Direct Assessment of Coronary Steal and Associated Changes of Collateral Hemodynamics in Chronic Total Coronary Occlusions

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Background—Coronary steal can occur in collateral-dependent myocardium during pharmacologically induced vasodilation. This study assessed coronary steal invasively in chronic total coronary occlusions (TCOs).

Methods and Results—In 35 consecutive patients with a percutaneous transluminal coronary angioplasty of a TCO (duration >4 weeks), coronary flow velocity (APV) by a Doppler wire and distal pressure (P_d) by a pressure wire were assessed in the collateral-dependent vascular bed before dilatation. Indexes of peripheral resistance (R_p) and for the collateral pathway, including the donor artery segment (R_c_P), were calculated. Changes of these parameters were assessed during intravenous adenosine (140 µg·kg⁻¹·min⁻¹). Adenosine caused a decrease of APV, ie, coronary steal, in 13 patients (37%; group S), an increase in 11 patients (group R), and no change in 11 patients (group N). Angiographic analysis of collateral pathways showed no difference between the groups, except that in group S all collateral connections were continuously visible but no large collaterals (>0.5 mm) were found. In group N, collaterals were least developed. The increase of APV in group R was associated with a decrease of R_p, whereas R_c_P remained unchanged. In contrast, group S showed no change in R_p but a significant increase of R_c_P, indicating an increased resistance of the donor segment.

Conclusions—Coronary steal is observed in about one third of TCOs and is associated with specific hemodynamic changes of R_p and R_c_P. Steal occurred only with well-developed angiographically visible collaterals but not with very large collaterals. (Circulation. 2002;106:435-440.)

Key Words: collateral circulation ■ coronary disease ■ vasodilation

Coronary collaterals can provide a perfusion reserve in case of increased myocardial oxygen demand.1,2 In some patients, microvascular vasodilation during exercise or pharmacological stimulation leads to a decrease of blood flow to the collateral-dependent myocardium, an observation described as coronary steal.3-9 This becomes clinically relevant when specific drugs are prescribed for patients with coronary artery disease.10

In humans, coronary steal can be detected noninvasively by perfusion scintigraphy.11-13 Direct assessment of the effect of vasodilators on the collateral circulation became possible through microsensors to record intracoronary flow velocity and pressure.14-16 These studies during balloon occlusion in the course of a percutaneous transluminal coronary angioplasty (PTCA) of nonocclusive lesions do not represent the hemodynamic situation in a totally collateral-dependent myocardium of a chronic total coronary occlusion (TCO), because there is a considerable difference between the baseline collateral function before recanalization and recruitable collateral function during balloon occlusion.17,18 The present study should assess, for the first time, the effect of pharmacologic vasodilation on collateral-dependent coronary flow distal to the occluded lesion and determine the hemodynamic changes of the collateral circulation associated with coronary steal and its relation to clinical parameters and angiographic collateral anatomy.

Methods

Patients

The study consisted of 35 consecutive patients with a TCO in whom an over-the-wire catheter could be advanced distal to the occlusion without predilatation of the occlusion. Inclusion criteria were the following: (1) duration of the occlusion >4 weeks; (2) TIMI 0 coronary flow; (3) spontaneously visible collaterals of either grade 2 (partial epicardial filling of the occluded artery) or 3 (complete epicardial filling of the occluded artery))19; and (4) written informed consent. The university ethics committee approved the study protocol.

Angiographic Analysis

Coronary angiograms of the collateral connections were obtained using a 7-inch field size, and the view with the least foreshortening was selected for analysis. Angiograms were stored on digital media in DICOM format (512×512 matrix). The pathway anatomy was...
Assessment of Collateral Hemodynamics

The PTCA was done as previously described. After the lesion was crossed by a 0.014-inch guide wire, an over-the-wire exchange catheter (Transit, Cordis) or low-profile balloon catheter (Ranger, Scimed) was advanced distal to the occlusion. The guide wire was exchanged for a pressure recording wire (PressureWire, RADI Medical Systems). The distal coronary pressure (Pd) was recorded together with the aortic pressure (Pao). Mean pressures were used for additional computation. The fractional collateral flow Qc/QD during hyperemia was calculated (Pd−Pao)/(Pao−Pra).21-23 where Pra as the right atrial pressure was substituted for by 5 mm Hg.

The pressure wire was then exchanged for the Doppler wire (FloWire, JoMed). An unaccounted contribution of antegrade flow along the exchange catheter was ruled out in all patients by lack of contrast passage along the over-the-wire catheter during proximal contrast injection into the recanalized artery and no effect on the distal Doppler signal. All Doppler flow signals were measured manually, as previously described. The velocity integral during systole and diastole and the duration of systole and diastole were measured to calculate the average peak velocity (APV).

Study Protocol

The baseline recordings started with distal pressure, followed by the Doppler flow velocity. These measurements were repeated during intravenous adenosine infusion (140 μg·kg⁻¹·min⁻¹), but because pressure recordings are less affected by the exact wire position than the Doppler flow signal, adenosine was started with the Doppler wire kept in a constant position after the baseline measurement. The APV was recorded until 3 minutes after the start of the adenosine infusion. During continuing infusion, the Doppler wire was exchanged for the pressure wire and Pd and Pao were obtained. Adenosine infusion was stopped after another 2 minutes, and Pd and Pao were recorded until they had returned to their baseline values. From the measurements during adenosine infusion, CFI and derived indexes were again calculated.

Study Groups

The collateral flow reserve is the ratio of APV during adenosine infusion and APV at baseline. The spontaneous variability of the Doppler collateral signal was 15%, as determined by continuous cascaded as proposed by Rockstroh and Brown: septal, atrial, branch-branch, and bridging collaterals. The size of collaterals was graded as discontinuous (size 0), continuous connection just visible (size 1, 0.1 to 0.3 mm), continuous of small side-branch size (size 2, 0.4 to 0.5 mm), and large (size 3, >0.5 mm) (Figure 2). The grading of the collateral size was confirmed by a caliper measurement of the minimum diameter of the collateral when it appeared maximally filled using a validated software (CAAS II, Pie Medical Imaging). The resolution was 1 pixel/0.12 mm. In case of several pathways per lesion, the one that was the first to opacify the recipient artery segment on a frame by frame analysis was considered the principal pathway for additional statistical analysis.

The quantitative angiographic analysis of left ventricular function was done with a standard software (LVA 4.0, Pie Medical Imaging). The regional wall motion was assessed by consensus of 2 experienced investigators blinded to the physiologic data. It was graded as either normal/moderately hypokinetic or severely hypokinetic/ akinetic.

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Figure 1. Schematic presentation of the electric analog model of coronary and collateral circulation (adapted from reference 9). Pao, APV, and Pd are recorded distal to the occlusion. The collateral blood flow is determined by the sum (RCP) of the resistances of the collateral (Rcoll) and the donor artery segment (Rdonor), proximal to the collateral takeoff and by the RP of the ipsilateral and contralateral artery.

Figure 2. Examples of collateral pathway grading. A, Septal collaterals from left anterior descending (LAD) to right posterior descending artery (PDA) (arrows) with discontinuation between arrow heads (size 0). B, Atrial collateral from right ventricular branch to LAD (arrows; size 2) and additional septal collaterals with continuous but small caliber (arrow heads; size 1). C, Atrial collateral from proximal circumflex branch to right posterior lateral branch (arrows; size 1). D, Singular septal collateral between LAD and PDA (arrows; size 2). E, Large branch-branch collateral between LAD and PDA (arrows; size 3). F, Large atrial collateral between circumflex and right ventricular branch (arrows; size 3).
analysis in 7 patients. Therefore, a change of the collateral flow reserve was considered significant from 1.0 by \( \pm 0.15 \). A collateral flow reserve \( \geq 0.85 \), or coronary steal, was observed in 13 patients (group S; Figure 3). A collateral flow reserve \( \geq 1.15 \) was observed in 11 patients (31%) (group R), and no significant change was observed in 11 patients (31%) (group N).

**Statistics**

Data are given as mean \( \pm \)SD. Comparisons of continuous variables among the 3 groups were done by ANOVA and a post-hoc Scheffé test. Categorical variables were compared by a Fisher’s exact test. Repeated-measures ANOVA was used to compare parameter changes during adenosine infusion. The correlation between 2 parameters was assessed by linear regression analysis. \( P < 0.05 \) was considered significant. All calculations were done on a personal computer with SPSS for Windows (Version 10.05, SPSS Inc).

**Results**

**Clinical Variables**

Patients had a right (60%) or left (40%) coronary occlusion similarly distributed within the study groups. There was no difference in age, sex, history of prior myocardial infarction, extent of coronary artery disease, regional dysfunction, and clinical symptoms, but there was a trend toward more diabetic patients in group N (Table 1). No differences in medication, such as \( \beta \)-blocker, were observed. A circumscript stenosis in the donor artery proximal to the takeoff of a collateral was observed in only 5 patients, 1 in group S and 2 each in the other groups.

**Collateral Flow Pattern and Flow Duration**

Patients of group S and group R showed a biphasic diastolic/systolic collateral flow pattern in 23 of 24 cases (96%) (Figure 3). Patients of group N showed this pattern in only 4 of 11 cases (36%) \( (P < 0.001) \). Collateral flow occurred during the complete cardiac cycle in 77% in group S and in 73% in group R but in only 27% in group N \( (P = 0.027) \). In group R the diastolic/systolic APV ratio tended to be higher than in group S (Table 2).

**Collateral Function During Adenosine Infusion**

Adenosine increased the heart rate in all groups \( (67 \pm 11 \text{ min}^{-1} \text{ to } 78 \pm 17 \text{ min}^{-1}; P < 0.001) \). The collateral flow reserve was \( 0.65 \pm 0.17 \) in group S, \( 1.40 \pm 0.16 \) in group R, and \( 1.02 \pm 0.10 \) in group N. Both baseline Doppler and pressure parameters and their changes during adenosine infusion are summarized in Table 2. Group N showed a lower diastolic APV and diastolic/systolic velocity ratio and a lower CFI compared with the other groups. The \( Q_d/Q_s \) was highest in group N. Flow and pressure indexes were not correlated \( (r = 0.16; P = 0.38) \). Collateral flow reserve was independent of the effect of adenosine on \( P_D \), which decreased even in group R while CFI increased (Figure 4).

There was a distinct difference in the effect of adenosine on \( R_P \) in groups R and S (Figure 5). \( R_P \) was slightly higher at baseline in group S, and it decreased significantly after adenosine only in group R. In contrast, \( R_{CP} \) remained unchanged in group R but increased in group S (Figure 5). No significant changes of \( R_P \) and \( R_{CP} \) were observed in group N (Table 2).

**Collateral Anatomy and Flow Reserve**

An average of \( 2.1 \pm 0.6 \) collateral pathways was observed in each patient, similar between groups. The principal pathways
were septal in 46%, atrial in 29%, branch-branch in 17%, and bridging in 9%, equally distributed between the study groups \((\chi^2, P=0.94)\). In group S, no large collaterals or small discontinuous connections were observed, and all had continuous collateral connections of size 1 (77%) and 2 (23%) (ie, between 0.1 and 0.5 mm). In group R, all collateral pathways and sizes were found, and only one had discontinuous connections. Of the 3 patients with very large collaterals (size 3), all were in group R; these were the collaterals with the lowest \(R_{CP}\) (3.7 ± 1.0 mm Hg · cm⁻¹ · s⁻¹) compared with smaller collaterals (8.5 ± 4.8 mm Hg · cm⁻¹ · s⁻¹). In group N, most patients had discontinuous or size 1 connections (82%).

**Discussion**

**Previous Studies on the Effect of Adenosine on Collateral Function**

Two groups studied the effect of adenosine on collaterals during balloon occlusion.14,16 They reported an increase of collateral flow, but a few patients also showed a decrease, ie, coronary steal. Another study assessed Doppler coronary flow velocity during intracoronary adenosine injection into the collateral-dependent artery; coronary steal occurred in 10% of patients.24 We observed a higher frequency of steal, which could be partly attributable to the fact that we applied adenosine systemically whereas it was injected locally into the collateral donor or recipient artery in previous studies.14,24 Another issue that limits the comparability with previous studies is that coronary steal was not always defined as a reduction of flow velocity, but as a drop of a pressure-derived collateral flow index.16 But because steal represents a reduction of flow, and distal coronary pressure \((P_D)\) may decrease even when flow increases during adenosine infusion (Figure 4), coronary steal should be defined by parameters of flow and not pressure.

**Comparison With Scintigraphic Studies**

The present study was done under baseline conditions similar to those in scintigraphic studies.1,4,5,12,13 Coronary steal was observed in one third of patients. Another one third had a collateral flow reserve \(>1\). When group S is compared with quantitative PET studies, the values of collateral flow reserve \((0.65\pm0.17)\) are in a similar range.5,13 In patients without

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**TABLE 1. Clinical Characteristics of Patients With TCOs**

<table>
<thead>
<tr>
<th></th>
<th>Group S</th>
<th>Group R</th>
<th>Group N</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Number</td>
<td>13</td>
<td>11</td>
<td>11</td>
<td></td>
</tr>
<tr>
<td>Age, years</td>
<td>60.7±12.8</td>
<td>59.6±9.6</td>
<td>63.8±10.9</td>
<td>0.66</td>
</tr>
<tr>
<td>Male, n (%)</td>
<td>8 (62)</td>
<td>10 (91)</td>
<td>8 (73)</td>
<td>0.39</td>
</tr>
<tr>
<td>Angina pectoris (CCS 0–4)</td>
<td>0/1/5/70</td>
<td>0/0/3/80</td>
<td>0/0/5/1</td>
<td>0.53</td>
</tr>
<tr>
<td>Heart failure (NYHA 0–4)</td>
<td>1/5/6/10</td>
<td>0/6/5/00</td>
<td>0/2/6/30</td>
<td>0.27</td>
</tr>
<tr>
<td>Diabetes, n (%)</td>
<td>3 (23)</td>
<td>1 (10)</td>
<td>5 (45)</td>
<td>0.15</td>
</tr>
<tr>
<td>Hypertension, n (%)</td>
<td>11 (85)</td>
<td>9 (82)</td>
<td>9 (82)</td>
<td>0.98</td>
</tr>
<tr>
<td>No. of diseased arteries, 1/2/3</td>
<td>4/6/3</td>
<td>7/2/2</td>
<td>5/5/1</td>
<td>0.46</td>
</tr>
<tr>
<td>Duration of occlusion: 3 months, n (%)</td>
<td>5 (38)</td>
<td>6 (55)</td>
<td>6 (55)</td>
<td>0.66</td>
</tr>
<tr>
<td>Prior Q-wave myocardial infarction, n (%)</td>
<td>7 (54)</td>
<td>7 (64)</td>
<td>8 (73)</td>
<td>0.63</td>
</tr>
<tr>
<td>Regional dysfunction, n (%)</td>
<td>10 (77)</td>
<td>7 (64)</td>
<td>8 (73)</td>
<td>0.77</td>
</tr>
<tr>
<td>Ejection fraction, %</td>
<td>62.5±11.2</td>
<td>68.6±18.8</td>
<td>60.1±18.9</td>
<td>0.47</td>
</tr>
<tr>
<td>LVEDP, mm Hg</td>
<td>18.0±8.0</td>
<td>15.8±6.0</td>
<td>20.1±10.2</td>
<td>0.55</td>
</tr>
</tbody>
</table>

CCS indicates Canadian Cardiovascular Society classification of chest pain; LVEDP, left ventricular end diastolic pressure; and NYHA, New York Heart Association classification of heart failure.

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**TABLE 2. Collateral Flow, Pressure, and Resistance Indexes in TCO Before and After Adenosine Infusion**

<table>
<thead>
<tr>
<th></th>
<th>Before Adenosine</th>
<th>After Adenosine</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>APV ↓ Occl, cm/s</td>
<td>11.3±6.0</td>
<td>7.6±4.8</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Diastolic/systolic APV ratio</td>
<td>0.82±0.42</td>
<td>0.71±0.62</td>
<td>0.72</td>
</tr>
<tr>
<td>(P_a), mean, mm Hg</td>
<td>109±16</td>
<td>94±11</td>
<td>0.006</td>
</tr>
<tr>
<td>(P_s), mean, mm Hg</td>
<td>42±11</td>
<td>29±6</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>CFI</td>
<td>0.36±0.21</td>
<td>0.24±0.16‡</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>(Q_a/\varnothing)</td>
<td>0.25±0.07</td>
<td>0.30±0.08</td>
<td>0.38±0.14*</td>
</tr>
<tr>
<td>(R_{ap}), mm Hg · cm⁻¹ · s⁻¹</td>
<td>8.35±5.15</td>
<td>17.63±17.17</td>
<td>0.02</td>
</tr>
<tr>
<td>(R_{ap}), mm Hg · cm⁻¹ · s⁻¹</td>
<td>5.75±4.54</td>
<td>7.42±5.76</td>
<td>0.18</td>
</tr>
</tbody>
</table>

Note: Values are means ± SD. \(P\) values for changes within groups; difference between group N and S, \(*P<0.05\); between group N and R, †\(P<0.05\); between group S and R, ‡\(P<0.05\).
steal, flow increased >4 times in selected patients without myocardial infarction and regional dysfunction,1,2 but in patients with regional dysfunction2 and multivessel disease,4,5,13 the flow reserve was similar to our study (1.40±0.16). Even though we measured collateral-dependent flow velocity in an epicardial artery and scintigraphy assesses myocardial perfusion, this quantitative agreement supports the validity of the invasive approach.

Direct Confirmation of the Collateral Network Model in Man

Gould and colleagues9,25 specified the following 3 assumptions required for the occurrence of steal: (1) the collateral resistance is not negligible; (2) the microvasculature distal to the occlusion, being already maximally dilated, lacks a vasodilation reserve, and (3) the epicardial resistance of the supply artery causes a pressure drop proximal to the collateral origin during adenosine-induced hyperemic flow. All 3 assumptions are supported by our data. First, coronary steal required well-developed collaterals,14,16,24 whereas collaterals in group N had a lower CFI, higher R CP, and predominantly a systolic collateral flow pattern, as evidence of reduced collateral function.18,26

As shown previously, collateral resistance is not negligible.14,16,18 The assumption that large collaterals, because of a low resistance, would not show steal25 was confirmed by our study, because large collaterals were only observed in group R but not in group S. The lack of a vasodilatory reserve is demonstrated by the unchanged R P in group S, whereas it was decreased significantly in group R. On the other hand, the higher R P in group N and its lack of response to adenosine could be related to the higher proportion of diabetics with a known prevalence of microvascular dysfunction.27

We did not record directly the coronary pressure in the donor artery at the collateral origin. The multiple coexisting collateral pathways make such an approach difficult and unreliable; in only 2 of 13 patients with steal we observed a singular collateral takeoff. The increase of RCP in patients with steal during adenosine infusion represented an increased resistance of the complete collateral supply pathway, which includes the donor artery segment proximal to the collateral takeoff (Figure 1). Assuming a constant collateral resistance, the increase of RCP indicated an increase of the donor segment resistance or, stated another way, decrease of distal coronary donor pressure. This does not require a circumscrip epicardial stenosis, as recently demonstrated by a pressure drop during hyperemia in angiographically normal segments of patients with coronary artery disease attributable to diffuse atherosclerosis.28,29

Collateral Pathway and Coronary Steal

Angiography is of limited value for assessing function of collaterals,23,30 but it illustrates their anatomy.31 Angiographic assessment of collaterals is mostly semiquantitative.32 A refined quantitative approach was recently suggested but requires high-quality cine films.20 With the limitation of a lower spatial resolution of digital media, we analyzed the collateral anatomy and size relative to the capillary bed to test an extension of the network model of coronary steal. It was suggested that prearteriolar collaterals would be required for coronary steal.25 This was confirmed in group S, where all collaterals showed continuous connections between donor and recipient segments. Another assumption was that an increase of flow would occur in barely visible postarteriolar collaterals that pass through the microvascular bed of the donor artery. Their flow capacity should increase with peripheral vasodilation of the donor microcirculation. This could not be confirmed in our study, because these postarteriolar collaterals were predominantly observed in group N and indicated a low collateral function. Visible connections were the principal requisite for the incidence of coronary steal but also for a positive collateral flow reserve. However, there was no anatomic pathway that predicted the response to adenosine.
Study Limitations
Flow and pressure recorded distal to an occlusion are approximations of collateral function, because they assess only that part of collateral perfusion that reaches the occluded epicardial artery. Collateral perfusion through intramyocardial pathways may be underestimated. An increase of collateral flow velocity during adenosine might lead to a shear stress–related vasodilation and, consequently, an underestimation of true volume flow. However, the flow velocities involved were far below those evoked during the observation of flow-mediated epicardial vasodilation. Nitroglycerine application would have avoided the influence of minor diameter changes, but because nitroglycerine itself influences collateral hemodynamics, we did not apply it during collateral flow measurements. Right atrial pressure (PRA) was not directly measured.

Clinical Implications
In patients with coronary artery disease, coronary steal can occur without a circumscript donor artery lesion but requires the presence of diffuse atherosclerosis that causes a pressure block. Collateral artery. Collateral perfusion through intramyocardial pathway may be underestimated. An increase of collateral flow velocity during adenosine might lead to a shear stress–related vasodilation and, consequently, an underestimation of true volume flow. However, the flow velocities involved were far below those evoked during the observation of flow-mediated epicardial vasodilation. Nitroglycerine application would have avoided the influence of minor diameter changes, but because nitroglycerine itself influences collateral hemodynamics, we did not apply it during collateral flow measurements. Right atrial pressure (PRA) was not directly measured. This influences the calculation of absolute values of pressure indexes, but because PRA does not considerably change during adenosine infusion, it would not decisively affect the assessment of the intraindividual response to adenosine.

References