Initial Experience of Retrograde Wire Approach in Coronary Chronic Total Occlusion Intervention

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ABSTRACT

Background and Objectives: Retrograde wire approach has been emerged as a useful tool to enhance success rate in coronary chronic total occlusion (CTO) intervention. Therefore, we tried to report the initial experience of retrograde approach and its clinical implication on CTO intervention. Subjects and Methods: From February 2007 to July 2008, retrograde approaches were performed in 28 patients with 31 CTO lesions out of 61 patients. A hydrophilic coated guidewire was inserted by using microcatheter or over-the-wire (OTW) balloon through the collateral channel (septal or epicardial artery) via several strategies. Results: Mean age of patients was 63.4 ± 11.6 years. Male and female were 20 and 8 patients, respectively. The target artery with CTO lesions included the right coronary artery (45.2%), the left anterior descending artery (51.6%), and the left circumflex artery (3.2%). The mean length of CTO lesion was 18.4 ± 16.4 mm. Overall technical success rate was 64.5%. The success rate of primary attempt was 78.9%, while the success rate of immediate and secondary attempt was 41.7%. Collateral channel dissections were observed in 3 patients and no patients among these patients developed cardiac tamponade. One patient had a silent non-Q wave myocardial infarction (MI) after the procedure. One failed patient died suddenly 3 days after the procedure. After percutaneous coronary intervention (PCI) procedure, no case was performed target vessel revascularization (TVR), urgent coronary artery bypass graft (CABG), and urgent PCI. Conclusion: Retrograde approach is an evolving technique to improve the success rate of CTO intervention. After the learning curve period, this technique could be the useful tool to enhance success rate in CTO intervention. (Korean Circ J 2009;39:228-235)

KEY WORDS: Coronary occlusion; Collateral circulation.

Introduction

It has been known that successful recanalization of coronary chronic total occlusion (CTO) of coronary arteries can reduce the need for subsequent coronary artery bypass surgery1 and also increase the long-term survival of CTO patients.2,4 Recently, the development of more effective techniques and devices increased primary success rate of percutaneous coronary intervention (PCI) for CTO, but the rate is still lower than PCI for non-CTO lesions.

Retrograde approach is a novel technique of PCI for CTO. It has been shown that retrograde wire approach can enhance the procedural success rate of PCI in CTO without increasing peri-procedural complication.3,6 The retrograde approach was first introduced by Kahn and Hartlzer in 19907 and was used as a route through saphenous vein grafts for coronary angioplasty in 17 patients with prior coronary bypass surgery. Recently, the technique of retrograde approach was developed and various strategies were pioneered by Japanese cardiologists.4,8 Strategies of retrograde approach are distinctly different to the use of saphenous vein grafts which was described by Kahn and Hartlzer7 because they selected more tiny septal or epicardial collaterals for retrograde approach. These tiny collateral networks as a route of retrograde approach permitted successful CTO recanalization to perform coronary angioplasty immediately or secondly following unsuccessful antegrade attempt.

In Korea, the success rate of PCI for CTO lesions with conventional techniques is been improving lately as developing of devices and guidewires by experienced CTO-operators. However, the success rate is still not high up to 90%. Retrograde approach has been in-
introduced recently in several centers, but the clinical experiences, technical overview and the results of this technique have not been yet reported. From February 2007 to July 2008, the authors performed retrograde wire approach as 1) a primary approach, 2) immediately after failed antegrade approach, or 3) as a second procedures following unsuccessful antegrade-attempt in 28 patients with 31 CTO lesions. Therefore, we sought to describe our initial experiences, technical detail and complications.

Subjects and Methods

Patient populations
From February 2007 to July 2008, 28 patients out of 61 CTO patients were performed using retrograde wire approach with various strategies by highly experienced CTO operators. Very small distal artery sizes with poor visualization were excluded and informed consent was obtained from all patients.

Definitions
Coronary chronic total occlusion was defined as an obstruction of native coronary artery with no luminal continuity and with the thrombolysis in myocardial infarction (TIMI) flow grade 0. The duration of CTO had to be more than 3 months. The duration of occlusion was defined as the interval from previous coronary angiogram in patients with total occlusion or from the first onset of clinical symptoms suggesting the ischemic heart disease in patients without a previous angiogram.

Technical success was defined as restoration of antegrade flow with TIMI flow grade 2 or 3 and residual stenosis of <30% by quantitative coronary angiography.

In-hospital major cardiac events were defined as death, myocardial infarction (MI), target vessel revascularization (TVR), urgent PCI, and urgent coronary artery bypass graft (CABG). MI was defined as more than 3 fold increase of creatine kinase-MB (CK-MB) upper limits.

The attempt of the retrograde approach was defined as the introduction of a guidewire into the collateral channels, which connecting with the target coronary artery distal to the target CTO lesion. The occlusion length was assessed from the beginning of the occlusion to distal antegrade or retrograde vessel filling from either bridging collaterals, non-target vessel collaterals or following simultaneous left and right coronary injection by quantitative coronary angiography (QCA).

Selection of variable strategies of retrograde approach
The first strategy was a primary attempt of retrograde approach. That is, if the operator feels that antegrade wire technique seems very difficult to penetrate CTO lesion in terms of anatomical factors (heavily calcified or tortuous, angulated lesion, ostial bifurcation lesion without visible stump) and the patient has good visible, continuous collaterals, the retrograde approach can be considered as a primary attempt.

The second strategy was an immediate attempt of retrograde approach after failure or difficulty in the processing of antegrade procedure. That is, if the initial antegrade approach induced intimal dissection or unsuccessful parallel wire technique and the anatomy of CTO lesion became more difficult to do further antegrade attempts, the retrograde approach was attempted immediately.

The third strategy was a second stage procedure in selected patients several months after conventional antegrade approach failure. That is, if the patient had a heavily calcified lesion or previous big dissections which were unable to penetrate donor occluded vessel despite one or more previous attempt of antegrade approach, the retrograde approach was attempted secondly.

Techniques of retrograde approach

Procedural preparation
All patients were pretreated with aspirin and clopidogrel. Heparin was used to maintain an activated clotting time (ACT) more than 300 seconds. Also, additional heparin 2,000 or 3,000 units were injected every hour according to the ACT level. Both femoral punctures with 7-8 French (Fr) guiding catheter (GC) were used commonly. In some cases, we performed radial or brachial route with femoral approach. Because of the long access route of retrograde approach from contralateral GC to CTO lesion, GC cutting method (usually 10-15 cm cut) was used for retrograde approach (Fig. 1).

Guide wire passage through collateral channels
Guide wire (GW) passage through collateral channels means that the GW cross through collateral channel to reach the place distal to the donor CTO lesion following the procedures below:

A microcatheter was inserted into the target collateral artery with the aid of floppy GWs, and then the GW was exchanged to a hydrophilic floppy GW to cross through the collateral channel. The tip of hydrophilic GW was shaped as anatomy of collateral artery previously, and then advanced along the route of collateral. Following the hydrophilic GW into the collateral as far as possible, the microcatheter was advanced until the tip of the GW, and then the GW was pulled out. Occasionally, superselective contrast injections with small volume of dye were performed using a 2.5 cc rock syringe to visualize the collateral connection. The hydrophilic GW and the microcatheter were advanced alternatively to the target artery distal to the CTO lesion. After successful passage of hydrophilic GW into the target co-
In the coronary artery, the microcatheter was attempted to be passed into the target artery. When the microcatheter did not cross through, the microcatheter was exchanged with a 1.25 mm or 1.3 mm over-the-wire (OTW) balloon, and then dilatations of balloon at 2-4 atm in the collateral channel were performed to advance GW and OTW into the target artery distal to the CTO lesion. After the microcatheter crossed through the collateral route to the distal cap of the CTO lesion, the retrograde GW was exchanged to a Miracle series GW to attempt penetration. If the distal cap was too hard for Miracle series GW, it was exchanged to a stiffer GW such as Conquest Pro 12.0 g.

Strategies of retrograde approach

The strategies of retrograde approach included mark wire (or landmark) technique, proximal kissing wire technique, loop method (or pure retrograde wire crossing technique), and controlled antegrade and retrograde sub-intimal tracking (CART) technique. Adjunctive strategies involved anchoring balloon technique and Tornus catheter assistant.

Mark wire (or landmark) technique

After successful crossing through collateral artery into the target artery distal to the CTO lesion, the retrograde GW which has not been attempted to penetrate the CTO lesion was just used as a landmark for antegrade GW manipulation. This technique would reduce the amount of contrast dye used (Fig. 2).

Proximal kissing wire technique

On the hypothesis that the distal cap of CTO lesion is softer than the proximal cap of the CTO lesion, the retrograde GW could advance easily to the proximal part within the CTO lesion. If the tip of retrograde GW came near the proximal cap of the CTO lesion, an antegrade GW was attempted to penetrate the proximal cap, and finally, both GW could met within the true lumen, and then the antegrade GW crossed through the occluded lesion into the distal part of the target artery (Fig. 3).

Loop method

If the antegrade GW was hard to cross through the CTO lesion in the previous “proximal kissing wire technique” or complex dissection was created due to balloon dilatation with reverse CART technique which usually made procedure of antegrade wiring difficult to cross through occluded site into the true lumen, we could consider the loop method (or so called “pure retrograde wire crossing technique”). In this situation, a retrograde GW was introduced into the antegrade GC crossing through the CTO lesion, and then a microcatheter was advanced over this GW into the GC. If the MC could not be passed through GC, 2.5 mm balloon was inflated in the GC antegrade to anchor the GW.
for advancing retrograde MC into the antegrade GC. Once the retrograde MC passed through antegrade GC, the retrograde GW was exchanged mostly to Choice PT 300 cm GW (Boston Scientific, USA). By pushing the Choice PT 300 cm GW into antegrade guiding catheter through the microcatheter, the distal tip of the GW could be caught manually in the part of opposite sheath, and then a balloon was put into the CTO lesion antegrade over the GW and inflated at the occluded site. Thereafter, an antegrade GW could be inserted easily into the distal true lumen of the target artery. Following successful antegrade wiring, the Choice PT 300 cm GW was removed with microcatheter (Fig. 4).

Controlled antegrade and retrograde subintimal tracking technique

If both antegrade and retrograde GW could not been advanced successfully with previous strategies, the retrograde GW was then inserted carefully into the subintimal space of the CTO lesion parallel with antegrade GW which also within the subintimal space of the same lesion. Thereafter, a balloon was put into the subintimal space of the lesion retrogradely over the retrograde GW and then dilated to make the target space for the antegrade GW access. The target space was distinctly connected with the distal true lumen, so that the tip of antegrade GW was back out and advanced forward the

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Fig. 2. An example of landmark technique. A: proximal left anterior descending coronary artery (LAD) occlusion. B: route of collateral artery. C: a hydrophilic GW reached to distal cap of LAD CTO lesion and antegrade GW successfully crossed through CTO lesion. D: final result after drug-eluting stent implantation. GW: guide wire, CTO: chronic total occlusion.

Fig. 3. An example of proximal kissing wire technique. A: proximal left anterior descending coronary artery (LAD) occlusion. B: retrograde GW reached to distal cap of the CTO lesion. C: retrograde GW advanced to proximal cap of the CTO lesion. D: successful antegrade GW crossing after retrograde GW advanced. E: final result after drug-eluting stent insertion. GW: guide wire, CTO: chronic total occlusion.
target space. It could be easily inserted into the distal true lumen (Fig. 5).

On the contrary, the balloon also was inserted antegradely into the subintimal space within the CTO lesion and inflated in order to make the target space for retrograde GW penetration. It refers to Reverse CART technique.

Anchoring balloon technique
To reach strong backup support for antegrade guiding catheter, a second balloon with small size was inserted and inflated in the side branch proximal to the CTO lesion. This could increase the penetration power of first balloon through the CTO lesion.

Tornus catheter assistance
If the lesion was too hard to cross with microcatheter and OTW, the Tornus catheter could be used.

Statistical analysis
Descriptive analyses were performed for the data. Continuous data were presented as mean ± SD and discrete variables were expressed as percentage.

Results
Baseline clinical characteristics
Mean age of patients was 63.4 ± 11.6 years. Male and female were 20 (71.4%) and 8 (28.6%) patients, respectively. The cardiac risk factor involved diabetes mellitus, hypertension, hyperlipidemia, and smoking. The percentage of risk factors was shown in Table 1. Six patients had prior MI history, 11 patients were underwent PCI previously, and no patients had prior CABG history. Seven patients had left ventricular dysfunction with left ventricle ejection fraction (LVEF) less than 40%. Eight, 12, and 8 patients had single-, double-, and tri-
plevessel disease, respectively. Comparison of baseline characteristics between two methods of antegrade and retrograde approach in the same period was also appeared in Table 1.

**Table 1. Baseline clinical characteristics of the patients**

| Variables                  | Retrograde approach (n=28) | Antegrade approach (n=33) | p   |
|---------------------------|---------------------------|---------------------------|-----|
| Age (yrs)                 | 63.4±11.6                 | 64.5±11.2                 | 0.36|
| Male/Female (n)           | 20/8                      | 24/9                      | 0.91|
| Risk factor, n (%)        |                           |                           |     |
| Diabetes mellitus         | 9 (32.1)                  | 14 (42.4)                 | 0.409|
| Hypertension              | 16 (57.1)                 | 19 (57.6)                 | 0.973|
| Hyperlipidemia            | 6 (21.4)                  | 8 (24.2)                  | 0.795|
| Smoking                   | 6 (21.4)                  | 9 (27.3)                  | 0.597|
| Prior MI, n (%)           | 6 (21.4)                  | 6 (18.2)                  | 0.751|
| Prior PCI, n (%)          | 11 (39.2)                 | 15 (45.5)                 | 0.627|
| Prior CABG, n (%)         | 0                         | 0                         |     |
| LVEF <40%, n (%)          | 7 (25)                    | 9 (27.3)                  | 0.841|
| No. of vessel disease, n (%) |                         |                           | 0.398|
| 1                         | 8 (28.5)                  | 7 (21.2)                  |     |
| 2                         | 12 (42.8)                 | 11 (33.3)                 |     |
| 3                         | 8 (28.5)                  | 15 (45.5)                 |     |

MI: myocardial infarction, PCI: percutaneous coronary intervention, CABG: coronary artery bypass graft, LVEF: left ventricular ejection fraction, VD: vessel disease

Angiographic characteristics of the chronic total occlusion lesion

The target artery with CTO lesions included the right coronary artery (RCA, 45.2%), the left anterior descending artery (LAD, 51.6%), and the left circumflex artery (LCX, 3.2%). The target CTO lesion showed the blunted stump in 7 (22.5%), the tapered stump in 14 (45.2%), occluded at side branch in 10 (32.3%), and the bridging collateral in 4 (12.9%). The mean length of CTO lesion was 18.4±16.4 mm. Two patients had visible calcification (moderate) and one patient had severe calcification in the CTO lesion. The mean fluoroscopic time was 56.7±25.9 minutes and the average amount of contrast dye used was 342.9±71.1 mL. Angiographic characteristics of the patients with retrograde approach revealed no difference compared with that in the patients with antegrade approach which was performed in the same period (Table 2).

Strategies and procedural characteristics

Primary, immediate, and secondary attempt of retrograde approach were performed in 19, 8, and 4 CTO lesions, respectively. Overall technical success rate was 64.5% (n=20). The success rate of primary attempt was 78.9% (n=15), while the success rate of immediate and secondary attempt was 41.7% (n=5).

Septal artery in 28 CTO lesions and epicardial artery in 3 CTO lesions were chosen as a collateral chan-
Table 2. Angiographic characteristics of the patients

| Variables                        | Retrograde approach (n=31) | Antegrade approach (n=33) | p     |
|----------------------------------|----------------------------|---------------------------|-------|
| Lesion site, n (%)               |                            |                           | 0.91  |
| LAD                              | 16 (51.6)                  | 13 (39.4)                 |       |
| RCA                              | 14 (45.2)                  | 13 (39.4)                 |       |
| LCX                              | 1 (3.2)                    | 7 (21.2)                  |       |
| Stump morphology, n (%)          |                            |                           | 0.148 |
| Blunted stump                    | 7 (22.5)                   | 2 (6.1)                   |       |
| Tapered stump                    | 14 (45.2)                  | 20 (60.6)                 |       |
| Occluded at side branch          | 10 (32.3)                  | 11 (33.3)                 |       |
| Bridging collateral present      | 4 (12.9)                   | 3 (9.1)                   | 0.625 |
| Calcification (%)                |                            |                           | 0.087 |
| None to mild                     | 28 (90.3)                  | 23 (69.7)                 |       |
| Moderate                         | 2 (6.5)                    | 9 (27.3)                  |       |
| Severe                           | 1 (3.2)                    | 1 (3.0)                   |       |
| Lesion length of CTO (mm)        | 18.4 ± 16.4                | 14.3 ± 10.7               | 0.058 |
| Fluoroscopy time (min)           | 56.7 ± 25.9                | 51.3 ± 22.9               | 0.43  |
| Amount of contrast dye (mL)      | 342.9 ± 71.1               | 34.3 ± 135.5              | 0.577 |

LAD: left anterior descending coronary artery, RCA: right coronary artery, LCX: left circumflex artery, CTO: chronic total occlusion

Table 3. Strategies and procedural characteristics of retrograde approach

| Variables                        | Value (n=31) |
|----------------------------------|--------------|
| Strategies of RA, n (%)          |              |
| Primary attempt                  | 19 (61.3)    |
| Immediate attempt                | 8 (25.8)     |
| Second attempt                   | 4 (12.9)     |
| Techniques used, n (%)           |              |
| Mark wire technique              | 5 (16.1)     |
| Proximal kissing wire technique  | 14 (45.2)    |
| Loop method                      | 5 (16.1)     |
| CART technique                   | 1 (3.2)      |
| Collateral channel, n (%)        |              |
| Septal artery                    | 28 (90.3)    |
| Epicardial artery                | 3 (9.7)      |
| Successful passage of RA         | 25 (80.6)    |
| Success rate of RA               | 20 (64.5)    |

RA: retrograde approach, CART: controlled antegrade and retrograde subintimal tracking

ner of retrograde approach. Successful passage of retrograde approach which reached to the distal cap of CTO lesion was done in 25 patients (80.6%), and retrograde GW was successfully crossed through CTO lesions in 20 patients. The mark wire technique, proximal kissing wire technique, loop method, and CART technique were performed in 5, 14, 5, and 1 patient, respectively (Table 3). During the later phase (n=16), the success rate of retrograde approach increased dramatically after learning period (n=15) (40% vs. 88%).

Complications and in-hospital major adverse cardiac event

Collateral channel dissections during the procedures were observed in 3 patients and no patient among these patients developed cardiac tamponade. The reasons of dissections were GW tip induced in 2 patients and 300 cm bare wire removal without microcatheter in one patient. One of these patients had a silent non-Q wave MI after the procedure possibly due to dissection or spasm of the septal channel. After PCI procedure, no case was performed TVR, urgent CABG, and urgent PCI. One patient with unstable angina died suddenly 3 days after the failure of the second attempt of the PCI procedure for CTO lesion of LAD. A week ago, two drug eluting stents were inserted at totally occluded mid right coronary artery in this patient.

Discussion

This study included our initial experience of retrograde approach in the CTO lesions. The success rate (78.9%) of combined attempt of retrograde and antegrade approach or pure retrograde approach is not higher than that of purely antegrade approach which was performed in the same period (84.8%). The reasons of lower success rate in retrograde approach seem to be due to difficult lesion morphology, previous failed CTO lesions, and learning curve of the procedure.

Saito et al. previously explained the reasons why retrograde approach work even after previous antegrade approach failed. First reason, previous failed attempts of antegrade approach induce intimal dissection, and as a result the anatomy becomes more difficult on future occasions for further antegrade attempts. Second, the distal cap of the CTO lesion is hypothesized to be softer than the proximal cap. Third, the entry point into the CTO lesions, which is inferred from the angiogra-
phic appearance, is sometimes wrong. In this situation, a Miracle 3.0 g GW can easily pass through the lesion into the proximal true lumen from a retrograde approach.

In the study, we also performed the retrograde approach technique immediately or secondarily after antegrade approach failed and 41.7% of patients were accomplished to successful recanalization of CTO lesion. This success rate of our initial experience is not so high because our technology was in evolving state. Also, we performed CART technique only in one case because of it needed more advanced technical experiences and devices. Several group also reported that the retrograde approach can help to achieve successful recanalization of CTO lesion in the patients with failure of antegrade approach.\(^5\)\(^6\)\(^7\) Therefore, we considered retrograde approach as another choice after failure of antegrade attempt.

During the procedure, three collateral channel dissections were induced and no patient among these patients developed cardiac tamponade. One patient had a silent non-Q wave MI possibly due to the septal artery dissection or spasm of the septal channel after the procedure of retrograde approach. After PCI procedure, no case was performed TVR, urgent CABG, and urgent PCI. One patient died suddenly 3 days after the failed retrograde approach, who was previously implanted 2 overlapping drug eluting stents in the other vessel a week before the 2nd procedure. The cause of death is not yet clear, but possibly due to subacute stent thrombosis.

Although several groups\(^5\)\(^6\)\(^7\)\(^8\) reported that retrograde approach had no more increase of complication than conventional antegrade approach, retrograde approach actually is technically demanding and it includes some risk of donor vessel dissection or thrombosis and/or cardiac event during the procedure.\(^1\)\(^2\) Therefore, the operator should concern about the anticoagulation and donor vessel status, such as spasm or dissection. Also, prolonged nitrate infusion with anticoagulation is necessary, even after the procedure.

In conclusion, retrograde approach could increase the success rate of PCI after antegrade approach failed and would become a useful tool of the recanalization of CTO lesions, but further experience and technical development are needed.

Acknowledgments

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