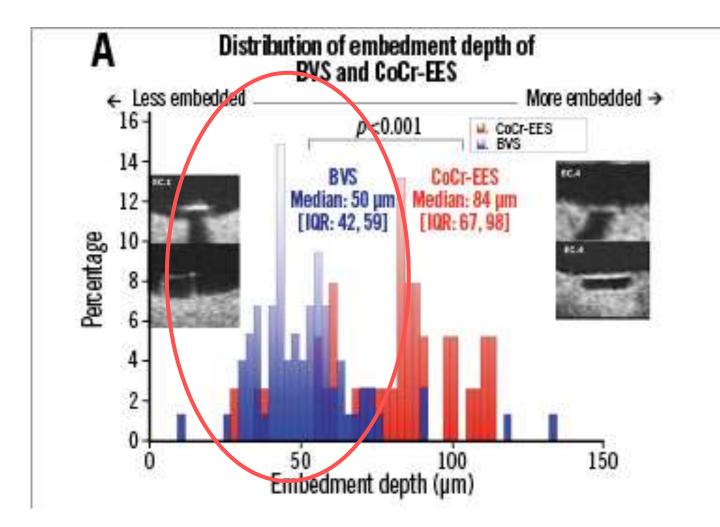
Scaffold thrombosis

226 patients with high-pressure post-dilatation (\geq 18atm): **0 case** 1773 patients without high-pressure post-dilatation (<18atm): **11 cases** (0.6%)

*Balloon size was larger than the nominal BVS diameter

Ali ZA, et al. Lancet. 2017 Aug 19;390(10096):760-772

Absorb Japan Trial -2year result-



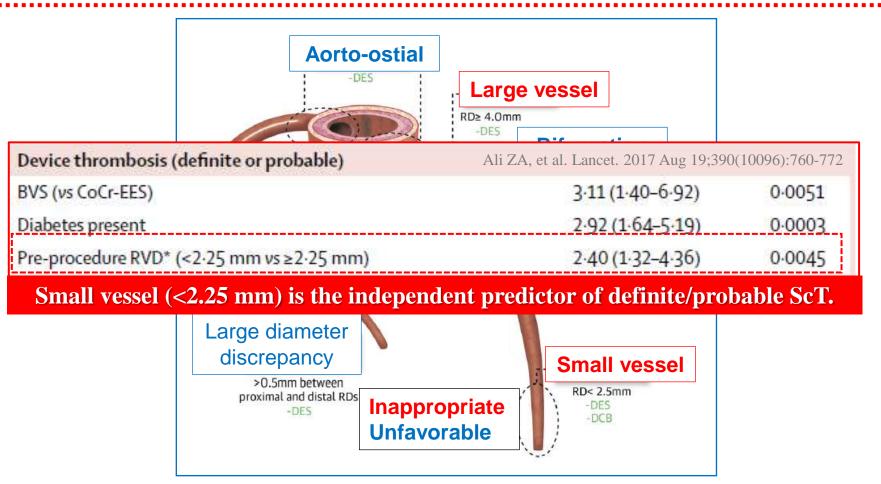
Absorb Japan Trial -2year result-

Online Table 3. Quantitative coronary angiographic results of OCT-1 subgroup (full analysis set).

	BVS	CoCr-EES	<i>p</i> -value	
Follow-up at 13 months				
Number of lesions	83	41		
Reference vessel diameter (mm)	2.68±0.4	2 70 0 50	0.01	
In-segment MLD (mm)	2.04±0.4	recoil		
In-device MLD (mm)	2.20±0.4	recon		
In-segment DS (%)	23.8±10.2	23.3±13.6	0.84	
In-device DS (%)	17.8±9.9			
In-segment binary restenosis	1 (1.2%	te lumer	1 IOS	
In device binary restenosis	1 (1.2%			
In-segment late lumen loss (mm)	0.17±0.29	0.06±0.33	0.072	
In-device late lumen loss (mm)	0.22±0.26	0.12±0.35	0.09	
Folion-up at 24 months				
Number of lesions	77	40		
Reference vessel diameter (mm)	2.68±0.43	2.77±0.50	0.35	
In-segment MLD (mm)	1.95±0.49	2.10±0.51	0.14	
In-device MLD (mm)	2.08±0.56	2.37±0.58	0.01	
In-segment DS (%)	27.7±12.2	24.7±10.2	0.16	
In-device DS (%)	23.3±13.0	14.8±11.9	0.0006	
In-segment binary restenosis	6 (7.8%)	1 (2.5%)	0.42	
In-device binary restenosis	4 (5.2%)	1 (2.5%)	0.66	
In-segment late lumen loss (mm)	0.27±0.38	0.12±0.32	0.029	
In-device late lumen loss (mm)	0.36±0.38	0.21±0.38	0.04	

What should be concerned?

② Strict lesion selection is required.



Tanaka A, Colombo A, et al. JACC Cardiovasc Interv. 2017 Mar 27;10(6):539-547.

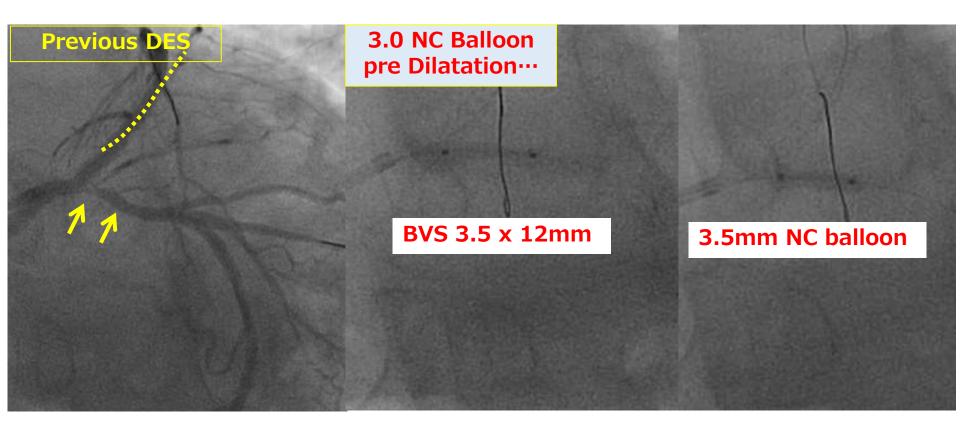
BVS is **not suitable for all types of lesions.** It should be particularly avoided to implant BVS for **small vessels (< 2.25mm).**

JACC: CARDIOVASCULAR IMAGING, VOL. 7, NO. 8, 2014

AUGUST 2014:843-50

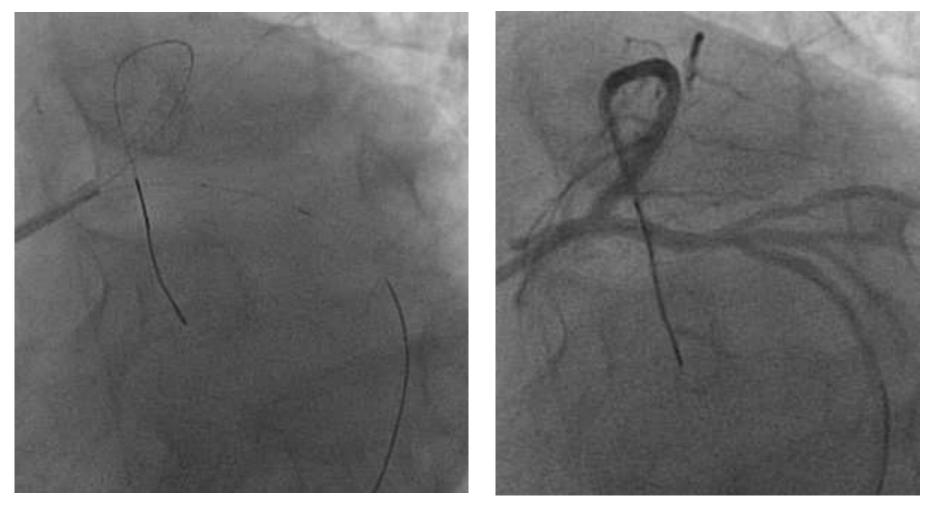
Toru Naganuma, MD Antonio Colombo, MD*

*EMO-GVM Centro Cuore Columbus



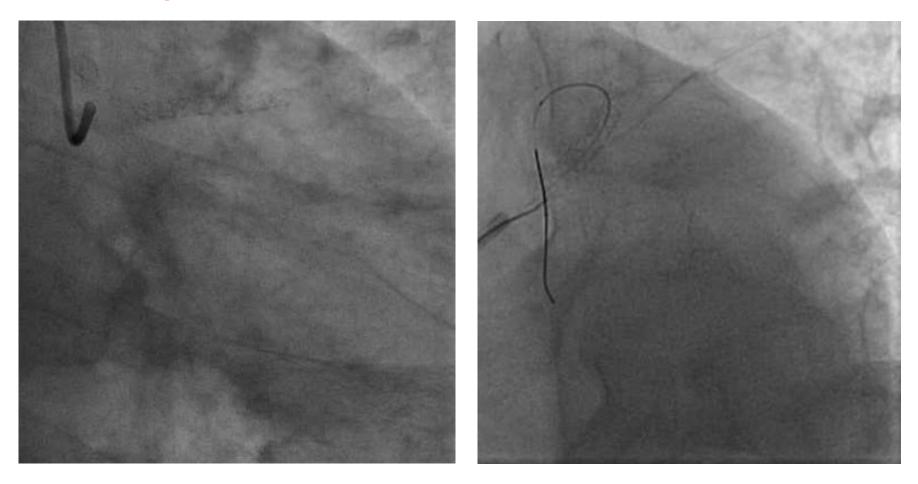
Toru Naganuma , Antonio Colombo at al JACC Int. 2015

Final Angiogram

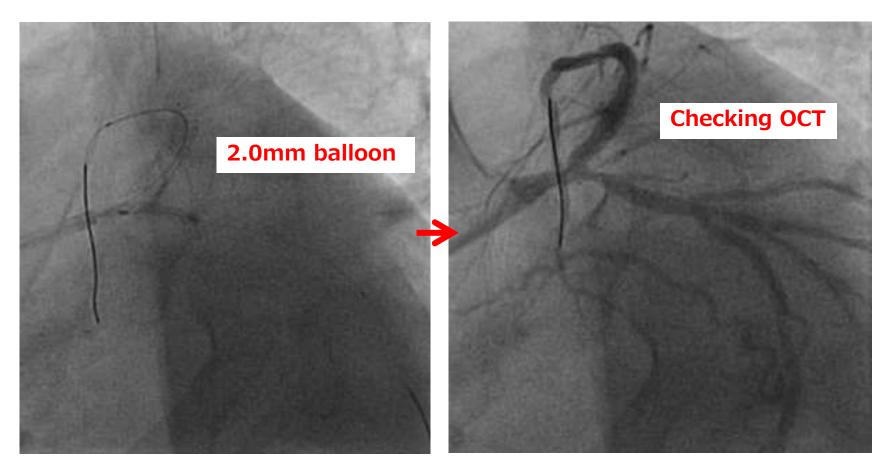


Toru Naganuma , Antonio Colombo at al JACC Int. 2015

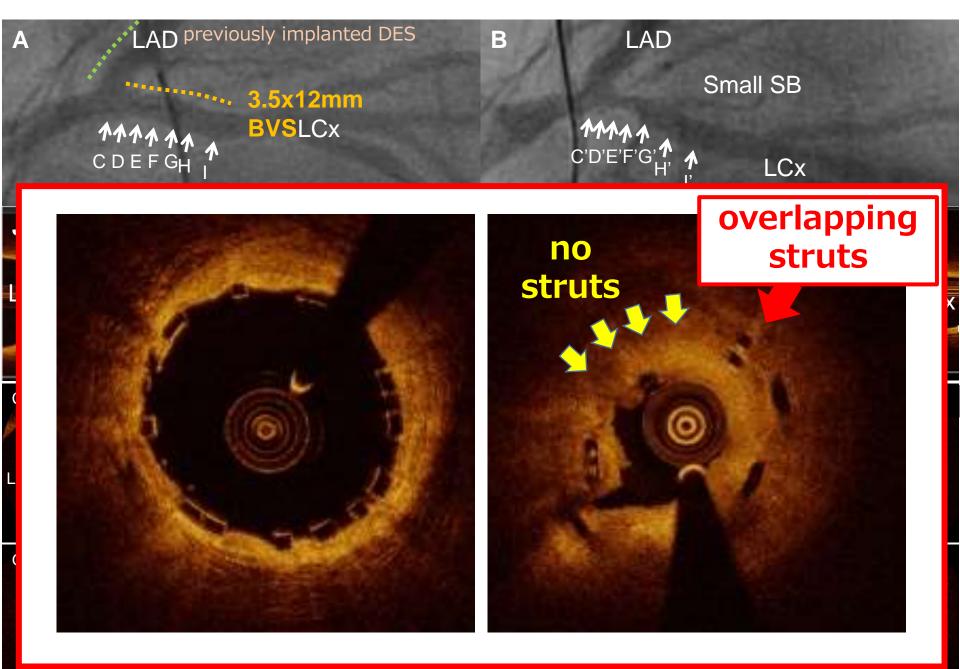
Only 6month later---Severe Restenosis in LCX ost.



Checking OCT at LCX ost …..



Toru Naganuma , Antonio Colombo at al JACC Int. 2015



Small SB

overlapping struts

Guide Post of BVS

- Important Reminder -

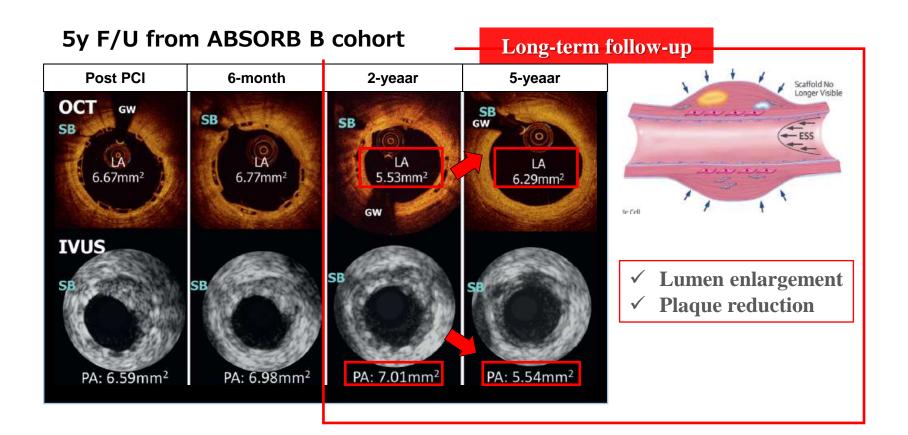
Implantation of BVS at ostium of LCX may be problematic.



Toru Naganuma M.D. FACC FESC

Catheter Cardiovasc Interv. 2013 JACC Imaging Vol 7 No.8 2014

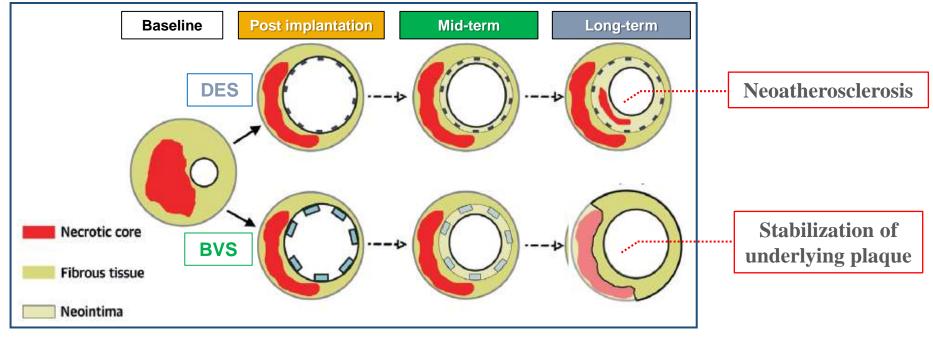
① BVS would contribute to positive vessel remodeling after complete bioresorption.



At long-term follow-up after BVS implant., positive vessel remodeling with plaque reduction has been reported, which could be favorable after CTO recanalization.

 BVS would contribute to less chronic inflammatory reaction after complete bioresorption.

5 Y OCT assessment from ABSORB A cohort

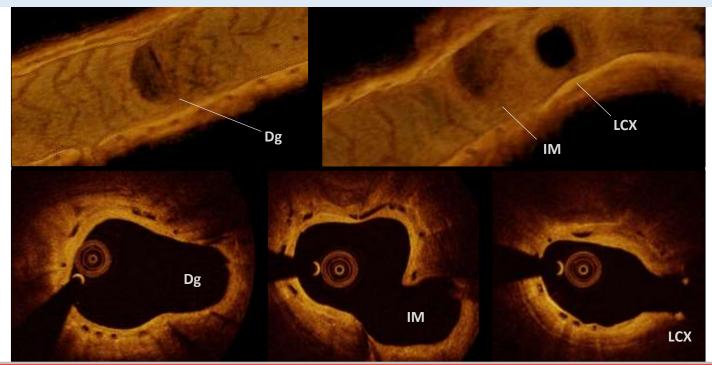


Karanasos A et al. J Am Coll Cardiol. 2014 Dec 9;64(22):2343-56

Homogeneous neointima after BVS implantation has been reported to contribute to seal underling high-risk plaque and additionally, develop less disease over time when compared to the neointima following metallic stents.

③ BVS has a potentially advantage in bifurcation area. 1

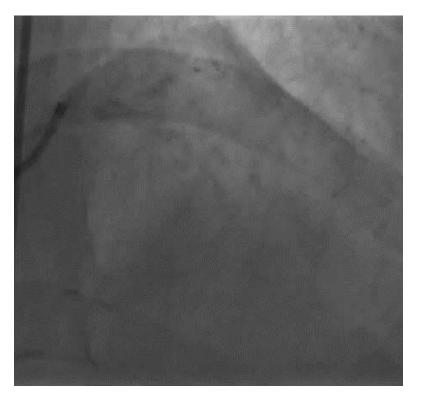
Case: BVS implant main branch without touching SB - No Final KISS ; 2Years Later OCT shows "beautiful Opening SB"-



If the side branch is not compromised, single cross-over stenting without any SB intervention could be considered.

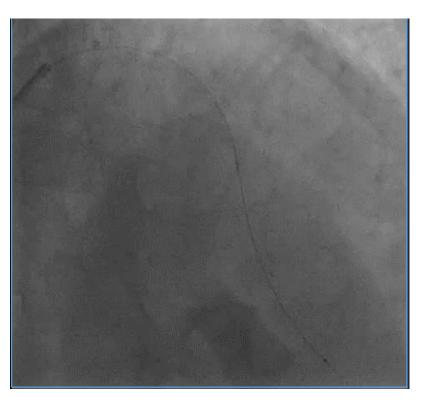
③ BVS has a potentially advantage in bifurcation area. 2

LAD/Dx simple disease



Moderate stenosis at Dx ostium

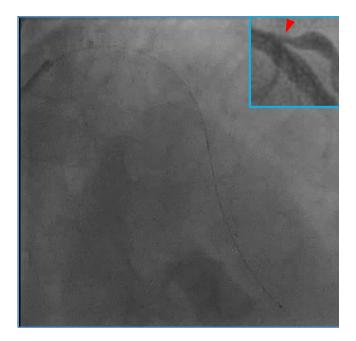
BVS implat. w/o KBT

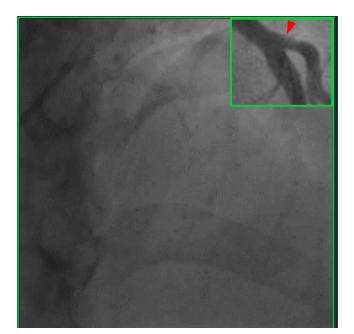


Improvement at ostium diagonal

③ BVS has a potentially advantage in bifurcation area. 2

Carina: going back to the original position





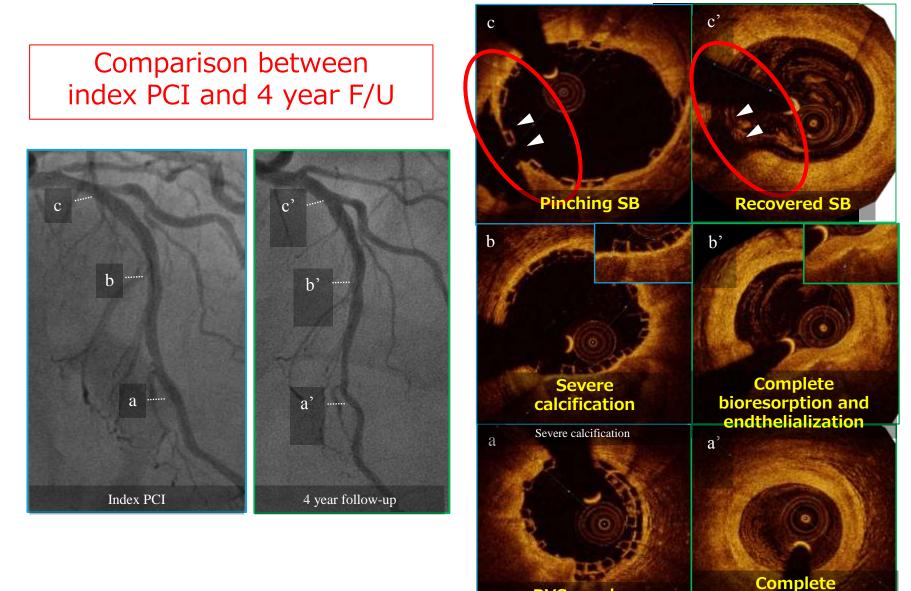
Just after index PCI

Pinching at Dx ostium

4 Y F/U

Improvement at Dx ostium

Carina was backed to the original position



BVS overlap bioresorption

At JIM ; I was a operator (Dr. Colombo' meeting in 2012)



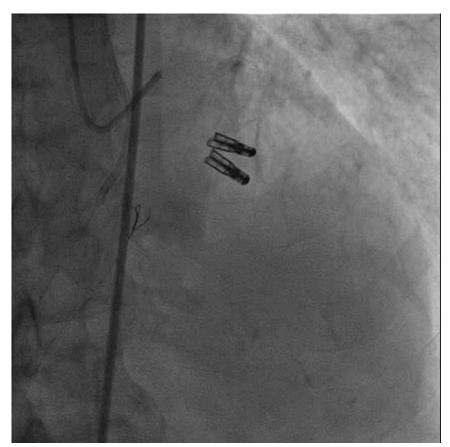
70's male LAD-CTO

LAD CTO

At Live Demo. In Milan 2012: I was a operator

70's male EF: 44% Normal renal function Post Mitra-Clip

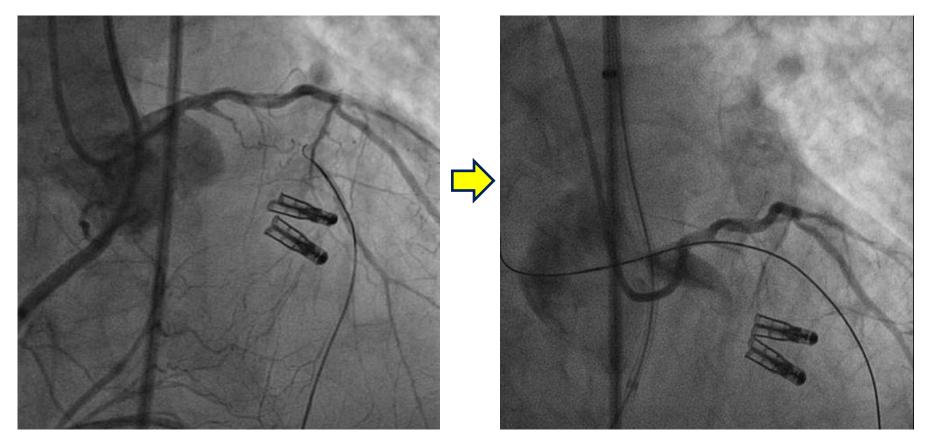
5th attempt LAD CTO from prox. to mid LAD



collaterals from 1) distal RCA (PD) via small septal branches 2) conus branch and 3) antegrade small bridge c

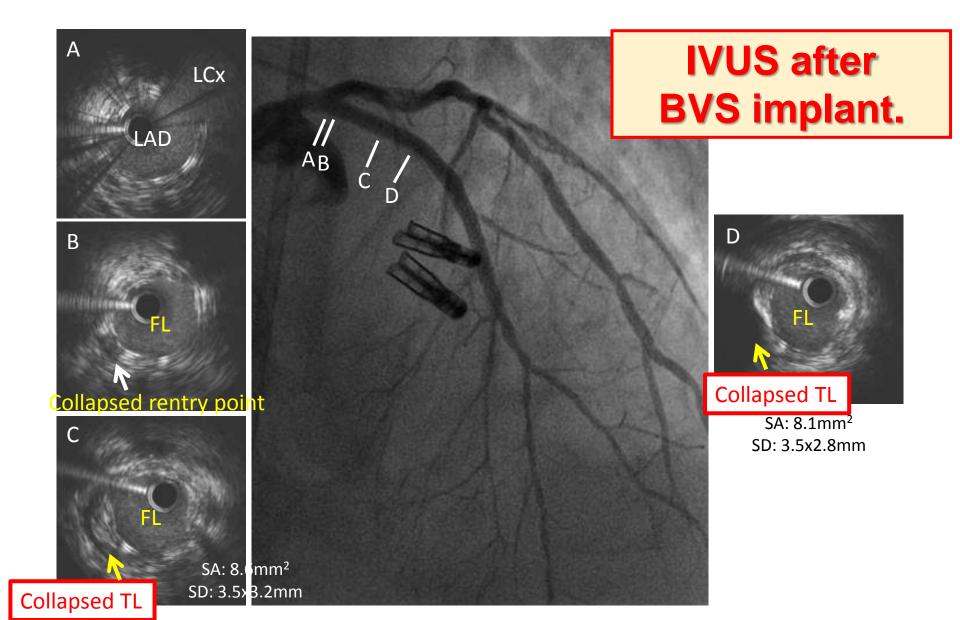
70's male LAD-CTO

Use Retrograde approach



After crossing the GW, Dilated with 2.5mm Balloon and implanted 2BVS(3.0mm and 3.5mm×18mm)

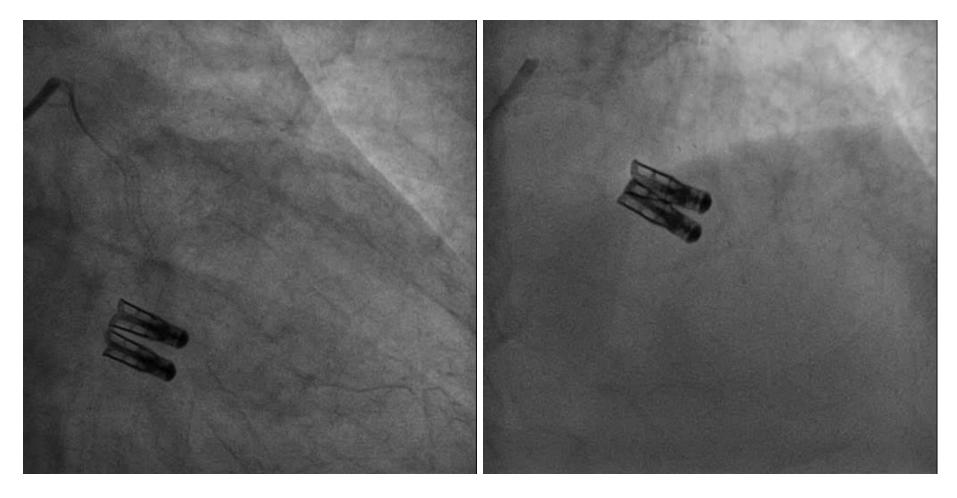
70's male LAD-CTO



One-year follow-up optical coherence tomography after implantation of bioresorbable vascular scaffolds for a chronic coronary total occlusion

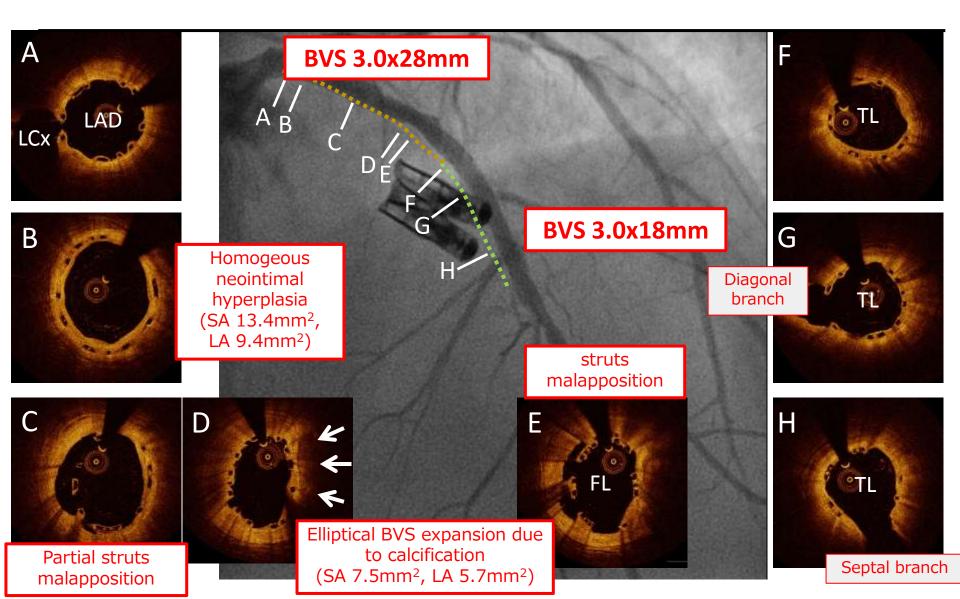
J Am Coll Cardiol Intv 2014 Toru Naganuma MD, Sunao Nakamura MD, Antonio Colombo MD et al

Angio. After 1 Year :Nicely Open !!!



OCT Findings 1 Year After Implant BVS

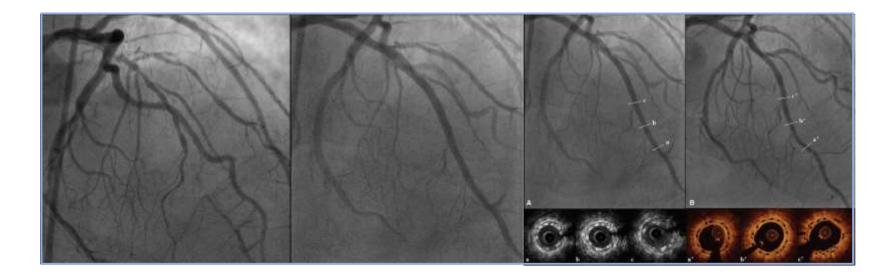
J Am Coll Cardiol Intv 2014 Toru Naganuma MD, Sunao Nakamura MD, Antonio Colombo MD et al



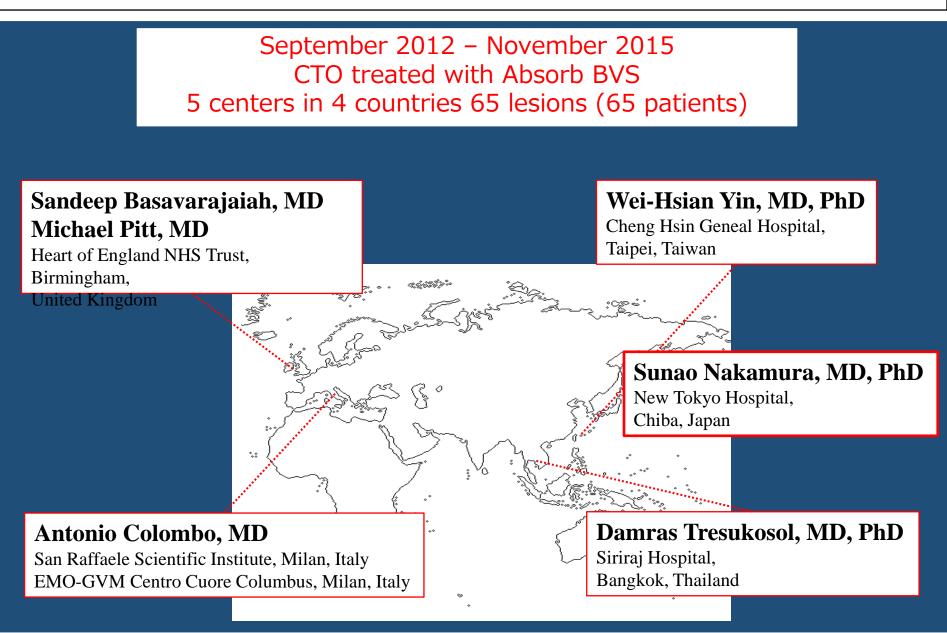
Coronary Interventions

Bioresorbable Vascular Scaffolds for the Treatment of Chronic Total Occlusions An International Multicenter Registry

Satoru Mitomo, MD; Toru Naganuma, MD; Yusuke Fujino, MD, PhD; Hiroyoshi Kawamoto, MD; Sandeep Basavarajaiah, MD; Michael Pitt, MD; Wei-Hsian Yin, MD, PhD; Damras Tresukosol, MD, PhD; Antonio Colombo, MD, PhD; Sunao Nakamura, MD, PhD



Bioresorbable Vascular Scaffolds for the Treatment of CTOs An International Multicenter Registry



Bioresorbable Vascular Scaffolds for the Treatment of CTOs An International Multicenter Registry

Study Population

- ✓ CTO PCI indication: symptomatic angina and/or ischemia on non-invasive functional test(s)
- ✓ Target vessel diameter: 2.5 4.0 mm
- ✓ DAPT: eligible for at least 12 month
- ✓ PCI strategy (antegrade or retrograde): Operator's discretion

Definition

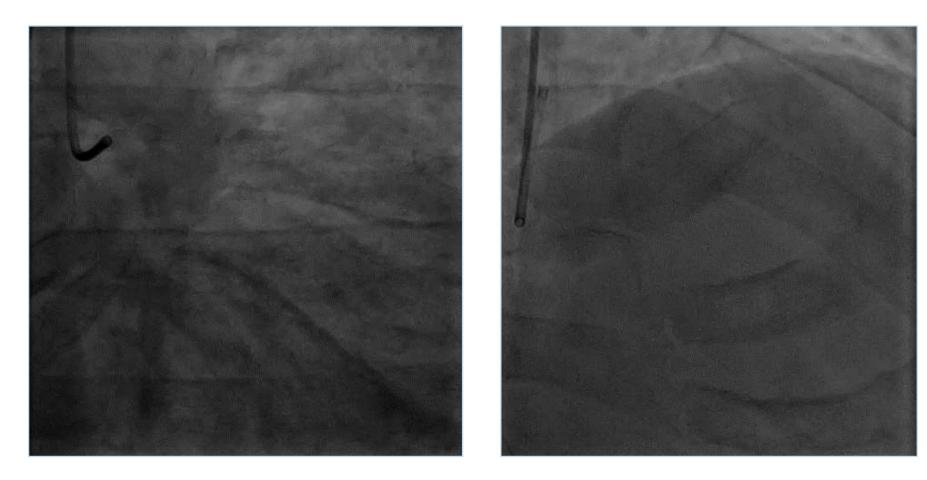
CTO: completely occluded vessel with Thrombolysis In Myocardial Infarction (TIMI) flow grade 0 through the affected segment of >3 months estimated duration Procedural success: <30% in-scaffold residual stenosis and TIMI flow grade >2 after BVS implantation

Primary outcome

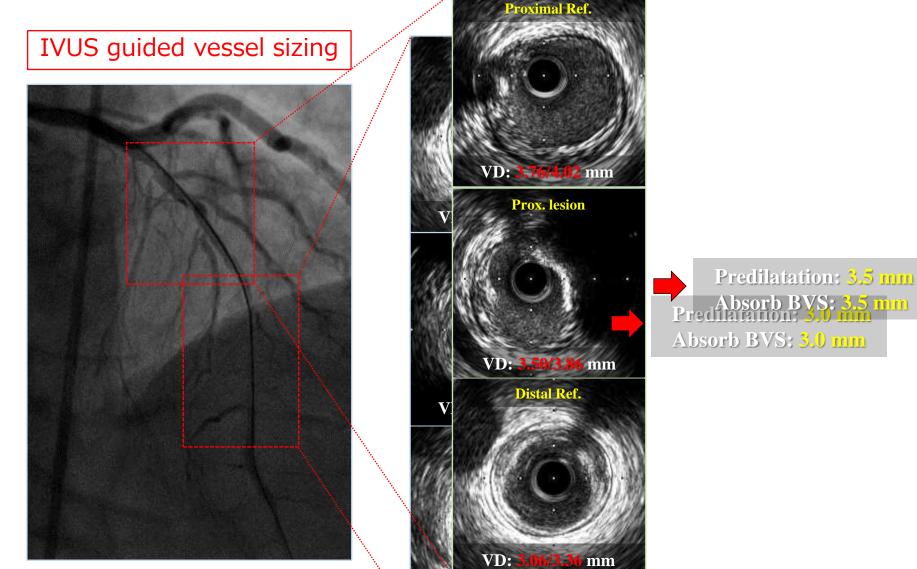
TLF: cardiac death, target vessel MI, and clinically driven TLR <u>Secondary outcomes</u>

All-cause death, clinically driven TVR, definite/probable ScT

45 year-old, male, Stable angina Coronary Rrisk F: hypertension, ex-smoker and Low bleeding risk



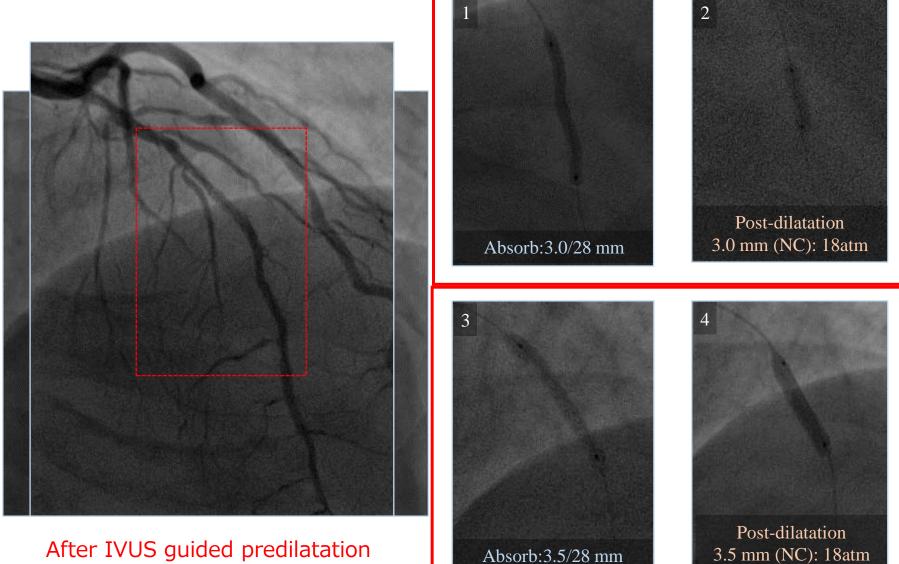
Mid LAD: CTO lesion : Ipsilateral collateral via septal (grade 3)



VD:

mm

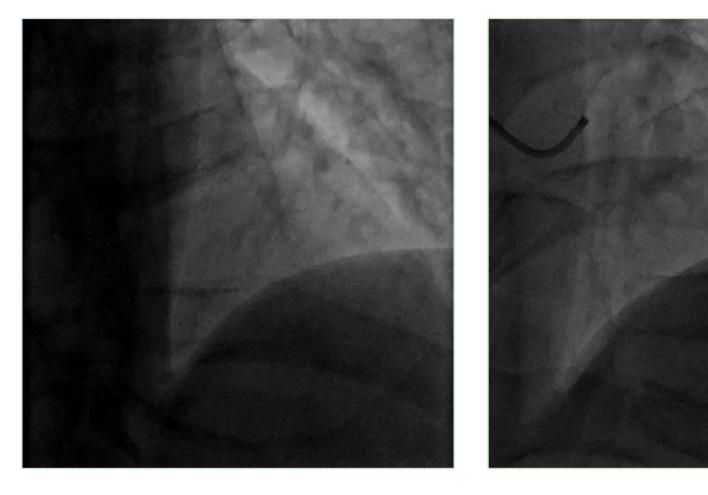
After antegrade wiring and recanalization with 1.5 mm balloon Absorb BVS: 3.5 mm



Pro.: 3.5 mm Distal-mid: 3.0 mm

Final at the index PCI

1-year follow-up Excellent angiographic results



Bioresorbable Vascular Scaffolds for the Treatment of Chronic Total Occlusions: An International Multicenter Registry.

Mitomo S¹, Naganuma T¹, Fujino Y¹, Kawamoto H¹, Basavarajaiah S¹, Pitt M¹, Yin WH¹, Tresukosol D¹, Colombo A¹, Nakamura S².

Patients Demographics

Lesion characteristics

	N=65
CTO vessel, n	
LMT	1 (1.5)
RCA	26 (40.0)
LAD	30 (46.2)
LCx	8 (12.3)
CTO location, n	
proximal	29 (44.6)
middle	31 (47.7)
distal	5 (7.7)
CTO lesion characteristics	
blunt type entry point, n	26 (40.1)
calcification, n	21 (32.3)
tortuosity, n	6 (9.2)
bent, n	13 (20.0)
bifurcation, n	23 (35.4)
bridge collateral, n	32 (49.2)
J-CTO score≧2, n	42 (64.6)
Vessel diameter by visual estimation, mm	2.97 ± 0.36
CTO length by visual estimation, mm	20.15 ± 2.97
Collaterals Rentrop grade>2	53 (81.5)
In-stent CTO, n	2 (3.1)
Number of diseased vessels	
Single vessel disease	31 (47.7)
2 vessel disease	22 (33.8)
3 vessel disease	12 (18.5)

Procedural Characteristics

	N=65		
Number of implanted BVS (/patient), n	1.8±0.7		
Patient treated with 1 BVS, n	24 (36.9)))	
Patient treated with 2 BVSs, n	31 (47.7)		
Patient treated with ≥ 3 BVSs, n	10 (15.4)	フ	
BVS diameter (/stent), mm	3.0±0.4	⊧0.4	
BVS total length (/patient), mm	47.6±19.9		
Pre-dilatation balloon diameter, mm	2.6±0.5	100 %	
Rotational atherectomy, n	3 (4.6)		
Post-dilatation balloon diameter, mm	3.3±0.3	100 %	
Max pressure for post-dilatation, atm	18.6±5.3		
Intravascular imaging, n			
IVUS, n	34 (52.3)	100 %	
OCT, n	31 (47.7)		
Antegrade approach, n	53 (81,5)		
Retrograde approach, n	12 (18.5)		
Double injection, n	51 (78.5)		
DES implantation in non-CTO lesions, n	17(26.2)		
Complete revascularization, n	52 (80.0)		

IVUS results

IVUS findings	N=34
Plaque morphology	
fibrous, n	11 (32.4)
soft, n	2 (5.9)
calcified, n	8 (23.5)
mixed (fibrous/ calcified), n	10 (29.4)
unclassified, n	3 (8.9)
Calcium arch (degree)*	120.0 (80.0-170.0)**
Proximal reference lumen area (mm ²)	9.36±2.41
Proximal reference lumen diameter (mm)	3.02 ± 0.38
Proximal reference vessel area (mm ²)	13.67±3.75
Proximal reference vessel diameter (mm)	3.65 ± 0.58
Distal reference lumen area (mm ²)	6.21±1.69
Distal reference lumen diameter (mm)	2.46 ± 0.30
Distal reference vessel area (mm ²)	9.03 ± 2.96
Distal reference vessel diameter (mm)	2.96 ± 0.48
Initial minimal vessel diameter (mm)	3.28 ± 0.52
Initial maximal vessel diameter (mm)	3.59 ± 0.47
Initial mean vessel diameter (mm)	3.44 ± 0.49
Final minimal scaffold area (mm ²)	7.74 ± 3.21
Final minimal scaffold diameter (mm)	2.71 ± 0.55
Final maximal scaffold diameter (mm)	3.11 ± 0.45
Final minimal vessel area (mm ²)	12.32±4.02
Final minimal vessel diameter (mm)	3.59 ± 0.66
Final maximal vessel diameter (mm)	3.87 ± 0.69
Edge dissection, n	3 (8.9)

OCT results

OCT findings	N=31
Calcification (>180 degrees), n	6 (19.4)
Thrombus, n	4 (12.9)
Proximal reference lumen diameter (mm)	3.31±0.71
Distal reference lumen diameter (mm)	2.29 ± 0.45
Final minimal lumen area (mm ²)	5.69±1.37
Final mean lumen area (mm ²)	7.81±1.87
Final minimal scaffold area (mm ²)	4.89±1.92
Final mean scaffold area (mm ²)	6.19±1.66
Final minimal scaffold diameter (mm)	2.54 ± 0.37
Final maximal scaffold diameter (mm)	3.51 ± 0.31
ISA at CTO lesion, n	3 (9.7)
Proximal edge ISA, n	2 (6.4)
Distal edge ISA, n	1 (3.2)
Scaffold disruption, n	3 (9.7)
Edge dissections, n	9 (29.0)

Primary endpoint:

target lesion failure (TLT); composite of cardiac mortality, target vessel MI, and clinical driven TLR

	in-hospital	follow-up
TLF, n	0	0
All cause mortality, n	0	0
Cardiac mortality, n	0	0
Target vessel MI, n		0
TLR, n	0	0
TVR, n	0	4 (13.5%)
definite/probable ScT, n	0	0

Median follow-up period 453 (IQR 230 - 703) days

Clinical outcomes

Median follow-up 453 days (IQR 230-703) Clinical follow-up 100 % Follow-up CAG: 22 cases (38.8%)

Primary endpoint TLF: 0 case

(TLF: cardiac death, target vessel MI, and clinically driven TLR)

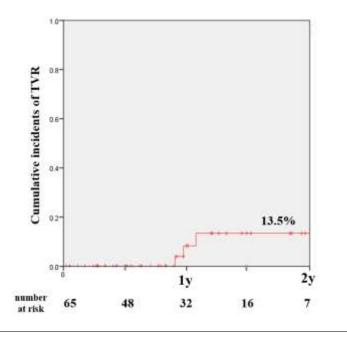
Secondary endpoint

Definite/probable ScT: 0 case

DAPT: only 1 patient stopped within 12 months because of gastric ulcer.

TVR: 4 cases

There were no cases of significant intra-scaffold restenosis, which required revascularization either clinically or angiographically.



Guide Post of BVS

- Important Reminder 6-

BVS implantation for the treatment of CTO seems feasible and safe. Appropriate lesion preparation, high-pressure post-dilatation, and the use of intravascular imaging are recommended to obtain the best possible final result.

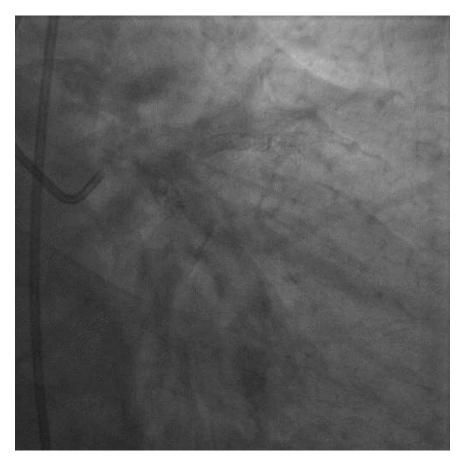


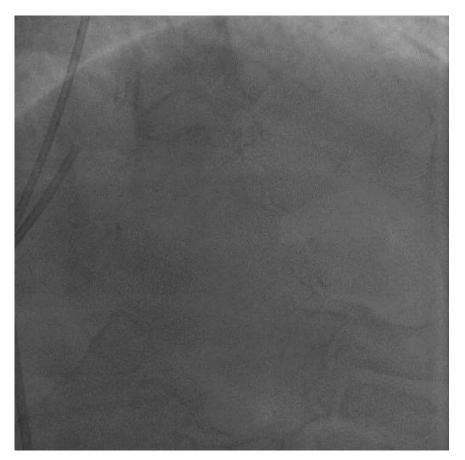
Satoru Mitomo M.D.

Circ Cardiovasc Interv. 2017;10

Satoru Mitomo, Sunao Nakamura, et al : Circ Cardiovasc Interv. 2017;10

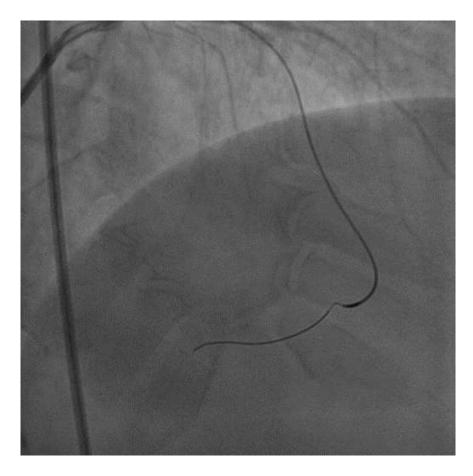
CTO lesion at RCA and well collateralized from LCA. Since bifurcation area is dull on image, it is difficult figure out the overview of RCA.

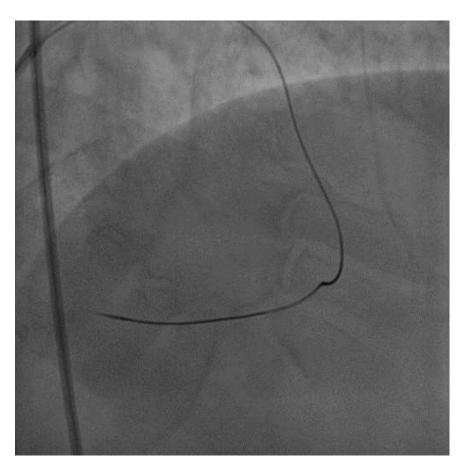




Satoru Mitomo, Sunao Nakamura, et al : Circ Cardiovasc Interv. 2017;10

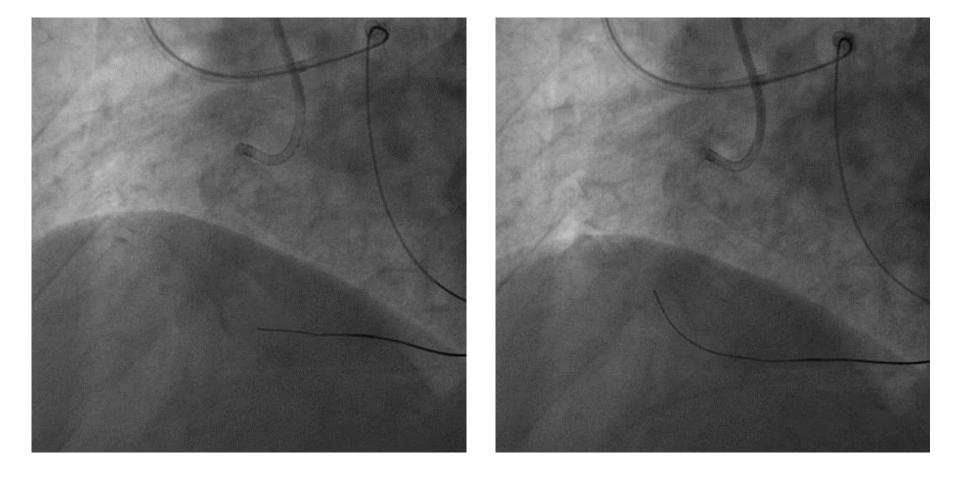
Since the RCA CTO starts from the ostium, Retrograde approach was opted from the beginning. Sion wire advanced from septum and GW was delivered to RCA distal.





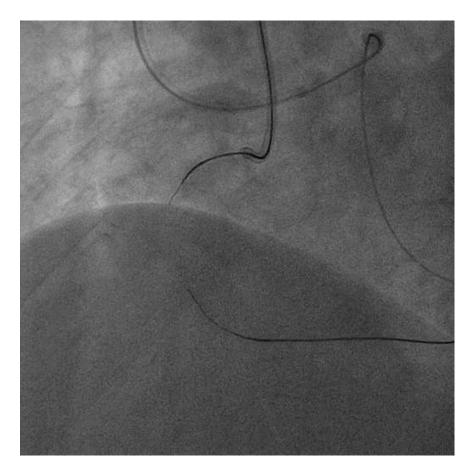
Satoru Mitomo, Sunao Nakamura, et al : Circ Cardiovasc Interv. 2017;10

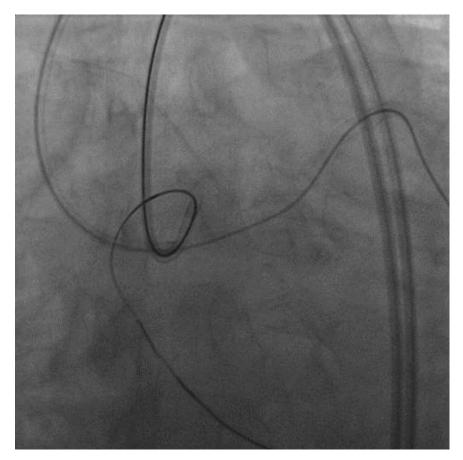
Now, you can see RCA CTO starts from the ost. And the bifurcation area looks dull shape, it was difficult to bring the wire to RCA proximal.



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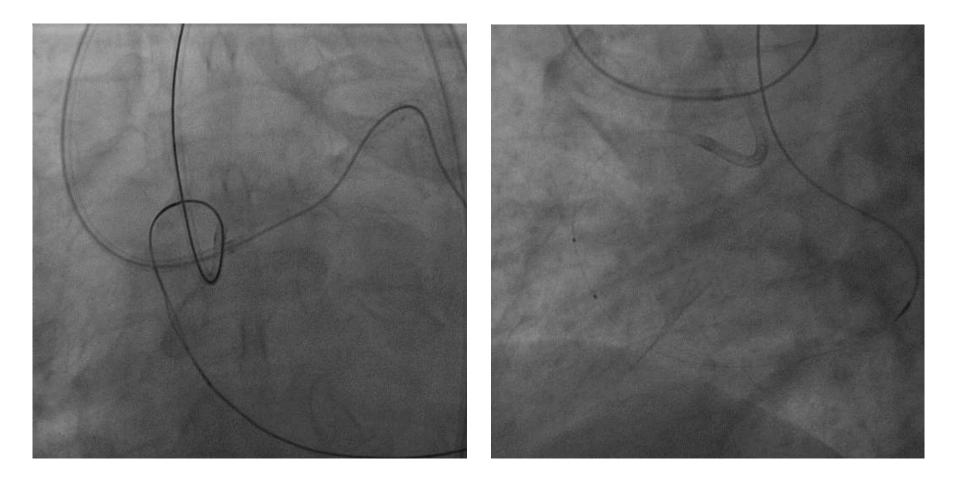
Next, we started antegrade approach. Selecting several angle views which shows both wires of retro and antegrade approach to confirm they are closing each other.





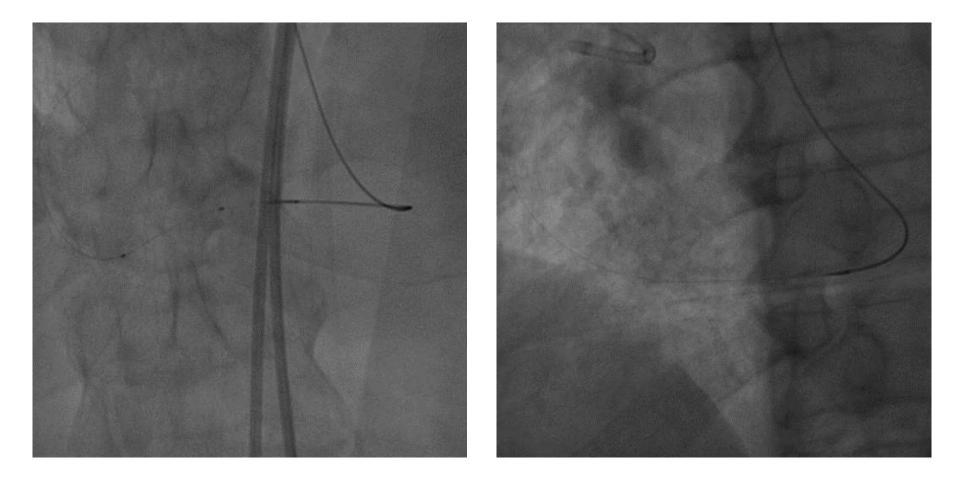
Satoru Mitomo , Sunao Nakamura, et al : Circ Cardiovasc Interv. 2017;10

Finally guidewire of retrograde approach was successfully externalized by employing Reverse Cart Technique. Ballooning was done from antegrade.



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Based on IVUS measurement, vessel diameter was around 3.0mm. BVS 3.0 x 23mm was implanted. BVS distal end was carefully positioned by using collateral angiogram.



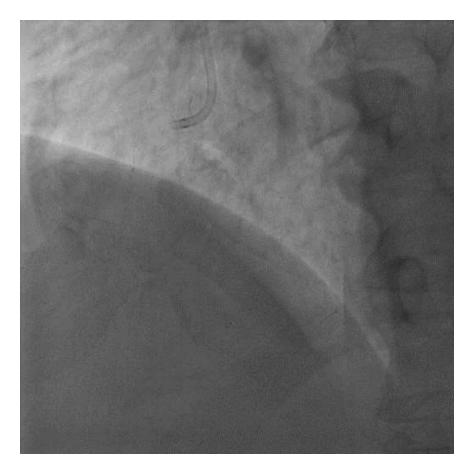
Satoru Mitomo, Sunao Nakamura, et al : Circ Cardiovasc Interv. 2017;10

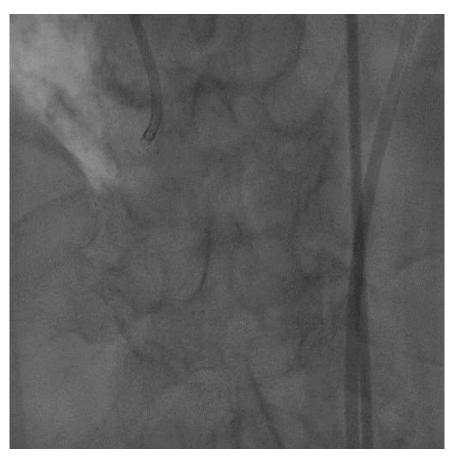
After this, 2 more same size of BVS and 1 BVS of 3.5mmx18mm were implanted followed by post dilatation with 0.5mm bigger size balloon with very high pressure.



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Final Angiogram





What the study adds

Feasibility of CTO PCI with BVS

Even in CTO lesions with high complexity (J-CTO \geq 2: 65%), if the specific implantation technique is adequately adopted, CTO PCI with BVS appears feasible.

- ✓ Intravascular guidance: 100%
- ✓ Predilatation: 100%
- ✓ Post-dilatation: 100%

Safety of CTO PCI with BVS

During follow-up (median 453 days), there were no cases of TLF and ScT (definite/probable).

 $\checkmark~$ 98.5% of the patients were on DAPT at 1-year.

What is future needed ??

Long-term F/U

The confirmations are required

- \checkmark whether vessel enlargement and plaque reduction could be observed.
- ✓ whether normal endothelialization could be observed.
- ✓ Even subintimal space, whether BRS could behave similarly to the other lesions.

Next Generation BRS

Further investigations are required

 ✓ Feasibility and safety of next generation BRSs (thinner struts, low thrombogenicity, different antiproliferative drugs, etc.) for the treatment of CTO lesions.

I strongly believe BRS future But ...

We are still on the way. ...



A long distance for touch down !!