Prediction of Acute Left Main Coronary Artery Obstruction by 12-Lead Electrocardiography
ST Segment Elevation in Lead aVR With Less ST Segment Elevation in Lead V1

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OBJECTIVES
We sought to determine the electrocardiographic (ECG) features associated with acute left main coronary artery (LMCA) obstruction.

BACKGROUND
Prediction of LMCA obstruction is important with regard to selecting the appropriate treatment strategy, because acute LMCA obstruction usually causes severe hemodynamic deterioration, resulting in a less favorable prognosis.

METHODS
We studied the admission 12-lead ECGs in 16 consecutive patients with acute LMCA obstruction (LMCA group), 46 patients with acute left anterior descending coronary artery (LAD) obstruction (LAD group) and 24 patients with acute right coronary artery (RCA) obstruction (RCA group).

RESULTS
Lead aVR ST segment elevation (>0.05 mV) occurred with a significantly higher incidence in the LMCA group (88% [14/16]) than in the LAD (43% [20/46]) or RCA (8% [2/24]) groups. Lead aVR ST segment elevation was significantly higher in the LMCA group (0.16 ± 0.13 mV) than in the LAD group (0.04 ± 0.10 mV). Lead V1 ST segment elevation was lower in the LMCA group (0.00 ± 0.21 mV) than in the LAD group (0.14 ± 0.11 mV). The finding of lead aVR ST segment elevation greater than or equal to lead V1 ST segment elevation distinguished the LMCA group from the LAD group, with 81% sensitivity, 80% specificity and 81% accuracy. A ST segment shift in lead aVR and the inferior leads distinguished the LMCA group from the RCA group. In acute LMCA obstruction, death occurred more frequently in patients with higher ST segment elevation in lead aVR than in those with less severe elevation.

CONCLUSIONS
Lead aVR ST segment elevation with less ST segment elevation in lead V1 is an important predictor of acute LMCA obstruction. In acute LMCA obstruction, lead aVR ST segment elevation also contributes to predicting a patient’s clinical outcome. (J Am Coll Cardiol 2001; 38:1348–54) © 2001 by the American College of Cardiology

Acute obstruction of the left main coronary artery (LMCA) is not frequently encountered (1). A large part of the myocardium of the left ventricle is perfused by the LMCA, and its acute obstruction thus causes severe hemodynamic deterioration, frequently resulting in rapid fatality (2,3). Prediction of acute LMCA obstruction is important with regard to selecting the appropriate treatment strategy and estimating the prognosis.

Recently, Engelen et al. (4) reported that lead aVR ST segment elevation was observed in acute obstruction of the left anterior descending coronary artery (LAD) proximal to the major septal branch, but not in acute LAD obstruction distal to the major septal branch. They concluded that lead aVR ST segment elevation associated with proximal LAD obstruction was caused by transmural ischemia of the basal part of the septum. This led us to assume that acute LMCA obstruction also causes lead aVR ST segment elevation through disturbance of major septal branch blood flow—that is, interruption of LAD blood flow. In fact, lead aVR ST segment elevation during angina pectoris attacks has been reported in patients with significant LMCA stenosis (5). The right coronary artery (RCA) also perfuses the septum through its septal perforator branches, to some extent, and acute RCA obstruction may possibly cause lead aVR ST segment elevation in some cases.

Both acute LMCA and LAD obstruction generally produce anterior wall ischemia, resulting in ST segment elevation in the precordial leads. One can easily assume that LMCA, but not LAD, obstruction is ordinarily associated with posterior wall ischemia. Acute LMCA obstruction may show ST segment shifts in the precordial leads different from those found in acute LAD obstruction, due to concomitant posterior wall ischemia, which produces reciprocal changes in the precordial leads.

Based on these reported findings, we hypothesized that
METHODS

Patients. From January 1988 to January 1998, 16 patients (14 men and 2 women; mean [±SD] age 65 ± 9 years [range 37 to 78]) were admitted to the hospital with acute LMCA obstruction (LMCA group). The 12-lead ECG findings in these patients were compared with those in 46 consecutive patients with the culprit lesion at location segment no. 6 (LAD group) and in 24 patients with the culprit lesion at location segment no. 1, according to the American Heart Association classification. All patients were admitted between December 1994 and January 1998. Patients with significant stenosis (>75% lumen diameter stenosis), other than that in the culprit lesion, were excluded from the LAD and RCA groups. Patients with acute left circumflex coronary artery (LCx) obstruction was excluded from the present study. The reasons for the exclusion were: 1) characteristic or diagnostic ST segment elevation in the precordial leads and lead aVR was not reported (6); and 2) LCx obstruction was anatomically unable to cause ischemia at the basal part of the septum, which has been thought to elevate the ST segment in lead aVR, as described earlier. Patients with a recurrent myocardial infarction (MI) were excluded from the analysis. The LAD group consisted of 41 men and five women whose ages ranged from 39 to 80 years (61 ± 9 years). The RCA group consisted of 18 men and six women whose ages ranged from 40 to 80 years (61 ± 10 years). There were no significant differences in gender or age among the three groups. All patients in the three groups were admitted within 12 h after the onset of the acute MI. The culprit lesion was determined by emergency coronary angiography and confirmed later by two cardiologists who were not aware of any ECG findings. The culprit lesion was defined when the lesion was totally occluded or showed severe stenosis. When the lesion showed severe stenosis, the lesion with flow delay (Thrombolysis in Myocardial Infarction [TIMI] flow grade 1), with angiographic findings that suggested local dissection or thrombus, was defined as the culprit lesion (7). Diagnosis of acute MI was established by ST segment elevation, defined subsequently, in more than two leads, associated with typical, severe anterior chest pain and confirmed by elevation of both serum creatine kinase (CK) and its MB isoenzyme (CK-MB) greater than two times the normal upper limit during the patient’s clinical course. The upper limits of the range (2 SD above the mean value) for the healthy age- and gender-matched control subjects for serum CK and CK-MB were 180 IU/l and 19 IU/l, respectively. The clinical and angiographic characteristics of the patients in the LMCA group are summarized in Table 1. In three patients in the LMCA group, serum levels of CK and CK-MB were not elevated more than two times the normal upper limit, as these patients died early.

Electrocardiography. The 12-lead ECGs recorded on admission before emergency coronary angiography were analyzed. A ST segment shift was determined as the mean value of five successive beats measured at 60 ms after the J point of the QR5 complex. ST segment elevation was defined as present when ST segment elevation was >0.05 mV in the limb leads and ST segment elevation was >0.1 mV in the precordial leads. The data for ST segment shifts were subjected to statistical analysis.

Initially, inter-observer and intra-observer differences were checked by using 10 randomly selected ECG samples from the three groups. Measurements were then performed by two observers who were not aware of any angiographic findings.

Analysis. In emergency coronary angiography, collateral circulation was classed into four grades according to the grading system of Rentrop et al. (8). First, the incidence of ST segment elevation was examined in all leads, including aVR, and compared among the three groups. Further, ST segment elevation in lead aVR and the precordial leads was compared between the LMCA and LAD groups, and the differences in ST segment elevation between leads aVR and V₁ were examined. For comparison between the LMCA and RCA groups, a comparison was made for the inferior leads in addition to lead aVR.

Statistics. Data are expressed as the mean value ± SD. For univariate analysis, the Fisher exact probability test was used to compare the prevalence of ST segment elevation between two groups. The unpaired Student t test was used to compare the extent of ST segment shift between two groups. We also performed stepwise linear multivariate discriminant analysis using a personal computer (Compaq, Prolina 5120, Houston, Texas) with the appropriate software. In this analysis, the dependent variables were those of the two groups being compared. Because leads II, III and aVF (i.e., the inferior leads) showed essentially the same prevalence of abnormalities on univariate analysis, lead aVF
### RESULTS

When evaluating the sample ECGs, the inter-observer and intra-observer differences in the limb leads averaged 0.01 ± 0.02 mV and 0.01 ± 0.03 mV, respectively. Similarly, the inter-observer and intra-observer differences in the precordial leads averaged 0.01 ± 0.04 mV and 0.00 ± 0.02 mV, respectively. Therefore, intra-observer and inter-observer variations were acceptably small and did not affect the validity of the results.

**Incidences of ST segment shift among the LMCA, LAD and RCA groups.** Figure 1 shows representative 12-lead ECGs at hospital admission for one patient from each group, and the incidence of ST segment shift (>0.05 mV in the limb leads and >0.1 mV in the precordial leads) on the 12-lead ECG is summarized in Figure 2. Lead aVR showed ST segment elevation in 88% (14/16) of patients in the LMCA group, whereas ST segment elevation was found in 43% (20/46) of patients in the LAD group and only 8% (2/24) of patients in the RCA group. Lead aVR, an inferior lead, clearly showed a higher incidence of ST segment elevation (96% [23/24]) in the RCA group than in the LMCA group (0% [0/16]) or in the LAD group (9% [4/46]). Similar incidences were seen in the other inferior leads (i.e., leads II and III).

**Comparison between the LMCA and LAD groups.**

**UNIVARIATE ANALYSIS.** Table 2 summarizes the results of a comparison of ST segment elevation in leads aVR and V1 between the LMCA and LAD groups. Significantly higher ST segment elevation in lead aVR was observed in the LMCA group than in the LAD group. Conversely, ST segment elevation in lead V1 was significantly lower in the LMCA group than in the LAD group. Lead V2 showed similar results (LMCA group vs. LAD group: 0.15 ± 0.21 mV; p < 0.01). Lead V3 tended to show similar results (0.27 ± 0.40 vs. 0.44 ± 0.35 mV; p > 0.05 and < 0.10).

As a consequence, 13 patients (81%) in the LMCA group and only 9 patients (20%) in the LAD group showed greater or equal ST segment elevation in lead aVR compared with that in lead V1. The finding that ST segment elevation in lead aVR was greater than or equal to that in lead V1 distinguished the LMCA group from the LAD group, with 81% sensitivity, 80% specificity and 81% accuracy.

**MULTIVARIATE ANALYSIS.** Stepwise linear multivariate discriminant analysis identified leads aVR and V1 as leads in which ST segment elevation significantly contributed, positively and negatively, respectively, to distinguishing the
LMCA group from the LAD group (Table 2). The results were completely in agreement with the results obtained by univariate analysis.

**Comparison between the LMCA and RCA groups.**

**UNIVARIATE ANALYSIS.** ST segment elevation in lead aVR (>0.05 mV) occurred with a significantly higher incidence in the LMCA group (88% [14/16]) than in the RCA group (8% [2/24]), and the elevation distinguished the LMCA group from the RCA group, with 88% sensitivity, 92% specificity and 90% accuracy. Conversely, the LMCA group clearly showed a lower incidence of ST segment elevation in leads II (6% [1/16]), III (13% [2/16]) and aVF (0% [0/16]) than the RCA group (92% [22/24], 96% [23/24] and 96% [23/24], respectively). The ST segment elevations in leads II, III and aVF were useful for distinguishing the LMCA group from the RCA group, with high sensitivity (92%, 96% and 96%, respectively), specificity (94%, 88% and 100%, respectively) and accuracy (93%, 93% and 98%, respectively). Lead aVF showed the highest sensitivity, specificity and accuracy for distinguishing the LMCA group from the RCA group.

**MULTIVARIATE ANALYSIS.** Stepwise linear multivariate discriminant analysis selected lead aVF rather than lead aVR as the lead whose ST segment shift contributed significantly to distinguishing the LMCA group from the RCA group, which was consistent with the results of univariate analysis in that lead aVF showed the highest sensitivity, specificity and accuracy.

**ST segment shift in lead aVR and patients’ clinical outcomes in the LMCA group.** Table 3 summarizes the findings about the relationship between the ST segment shift in lead aVR and patients’ clinical outcomes in the LMCA group. Death occurred more frequently in patients with higher ST segment elevation in lead aVR than in those with less severe elevation. When ST segment elevation of 0.15 mV was used as the cut-off value, death was predicted with 75% sensitivity, 75% specificity and 75% accuracy.

**Figure 1.** Representative 12-lead electrocardiogram tracings at admission in a patient in (A) the left main coronary artery (LMCA) group, (B) the left anterior descending coronary artery (LAD) group and (C) the right coronary artery (RCA) group. In the patient in the LMCA group, ST segment elevation is apparent in lead aVR. In the patient in the LAD group, marked ST segment elevation in the precordial leads is seen, whereas a ST segment shift in lead aVR is negligible. In the patient in the RCA group, ST segment elevation in the inferior leads is marked.
Differences in ST segment elevation between lead aVR and lead V1 did not contribute significantly to predicting mortality.

Stepwise multivariate linear discriminant analysis also showed that ST segment elevation in lead aVR significantly contributed to predicting patients’ clinical outcomes in the LMCA group (Table 3).

DISCUSSION

The present study revealed that high ST segment elevation in lead aVR, compared with lead V1, was a useful indicator for predicting acute LMCA obstruction, which requires immediate aggressive treatment. In acute LMCA obstruction, the ST segment elevation in lead aVR also contributed to predicting the patients’ clinical outcomes. Thus, the present results are clinically useful for selecting the treatment strategy for patients with acute MI.

Total obstruction or severe stenosis with flow delay in the LMCA lesion was demonstrated by emergency coronary angiography in all patients in the LMCA group. Similarly, neither the LAD group nor the RCA group included patients with significant stenotic lesion(s) other than the culprit lesion. ST segment elevation after acute MI changes with time. This study included only patients who were admitted within 12 h (3.2 ± 2.7 h) from the onset of acute MI, during the period when acute ST segment elevation is

Figure 2. The incidences of ST segment elevation in each lead in the left main coronary artery (LMCA) group (n = 16; solid bars), left anterior descending coronary artery (LAD) group (n = 46; hatched bars) and right coronary artery (RCA) group (n = 24; dotted bars). *p < 0.05. **p < 0.01.

Table 2. Results of Univariate and Multivariate Analyses for Distinguishing Between the LMCA and LAD Groups

<table>
<thead>
<tr>
<th>ST Segment Shift</th>
<th>LMCA Group</th>
<th>LAD Group</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Lead aVR (mV)</td>
<td>0.16 ± 0.13</td>
<td>0.04 ± 0.10</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Lead V1 (mV)</td>
<td>0.00 ± 0.21</td>
<td>0.14 ± 0.11</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Leads aVR–V1 (mV)</td>
<td>−0.16 ± 0.25</td>
<td>−0.09 ± 0.13</td>
<td>&lt;0.0001</td>
</tr>
</tbody>
</table>

Multivariate Analysis (Stepwise Linear Multiple Discriminant Analysis)

<table>
<thead>
<tr>
<th>Selected Factor</th>
<th>Discriminant Coefficient</th>
<th>Mahalanobis’ Sum of Squares</th>
<th>Partial F Value</th>
<th>p Value</th>
<th>F Value of Discriminant Equation</th>
</tr>
</thead>
<tbody>
<tr>
<td>Lead aVR</td>
<td>0.747</td>
<td>2.63</td>
<td>3.31</td>
<td>0.074</td>
<td>8.53 (p &lt; 0.01)</td>
</tr>
<tr>
<td>Lead V1</td>
<td>−0.817</td>
<td>2.34</td>
<td>5.7</td>
<td>0.02</td>
<td></td>
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</tbody>
</table>

Results of discrimination (percent correct): LMCA group: 63%; LAD group: 96%. Sensitivity = 63%, specificity = 96% and accuracy = 87%. Univariate data are presented as the mean value ± SD.

LAD = left anterior descending coronary artery; LMCA = left main coronary artery.
observed. In all patients studied, the acute MI was the first MI. Univariate and multivariate analyses showed essentially identical results. The selection of these patients and the methods of analysis were such that the ECG findings reflected the location of the myocardium perfused by each coronary artery, so they could be compared among the three groups.

**Lead aVR ST segment elevation.** The present study found lead aVR ST segment elevation in 43% (20/46) of the patients in the LAD group; this incidence is in good agreement with the incidence reported by Engelen et al. (4) in patients with acute LAD obstruction in which the culprit lesion was located proximal to the first major septal branch. These investigators concluded that lead aVR ST segment elevation in acute, proximal LAD occlusion is the result of transmural ischemia of the basal part of the septum, where the injury's electric current is directed toward the right shoulder. It is certainly reasonable to theorize that acute LMCA obstruction also causes ischemia of the basal part of the septum through disturbance of LCx blood flow—that is, interruption of the proximal LAD blood flow. This would account for lead aVR ST segment elevation in patients who do not have LAD obstruction, but not LAD obstruction, ordinarily results in ischemia of the posterior wall through disturbance of LCx blood flow. It is reasonable to assume that the electrical force in posterior wall ischemia counterbalances the ischemia-induced electrical force in the anterior wall. In fact, several reports have shown reciprocal changes in the precordial leads (V1 and V2) induced by posterior wall ischemia that was caused by LCx obstruction (9,10). The most likely interpretation of less ST segment elevation in lead V1 in the LMCA group compared with the LAD group is that it is the result of the electrical force induced by posterior wall ischemia, associated with LMCA obstruction counterbalancing the ischemia-induced electrical force in the anterior wall.

**Previous reports of acute LMCA obstruction.** A total of 42 patients with acute LMCA obstruction have been described in 21 reports published in English, according to our search results (1–3,11–29). The ECG findings were not described in four reports, representing 10 patients (1,13,16,26). ST segment changes consistent with anterior wall MI were reported in 25 of the remaining 32 patients. ST segment depression in the precordial leads was reported in seven patients and right bundle branch block in two patients. None of these reports described the findings of lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR. Pahlm et al. (30) investigated whether 35 ECG interpreters often ignore findings in lead aVR.
Study limitations. One of the limitations of our study is that it included a relatively small number of patients, because acute LMCA obstruction is not common. Careful patient selection and analysis by multiple methods (i.e., univariate and multivariate analyses), however, may have at least partly compensated for this limitation.

Conclusions. The present study showed that in patients with acute MI, careful attention to lead aVR ST segment elevation to predict acute LMCA obstruction is clinically important with respect to selection of the treatment strategy. The ST segment elevation in lead aVR also contributed to predicting patients’ clinical outcomes in acute LMCA obstruction.

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