

# Comparison of Drug-eluting Coronary Stents, Bare Coronary Stents and Self-expanding Stents in Angioplasty of Middle Cerebral Artery Stenoses

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**Objective :** The purpose of this study is to investigate the results of treatment using stent-angioplasty for symptomatic middle cerebral arterial (MCA) stenosis and comparison of in-stent restenosis between drug-eluting stents (DES), bare metal coronary stents (BMS) and self-expanding stents (SES).

**Materials and Methods :** From Jan. 2007 to June. 2012, 34 patients (mean age  $\pm$  standard deviation: 62.9  $\pm$  13.6 years) with MCA stenosis were treated. Inclusion criteria were acute infarction or transient ischemic attacks (TIAs) and angiographically proven symptom related severe stenosis. Stents used for treatment were DES (n = 8), BMS (n = 13) and SES (n = 13). National Institutes of Health Stroke Scale (NIHSS) at admission was 2.5  $\pm$  3.1 and mean stenosis rate was 79.0  $\pm$  8.2%. Assessment of clinical and angiographic results was performed retrospectively.

**Results :** Among 34 patients, periprocedural complications occurred in four cases (11.8%), however, only two cases (6.0%) were symptomatic. All patients were followed clinically (mean follow-up period; 40.7  $\pm$  17.7 months) and 31 were followed angiographically (91.2%. 13.4  $\pm$  8.5 months). There was no occurrence of repeat stroke in all patients; however, mild TIAs related to restenosis occurred in three of 34 patients (8.8%). The mean NIHSS after stent-angioplasty was 1.7  $\pm$  2.9 and 0.8  $\pm$  1.1 at discharge. The modified Rankin score (mRS) at discharge was 0.5  $\pm$  0.9 and 0.3  $\pm$  0.8 at the last clinical follow-up. In-stent restenosis over 50% occurred in five of 31 angiographically followed cases (16.1%), however, all of these events occurred only in patients who were treated with BMS or SES. Restenosis rate was 0.0% in the DES group and 20.8% in the other group ( $p = 0.562$ ); it did not differ between BMS and SES (2/11 18.2%, 3/13 23.1%,  $p = 1.000$ ).

**Conclusion :** Stent-angioplasty appears to be effective for symptomatic MCA stenosis. As for restenosis, in our study, DES was presumed to be more effective than BMS and SES; meanwhile, the results did not differ between the BMS and SES groups.

**Keywords** Intracranial stenosis, Middle cerebral artery, Restenosis, Stent-angioplasty, Drug-eluting stent, Self-expanding stent

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## INTRODUCTION

Intracranial atherosclerotic stenosis is responsible for

approximately 5% to 10% of all strokes in mixed patient populations, and in the Asian population, this is even the most commonly found vascular lesion.<sup>10)</sup> Endovascular therapy using primary angioplasty or stenting has been used in treatment of medically refractory patients with high-grade intracranial atherosclerotic stenoses.<sup>5)22)</sup> However, endovascular management of intracranial stenosis is associated with a significant number of potential complications, including acute in-stent thrombosis, thromboembolism, vessel dissection, and even rupture. To date, according to some reports, these complications during angioplasty have varied widely, ranging from 0% to more than 30%.<sup>6)7)12)13)</sup> Therefore, current treatment decisions are usually based on results of case series for each center. Prior to introduction of self-expanding stents (SES) for intracranial stent-angioplasty, the only available stents for intracranial stenotic disease were coronary balloon mounted (balloon-expandable) stents. Since that time, many studies of intracranial stent-angioplasty have proceeded in earnest.<sup>3)8)11)14)18)27)30)38)</sup>

The main trunk of the middle cerebral artery (MCA) represents one of the most common sites of symptomatic intracranial atherosclerosis.<sup>4)</sup> However, stent-angioplasty for MCA lesions remains challenging because of vascular tortuosity, small vessel diameter, and concern for perforator injury. In addition, MCA lesions are known to be highly prone to restenosis after stent-angioplasty.<sup>32)</sup>

The authors assessed the treatment results, periprocedural complications, and angiographic results of stent-angioplasty for symptomatic MCA stenosis, and then attempted to compare the results of restenosis between drug-eluting stents (DES), bare metal coronary stents (BMS), and SES.

## MATERIALS AND METHODS

### Patient characteristics and Stent selection

From January 2007 to June 2012, 35 patients (23 men, 12 women) underwent stent-angioplasty for

MCA stenosis. Patients ranged from 34 to 84 years of age (mean  $\pm$  standard deviation:  $63.3 \pm 13.6$  years). We required that candidates for stent-angioplasty have acute or subacute symptomatic MCA territorial infarction (i.e. borderzone infarction) or repeated TIAs, and severe MCA stenotic rate over 70% related to symptoms confirmed with catheter angiography. Patients with asymptomatic stenosis or mild stenosis under 70% were excluded from endovascular treatment. Because of actual assessment for clinical and angiographic results of stent-angioplasty, an additional inclusion criterion for this study was successful deployment of a stent on the lesion. Therefore, 34 patients (23 male, 11 female, mean age:  $62.9 \pm 13.6$  years) in whom stents were successfully deployed were included in this study. Stents used for treatment were DES (n = 8, 23.5%), BMS (n = 13, 38.2%), and SES (n = 13, 38.2%). The mean National Institutes of Health Stroke Scale (NIHSS) at admission was  $2.5 \pm 3.1$ , and mean stenosis rate was  $79.0 \pm 8.2\%$ . Follow-up (FU) NIHSS was evaluated immediately after the procedure, and at discharge. Evaluation of modified Rankin Score (mRS) was performed at discharge and was last performed in the Out-Patient Department. Clinical status (including NIHSS and mRS) and angiographic results were assessed retrospectively. We then compared the results between DES, BMS, and SES.

Until Apr. 2010, DES or BMS was implanted for intracranial stenosis, including MCA stenosis, and, after that time, due to the availability of stents in domestic medical practice, most patients were treated with SES (Wingspan stent, Boston Scientific, Natick, MA).

### Pre- and post-operative medications and Endovascular techniques

All patients undergoing stent-angioplasty were given dual antiplatelet therapy for at least three days before the procedure, which consisted of aspirin (100 mg - 300 mg/d orally) and clopidogrel (75 mg/d orally), except for one patient, who underwent stent-angioplasty as an emergency treatment after a loading dose of dual antiplatelet medication (aspirin 300 mg

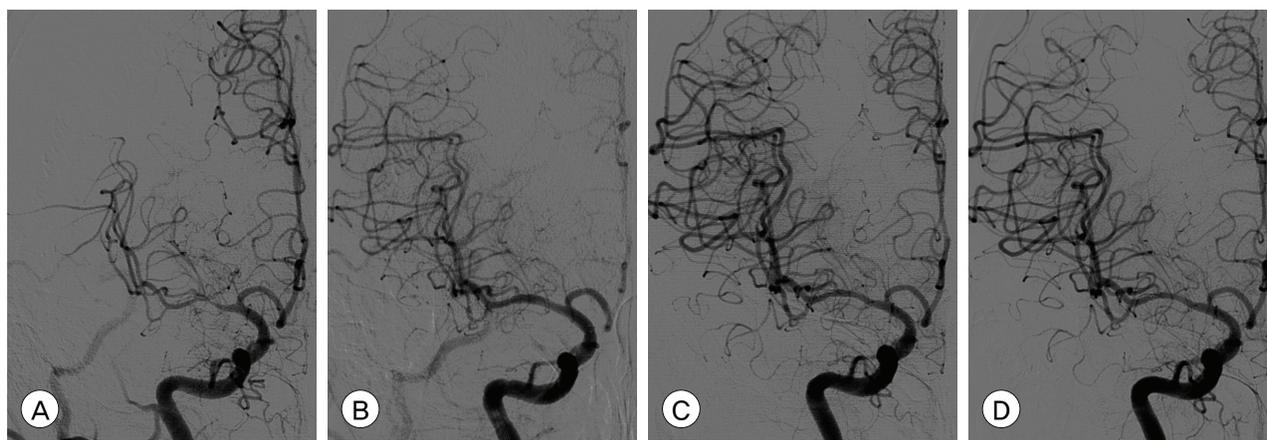
and clopidogrel 375 mg). All endovascular procedures were performed under local anesthesia by a neurosurgeon and a neuroradiologist. A 6 or 7 French guiding catheter was inserted into the proximal internal cerebral artery (ICA) after a femoral puncture. A 0.014 inch (0.35 mm) wire was then introduced through the guiding catheter with or without use of a microcatheter. In cases of coronary balloon mounted stents, including DES and BMS, after the wire had passed through the stenotic lesion under the road-map, the stent was delivered to the lesion along the wire and placed across the stenotic lesion by inflation of the balloon. The stent was deployed at the same time when the stenosis was dilated by inflation of the balloon. In cases of self-expanding stents (Wingspan stent), after the wire had passed through the lesion, a Gateway balloon was delivered to the lesion first, and inflated for pre-stent dilatation. A Wingspan stent was delivered to the lesion and placed across the lesion by unsheathing the stent-delivery catheter system after ballooning-dilatation. This procedure was performed under anticoagulation treatment with intravenous heparin, which was followed by nadroparin calcium (Fraxiparin; 2850 IU anti-Factor Xa, Sanofi Winthrop Industrie, Ambares, FR) dosed at 0.3 ml SC

q 12 hours for at least one day following the procedure. Dual antiplatelet therapy was continued after the procedure and then switched to monotherapy after at least 12 weeks (aspirin 100 mg/d orally).

#### Measurement of MCA stenosis and restenosis

All digital subtraction angiograms were analyzed by a neurosurgeon. The angiographic view that best demonstrated the stenotic lesion was identified during pre-stenting angiography. This optimal view was used as the working view for stent placement and immediate post-stenting angiography. The angle of the chosen working view was recorded and reproduced during FU angiography (Fig. 1). The degree of stenosis was calculated as the percent stenosis from the catheter angiogram using the following formula:

Percent stenosis (%) =  $100 \times (1 - S/N)$ , where S is the diameter of the residual lumen at the point of maximum stenosis and N is the width of the disease-free distal MCA (M1 segment) at the point where the walls were approximately parallel. Residual stenosis was quantified according to the vessel diameter achieved immediately after stent-angioplasty, and restenosis was calculated by comparing the lesion diameter of the follow up angiogram and that of the



**Fig. 1.** Catheter angiograms of a 66-year-old male patient who presented with repeated transient ischemic attacks of left arm weakness. (A) Preoperative right internal cerebral artery (ICA) angiogram demonstrates severe stenosis at the right middle cerebral artery (MCA) and flow compromise of the MCA territory. (B) Right MCA was fully recanalized after Wingspan stent (Boston Scientific, Natick, MA, USA) deployment. (C) Follow-up angiogram three months after stent-angioplasty. M1 segment in which stent was implanted is more remodeled than immediately after stent-angioplasty. (D) Follow-up angiogram 13 months after stent-angioplasty. No change is seen when compared with the three month follow-up angiogram.

postoperative angiogram immediately after treatment. Restenosis was categorized into two groups: insignificant to mild restenosis (0-49%), and moderate to severe restenosis (50-100%).

### Statistical analysis

Between-group comparisons were made according to the types of stents used. Age, sex, past history of medical disease (hypertension, diabetes mellitus, and hypercholesterolemia), angiographic data, including diameter of the disease-free distal MCA (reference diameter), length of stenosis, stenosis rate (preoperatively, postoperatively, and at FU angiography), and clinical status such as NIHSS and mRS were analyzed. We compared continuous variables using Student's *t*-tests and categorical variables using Pearson's Chi-Square tests. Comparisons were made according to confidence intervals of 95%. All statistical tests were two-sided and all analyses were performed using statistical software (SPSS for Windows, 15.0 standard version, IBM, NY). Values are expressed as mean  $\pm$  standard deviation. A probability value less than 0.05 was considered statistically significant.

## RESULTS

Of 35 patients, intracranial stents were successfully deployed at the first trial in 34 patients. In one patient, we attempted to perform stent-angioplasty using a Wingspan stent; stent delivery failed after successful pre-stent balloon angioplasty due to a severely tortuous ICA course. Therefore, we assessed the treatment results of stent-angioplasty with these 34 patients who were treated successfully. Overall results are shown in Table 1. NIHSS on the day after angioplasty showed improvement in most patients ( $2.5 \pm 3.1$  at admission vs.  $1.7 \pm 2.9$  after the procedure,  $p = 0.014$ ) and then continued to show improvement ( $0.8 \pm 1.1$ , at discharge,  $p = 0.001$ ) and so did mRS ( $0.5 \pm 0.9$  at discharge to  $0.3 \pm 0.8$  at last FU,  $p = 0.003$ ). Procedure-related complications occurred in four cases (11.8%); however, they were symptomatic in only two cases (5.8%). One was a case of MCA rupture after inflation of ballooning deployment of the coronary stent, and the other, a case of thromboembolism, appeared to be related to the procedure with a coronary stent not even detected angiographically during the

**Table 1. Comparisons of all patients who had undergone stent-angioplasty and the patients who have been conducted angiographic follow-up either**

Comparison variables	All patients treated (n=34)	Patients followed by angiography (n=31)	P value	
Age	62.9 $\pm$ 13.6	63.6 $\pm$ 13.6	0.837	
Reference diameter (mm)	2.5 $\pm$ 0.2	2.5 $\pm$ 0.3	0.959	
Preop. stenosis rate (%)	79.0 $\pm$ 8.2	78.9 $\pm$ 8.1	0.955	
Preop. stenosis rate (%)	6.7 $\pm$ 2.4	6.8 $\pm$ 2.4	0.885	
Used stents	DES	8 (23.5%)	0.954	
	BMS	13 (38.2%)		
	SES	13 (38.2%)		
Postop. residual stenosis rate (%)	4.0 $\pm$ 9.2	4.3 $\pm$ 9.6	0.922	
NIHSS	at admission	2.5 $\pm$ 3.1	2.5 $\pm$ 3.2	0.865
	Postop	1.7 $\pm$ 2.9	1.8 $\pm$ 3.0	0.865
	Discharge	0.8 $\pm$ 1.1	0.8 $\pm$ 1.1	0.881
mRS	Discharge	0.5 $\pm$ 0.9	0.6 $\pm$ 0.9	0.933
	At last FU	0.3 $\pm$ 0.8	0.3 $\pm$ 0.8	0.996
Clinical FU period (months)	40.7 $\pm$ 17.7	40.3 $\pm$ 18.1	0.928	

Preop= preoperative; Postop= postoperative; DES= drug-eluting stents; BMS= bare metal stents;

SES= self-expanding stents; NIHSS= National Institutes of Health Stroke Scale; mRS= modified Rankin scale; FU= follow up

procedure. Although the former was healed completely with the repetition of balloon inflation and deflation after stent deployment, a small amount of SAH and multiple scattered infarctions were developed in the MCA territory. And, in the latter, the aggravated hemiparesis showed improvement to a nearly previous state six months after the procedure. Two asymptomatic complications described above were proximal vessel (i.e. cervical segment of ICA) dissections caused by guiding catheters. In one case, it was healed spontaneously after the procedure, and the other was treated with another stent deployment without clinical sequela.

There were no cases of repeated infarction during the FU periods; however, mild TIAs occurred in three patients within one year after stent-angioplasty ( $7.2 \pm 5.5$  months). All of these patients' symptoms were mild but identical to preoperative ones, and proved to be related to severe in-stent restenosis over 70% on FU angiography. We attempted to retreat these patients with a drug-eluting balloon (DEB) in two of three patients, and with another stent in the other patient, because the stenotic lesion was located beside the center of the previously implanted stent. All of these patients have been in a symptom-free state for

at least seven months after retreatment (40 and 14 months after retreatment with DEB, seven months with another stent, respectively).

### Angiographic findings

The mean percent stenosis before the procedure was  $79.0 \pm 8.2\%$ , and the reference diameter (vessel diameter just distal to lesion) was  $2.5 \pm 0.2$  mm, and the length of the lesion was  $6.7 \pm 2.4$  mm. The residual stenosis was  $4.0 \pm 9.2\%$  immediately after stent deployment. FU angiograms were obtained in 31 of 34 patients (91.2%) at a mean of  $13.4 \pm 13.4$  months after the procedure. There were no differences in the group characteristics between whole patients who had undergone stent-angioplasty and the patients who have been conducted angiographic FU either (Table 1).

The mean percent of restenosis was  $19.2 \pm 29.1\%$  on FU angiograms. Each of the documented restenosis occurred as "in-stent" restenosis, as a sequela of thickening of the intima into the stent strut, not as a result of stent recoil or stent kinking. On the other hand, the overall rate of moderate-to-severe restenosis ( $\geq 50\%$ ) was 16.1% (five of 31 cases). Two cases were patients treated with coronary balloon-expandable stents, and the other three patients were treated with the SES (2/18, 11.1% vs. 4/13, 23.1%,  $p = 0.625$ ). Therefore, we

**Table 2. Comparisons of patients treated with coronary balloon-expandable stents (including drug-eluting stents and bare metal stents) vs. self-expanding stents.**

Comparison variables	Balloon-expandable stents (n=18)	Self-expanding stents (n=13)	P value	
Age	$61.2 \pm 15.0$	$66.9 \pm 11.0$	0.251	
Reference diameter (mm)	$2.5 \pm 0.1$	$2.5 \pm 0.4$	0.865	
Preop. stenosis rate (%)	$79.9 \pm 9.4$	$77.5 \pm 6.4$	0.426	
Lesion length (mm)	$6.5 \pm 2.0$	$7.2 \pm 3.0$	0.435	
Postop residual stenosis rate (%)	$0.0 \pm 0.0$	$10.4 \pm 12.8$	0.013	
Percent restenosis rate (%)	$13.3 \pm 28.7$	$27.3 \pm 28.7$	0.189	
Restenosis rate	2/18 (11.1%)	3/13 (23.1%)	0.625	
Angiographic FU interval (months)	$14.3 \pm 10.1$	$12.1 \pm 5.7$	0.504	
	at admission	$2.6 \pm 3.2$	$2.2 \pm 3.3$	0.752
NIHSS	Postop	$1.9 \pm 3.1$	$1.6 \pm 3.0$	0.810
	Discharge	$0.8 \pm 1.2$	$0.8 \pm 1.2$	0.880
mRS	Discharge	$0.5 \pm 1.0$	$0.6 \pm 0.9$	0.738
	At last FU	$0.4 \pm 1.0$	$0.3 \pm 0.6$	0.610

**Table 3. Comparisons of patients treated with drug-eluting stents vs. bare metal stents including coronary balloon-expandable stents and self-expanding stents.**

Comparison variables	Drug-eluting stents (n=7)	Bare metal stents (n=24)	P value
Age	65.6 ± 14.6	63.0 ± 13.5	0.667
Reference diameter (mm)	2.6 ± 0.1	2.5 ± 0.3	0.629
Preop. stenosis rate (%)	72.1 ± 7.1	80.1 ± 7.6	0.011
Lesion length (mm)	6.9 ± 1.3	6.8 ± 2.7	0.909
Postop residual stenosis rate (%)	0.0 ± 0.0	5.6 ± 10.6	0.017
Percent restenosis rate (%)	1.7 ± 4.5	24.3 ± 31.2	0.002
Restenosis rate	0/7 (0.0%)	5/24 (20.8%)	0.562
Angiographic FU interval (months)	12.4 ± 7.6	13.7 ± 8.9	0.724
	at admission	2.6 ± 3.5	0.588
NIHSS	Postop	1.7 ± 2.6	0.810
	Discharge	0.9 ± 1.5	0.886
mRS	Discharge	0.7 ± 1.5	0.598
	At last FU	0.7 ± 1.5	0.411

compared the results between the groups of patients who were treated with coronary balloon-expandable stents and SES (Table 2). In this comparison, we found that only one variable differed i.e. postoperative residual stenosis rate, even though the difference of restenosis rate itself did not reach statistical significance. There were no cases of restenosis in patients treated with DES. Thus, we compared patients treated with DES and the others (Table 3). Some variables differed between these two groups. Postoperative residual stenosis rate and percent restenosis rate differed significantly. We then compared the BMS and SES, because both stents have the same characteristics in terms of the bare metal stent, but not the same design, such as balloon-expandable stents vs. self-expanding stents, which should be deployed after sub-optimal balloon angioplasty (Table 4). In this comparison, the only significant difference was postoperative residual stenosis. However, in comparison of this between DES vs. BMS, it was nearly the same, but the percent restenosis rate was different, although it did not reach statistical significance (DES vs. BMS, 1.7 ± 4.5 vs. 20.7 ± 35.1,  $p = 0.105$ ) (Table 5). Meanwhile, in cases of SES, both were significantly different from

DES (Postoperative residual stenosis: 10.4 ± 12.8,  $p = 0.013$ , Percent restenosis rate: 27.3 ± 28.7%,  $p = 0.008$ ) (Table 5). No association was observed between development of restenosis and past history of medical disease, including hypertension ( $p = 0.394$ ), diabetes mellitus ( $p = 1.000$ ), and hypercholesterolemia ( $p = 1.000$ ).

## DISCUSSION

The success rate of stent-angioplasty was 97.1% in this study, and the procedural complication rate was 5.7% in the case of symptomatic patients. No infarction or hemorrhage was observed over one-year follow-up periods. The reported success rate and procedural risk of intracranial angioplasty varied according to the reports.<sup>9)36)</sup> Yoon et al.<sup>36)</sup> reported risk of disabling stroke or death of 6% (two of 32 patients), and mortality rate of 3% (one of 32 patients). Marks et al.<sup>20)</sup> reported a rate of periprocedural death and stroke of 8.3%, and, in a Taiwan study, Jiang et al.<sup>15)</sup> reported a complication rate up to 11.8%. In a systemic review in 2009, the success rate of stent angioplasty for intracranial arterial stenosis was reported from 71.4% to 100%, and the periprocedural minor or

**Table 4. Comparisons of patients treated with coronary bare metal stents vs. self-expanding stents.**

Comparison variables	Coronary bare metal stents (n=11)	Self-expanding stents (n=13)	P value
Age	58.4 ± 15.2	66.9 ± 11.0	0.124
Reference diameter (mm)	2.5 ± 0.8	2.5 ± 0.4	0.981
Preop. stenosis rate (%)	84.8 ± 7.1	77.5 ± 6.4	0.014
Lesion length (mm)	6.3 ± 2.3	7.2 ± 3.0	0.401
Postop residual stenosis rate (%)	0.0 ± 0.0	10.4 ± 12.8	0.013
Percent restenosis rate (%)	20.7 ± 35.1	27.3 ± 28.7	0.615
Restenosis rate	2/11 (18.2%)	3/13 (23.1%)	1.000
Angiographic FU interval (months)	15.5 ± 11.7	12.2 ± 5.7	0.372
at admission	3.1 ± 3.7	2.2 ± 3.3	0.556
NIHSS			
Postop	2.0 ± 3.6	1.6 ± 3.0	0.778
Discharge	0.8 ± 1.0	0.8 ± 1.2	0.913
mRS			
Discharge	0.4 ± 0.5	0.3 ± 0.9	0.407
at last FU	0.2 ± 0.4	0.2 ± 0.6	0.820

major stroke and death rates ranged from 0% to 50% with a median of 7.7%.<sup>13)</sup> Meanwhile, Zhang et al.<sup>37)</sup> reported feasible results of stent-angioplasty for MCA stenosis. Although the size of the study was relatively smaller than that of the previous reports, in comparison, the author's result appears to show an equivalent or slightly better clinical outcome.

As for the restenosis, in the Stenting of Symptomatic Atherosclerotic Lesions in the Vertebral or Intracranial Arteries (SSYLVA) Study, the rate of successful stent placement was 95%, however, restenosis rate of ≥ 50%

at six months was 32.4%.<sup>31)</sup> Recently, the incidence of in-stent restenosis and retreatment of this condition were intensively investigated from data of the US Wingspan Registry, and the rate of restenosis was reported as 29.8%, and an additional 4.5% of stent thromboses occurred during an average follow up period of 5.9 months.<sup>19)</sup> The rate of recurrent stroke in the subgroup with in-stent restenosis was 6.8% and the rate of TIA was 13.8%.<sup>19)</sup> In this study, overall restenosis over 50% was 16.1% (five of 31 cases) and only 9.7% (3/31) for symptomatic cases at a mean an-

**Table 5. Comparisons of patients treated with coronary drug-eluting stents vs. coronary bare metal stents and self-expanding stents.**

Comparison variables	Coronary drug-eluting stents (n=11)	Coronary bare metal stents (n=11)	P value	Self-expanding stents (n=13)	P value
Age	65.6 ± 14.6	58.4 ± 15.2	0.336	66.9 ± 11.0	0.819
Reference diameter (mm)	2.6 ± 0.1	2.5 ± 0.8	0.345	2.5 ± 0.4	0.659
Preop. stenosis rate (%)	72.1 ± 7.1	84.8 ± 7.1	0.002	77.5 ± 6.4	0.105
Lesion length (mm)	6.9 ± 1.3	6.3 ± 2.3	0.460	7.2 ± 3.0	0.796
Postop residual stenosis rate (%)	0.0 ± 5.6	0.0 ± 0.0	-	10.4 ± 12.8	0.013
Percent restenosis rate (%)	1.7 ± 4.5	20.7 ± 35.1	0.105	27.3 ± 28.7	0.008
Restenosis rate	0/7 (0.0%)	2/11 (18.2%)	0.359	3/13 (23.1%)	0.521
Angiographic FU interval (months)	12.4 ± 7.6	15.5 ± 11.7	0.540	12.2 ± 5.7	0.947
at admission	0.9 ± 2.3	3.1 ± 3.7	0.445	2.2 ± 3.3	0.797
NIHSS					
Postop	1.7 ± 2.6	2.0 ± 3.6	0.858	1.6 ± 3.0	0.942
Discharge	0.9 ± 1.5	0.8 ± 1.0	0.947	0.8 ± 1.2	0.885
mRS					
Discharge	0.7 ± 1.5	0.4 ± 0.5	0.478	0.3 ± 0.9	0.852
At last FU	0.7 ± 1.5	0.2 ± 0.4	0.390	0.2 ± 0.6	0.313

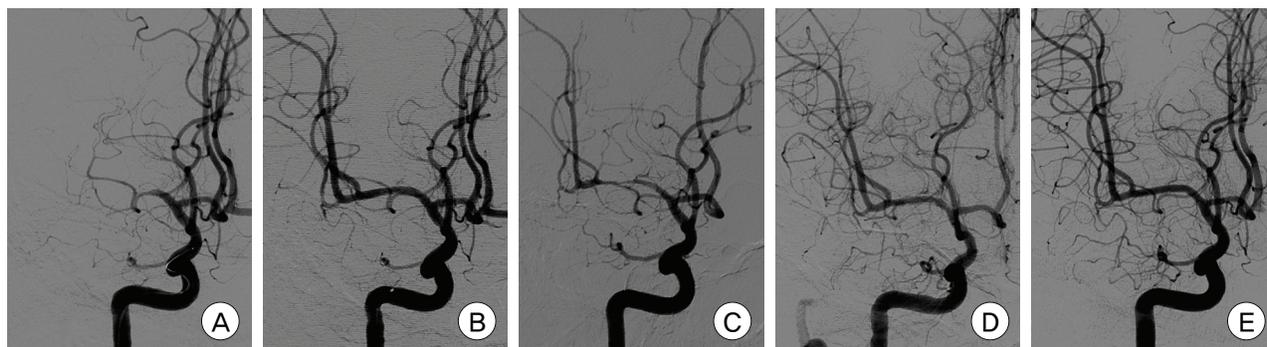
giographic follow up period of 13.4 months. However, due to the diverse composition of the kinds of stents, these figures should not be taken at face value. All cases of restenosis were bare metal stent, including coronary BMS and SES, and in case of SES (Wingspan stent), restenosis rate was 23.1%. This result is not much better than that of previous reports. However, on the contrary, restenosis did not occur in cases with DES, although the number of cases was very small (Fig. 2). DES has been known to be effective for reducing the risk of in-stent restenosis due to its controlled local release of antiproliferative agents, as compared with BMS in coronary stenting.<sup>23)28)29)</sup> Off-label use of drug-eluting coronary stents for treatment of intracranial atherosclerotic lesions has recently been reported.<sup>1)6)27)</sup> Park et al.<sup>24)</sup> reported promising results on drug-eluting stents for intracranial stenosis. The DES appears to be a good option for prevention of in-stent restenosis, however, technical difficulty due to stent stiffness remains a serious problem in generalizing and drawing definitive conclusions regarding the safety and effectiveness for in-

tracranial lesions, especially distal ICA or MCA lesions, although restenosis itself was solved in part with these stents.<sup>16)33)</sup>

Findings of this study revealed a tendency of a similar rate of restenosis in cases of coronary BMS and SES (18.2% at BMS, 23.1% at SES,  $p = 1.000$ ) (Table 4). The only meaningful difference was immediate postoperative residual stenosis rate. Percentage of residual stenosis in cases of balloon expandable stents such as DES and BMS was nearly zero, however, in case of SES, it was over 10%. This might be due to the pre-stent balloon angioplasty without simultaneous stenting of the Gateway Wingspan system. Technically, it places restrictions on sufficient dilatation of the lesions because of concern regarding serious arterial injury, such as dissection and even rupture. However, because of the similar results of the coronary BMS and SES, it might not be a risk factor. On the other hand, Shin et al.<sup>26)</sup> pointed out the speed of inflation as a risk factor for in-stent restenosis in Wingspan stenting. Although this explanation could have important implications for Wingspan stenting, it may



**Fig. 2.** Catheter angiograms of a 68-year-old female patient who presented with MCA borderzone infarct. (A) Preoperative left ICA angiogram demonstrates severe stenosis at M1 segment and definite flow compromise caused by the stenosis. (B) Postoperative angiogram after stent-angioplasty with a drug-eluting coronary stent (Endeavor, Medtronic Vascular, Santa Rosa, CA) shows no residual stenosis and fully recovered MCA flow. (C) One year follow-up angiogram shows no restenosis compared with immediate postoperative angiogram.



**Fig. 3.** Catheter angiograms of a 47-year-old male patient who presented with right MCA borderzone infarct. (A) Preoperative right ICA angiogram demonstrates MCA occlusion. (B) Postoperative angiogram after stent-angioplasty with a bare metal coronary stent (Coroflex, B. Braun, DE) shows no residual stenosis and fully recovered MCA flow. (C) Three month follow-up angiogram shows severe in-stent restenosis up to 80% with mild flow compromise. (D) Follow-up angiogram immediately after retreatment (balloon angioplasty) with a drug-eluting balloon (SeQuent<sup>®</sup> Please, B.Braun, DE) shows fully recovered MCA flow compromise, but insignificant residual stenosis remained. (E) One year follow-up angiogram after retreatment demonstrates a fully dilated M1 segment without any residual stenosis or restenosis.

not be applicable in the same way in cases of stenting with balloon-expandable coronary stents, because their use required significantly less stenting time (approximately 20 to 30 seconds in the author's series). Further studies should be required in order to clarify the association between in-stent restenosis and relevant risk factors, including technical aspects mentioned above, and other medical history.

In cases of restenosis, we attempted to administer retreatment only in symptomatic patients (three of five cases). In one case, retreatment was administered using another Wingspan system, because the stenotic lesion was located beside the center of the previously implanted stent. We treated the other two patients with the DEB (Fig. 3). All of these patients have been in a symptom-free state for at least seven months since retreatment (40 and 14 months after retreatment with a drug-eluting balloon, seven months with another stent, respectively). Several reported options, such as balloon angioplasty, another stent-angioplasty, etc, can be adopted for retreatment of restenosis. Currently, balloon angioplasty has been adopted in the majority of cases.<sup>12)</sup> Recently, the combination of mechanical balloon dilatation and local drug application was confirmed to be very efficient for treatment

of coronary in-stent restenosis, and Vajda et al.<sup>34)</sup> reported successful treatment results of a DEB for the patients with in-stent restenosis after stent-angioplasty for intracranial stenotic diseases. In addition, they introduced another combination of DEB followed by the SES, instead of a current bare balloon with a bare stent, such as a Gateway balloon and Wingspan stent.<sup>35)</sup> Although conduct of further studies is needed, the DEB appears to be a promising alternative option for in-stent restenosis.

## CONCLUSIONS

In this study, stent-angioplasty appeared to be safe and effective for MCA stenotic disease, however, a limitation of the current study was that the number of patients included was relatively small. It should prompt further studies on a larger scale.

The rate of in-stent restenosis was 16.1% as a whole, however, DES showed better results for prevention of restenosis, compared with BMS or SES. In this study, although the postoperative residual stenosis was one of the most noticeable differences between coronary balloon-expandable stents and SES, it appears to have little direct relationship with in-stent restenosis.

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