Postural Changes in Left Ventricular and Mitral Valvular Dynamics in the Systolic Click – Late Systolic Murmure Syndrome

By Mary E. Fontana, M.D., Charles F. Wooley, M.D., Richard F. Leighton, M.D., and Richard P. Lewis, M.D.

SUMMARY
Marked changes in the auscultatory pattern with posture have been noted in patients with mid-systolic clicks and/or late systolic murmurs (MSC-LSM). MSC tend to move earlier in systole and LSM become longer and often louder when patients assume upright posture. Systolic prolapse of the mitral leaflets with mild regurgitation account for MSC-LSM; earlier and greater prolapse with more and prolonged regurgitation associated with a reduced left ventricular volume (LVV) in the upright position would explain the auscultatory changes.

Twenty-two patients with MSC-LSM were studied supine and at 45° head-up tilt, recording intracardiac pressures, cardiac outputs, systolic time intervals, and performing LV cineangiography. Systolic prolapse of one or both mitral leaflets was demonstrated in all patients. Left ventricular end-diastolic and end-systolic volumes both decreased significantly at 45° in all sixteen technically satisfactory studies. Greater mitral prolapse was noted upright in 12 of 14 studies with enough sinus beats to judge. The amount of mitral regurgitation was mild in all, and changes in amount from supine to upright posture could not be discerned angiographically. The findings suggest that the auscultatory changes occurring with upright posture in patients with MSC-LSM are related to greater prolapse of the mitral leaflets which is associated with a small LVV in the upright position.

Additional Indexing Words:
Mitral regurgitation
Mitral prolapse
Ventricular volume

Mid-systolic clicks and late systolic murmurs are associated with systolic prolapse of one or both mitral valve leaflets into the left atrium. Others have noted1-13 and we have previously emphasized4 the marked changes in the auscultatory pattern that occur with changes in posture. The systolic clicks tend to move toward and/or merge with the first sound; and the late systolic murmurs tend to become longer, often louder, and frequently holosystolic with the assumption of upright posture. We believe that the changes in auscultatory findings result from earlier and greater prolapse of the mitral leaflets, allowing longer and more pronounced mitral regurgitation, and that the primary determinant is the reduction in left ventricular volume that occurs in the upright position. Other authors have suggested this mechanism as well,1, 5-7, 10 but objective proof is lacking. This study was designed to document the alterations in left ventricular and mitral valvular dynamics occurring in patients with mid-late systolic murmurs and/or clicks in the upright position.

Materials and Methods
Twenty-two symptomatic patients with mid-systolic clicks and/or late systolic murmurs were studied. The changes in auscultatory findings with changes in posture were documented in each patient by phonocardiograms recorded in the supine, sitting, standing, and squatting positions. The patients were then placed on the table in the catheterization laboratory and tilted upright until definite click mobility toward the first sound and/or murmur prolongation were produced. It became apparent that 45° head-up tilt was sufficient to produce obvious changes in the auscultatory findings, so this degree of tilt was routinely used. Although the auscultatory changes were not so dramatic as those occurring when the patients were standing, they were judged sufficient for the purpose of this study. It would have been technically difficult and would have increased patient risk to perform these studies in the standing position.

The day following the phonocardiographic studies, right and left heart catheterization with left ventricular cineangiography and systolic time intervals were performed in the supine and 45° head-up tilt positions. A microphone

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The amount of prolapse was determined by outlining the projected image of the cineangiogram frames where the prolapse appeared maximal. Lines were drawn connecting the point of intersection of the leaflets with the ventricular image. This area was then planimetered.

was placed on the chest to record the second sound for the systolic time intervals and also the clicks and murmurs whenever possible. Systolic time intervals were measured by the method of Weissler et al. Cardiac outputs were determined by the indicator dilution technique. Cineangiography was performed in the 30° right anterior oblique (RAO) and 55° left anterior oblique (LAO) positions supine, and in the 30° RAO position, upright, using meglumine diatrizoate and sodium diatrizoate. Upright LAO injections were not done because the total amount of contrast medium required would have been excessive. All recordings were made on a Sanborn 550M recorder. Pressure measurements, cardiac outputs, and systolic time intervals were measured in the supine position first. Upright systolic time intervals, pressure measurements, and cardiac outputs were obtained beginning six minutes after the

Figure 1


assumption of 45° head-up tilt. Upright cineangiography was then performed. The patients were then returned to the supine position and the supine cineangiograms were performed a minimum of fifteen minutes after returning to 90° tilt and a minimum of twenty minutes after the upright cineangiogram. Ventricular volume determinations were made from the 30° RAO cineangiograms by the area-length method. The portion of the image of the prolapsed leaflet that projected beyond the ventricular image was excluded from the drawing of the end-systolic area. Only normal sinus beats were used for calculation. The amount of mitral leaflet prolapse was determined by outlining the projected image on a Tagarno projector screen of a frame where prolapse appeared maximal (fig. 1). A line was then drawn connecting the points of intersection of the prolapsed leaflet with the ventricular image. The enclosed area was then planimetered. The beats used for analysis were the same ones used for volume determinations.

Results

The precatheterization phonocardiograms demonstrated click mobility toward the first sound and/or murmur prolongation and/or intensification in all patients when they assumed the upright position (table 1). As can be seen in table 1, most of the patients developed pansystolic murmurs associated with merging of the click(s) with the first sound in the standing position. An example is shown in figure 2. Prompt squatting routinely reproduced the supine auscultatory findings. Upon standing from the squat-

Figure 2

A representative phonocardiogram from a patient in this series recorded the day prior to the catheterization. In the left panel, a supine recording demonstrates two mid-systolic clicks. The grade II/VI late systolic murmur was only faintly recorded. In the sitting position (second panel), the clicks are clearly earlier in systole and the murmur is nearly holosystolic and increased in intensity. In the standing position (third panel), the clicks have essentially merged with the first sound and the murmur is pansystolic. The heart rates for the three positions are 94, 110, 124, respectively. Prompt squatting (right panel) essentially reproduces the supine findings. S1 = first heart sound; SC = systolic click; S2 = second heart sound; SM = systolic murmur; 3LSB = 3rd intercostal space, left sternal border; F-400 = frequency cut-off 400 HZ.
### Table 1

**Postural Changes in MSC-LSM Syndrome**

<table>
<thead>
<tr>
<th>Patient</th>
<th>Prescatheterization phonation</th>
<th>Catheterization, 45° tilt</th>
<th>Δ Q-SC or SM on tilt†</th>
<th>Δ Heart rate</th>
<th>Comment</th>
</tr>
</thead>
<tbody>
<tr>
<td>RD</td>
<td>?MSC, II LSM (74)†</td>
<td>SC → S₁, IV HSM, Whoop (71)</td>
<td>—</td>
<td>77 → 68</td>
<td>Incomplete study, technically poor phon; symptomatic ↓ BP</td>
</tr>
<tr>
<td>MK</td>
<td>2 LSC, III LSM (92)</td>
<td>SC → S₁, IV HSM (95)</td>
<td>80 msec.</td>
<td>77 → 97</td>
<td>Complete study</td>
</tr>
<tr>
<td>EP</td>
<td>II LSM (81)</td>
<td>IV HSM (111)</td>
<td>120</td>
<td>72 → 91</td>
<td>Incomplete study, symptomatic ↓ BP</td>
</tr>
<tr>
<td>CVM</td>
<td>LSC, I-II LSM (71)</td>
<td>SC → S₁, II HSM (87)</td>
<td>110–140</td>
<td>71 → 83</td>
<td>Complete study, asymptomatic ↓ BP</td>
</tr>
<tr>
<td>RD</td>
<td>MSC, II LSM (94)</td>
<td>SC → S₁, IV HSM (124)</td>
<td>50</td>
<td>66 → 85</td>
<td>Complete study</td>
</tr>
<tr>
<td>AH</td>
<td>LSC, II LSM (79)</td>
<td>SC → S₁, II HSM, Whoop (113)</td>
<td>40</td>
<td>75 → 91</td>
<td>Complete study, asymptomatic ↓ BP</td>
</tr>
<tr>
<td>LB</td>
<td>MSC, I LSM (83)</td>
<td>SC → S₁, III HSM, Whoop (115)</td>
<td>140</td>
<td>92 → 109</td>
<td>Complete study</td>
</tr>
<tr>
<td>MC</td>
<td>LSC, II LSM (122)</td>
<td>SC → S₁, IV HSM, Whoop (143)</td>
<td>—</td>
<td>95 → N.D.</td>
<td>Incomplete study, asymptomatic ↓ BP</td>
</tr>
<tr>
<td>DP</td>
<td>II MSM (79)</td>
<td>III MSM (115)</td>
<td>—</td>
<td>63 → 70</td>
<td>Murmur too soft to record; asymptomatic ↓ BP</td>
</tr>
<tr>
<td>BF</td>
<td>MSc, II LSM (Insp. Only) (94)</td>
<td>SC → S₁, II LSM (125)</td>
<td>30–70</td>
<td>92 → 105</td>
<td>Complete study</td>
</tr>
<tr>
<td>LH</td>
<td>LSC, I LSM (98)</td>
<td>SC → S₁, IV HSM (122)</td>
<td>90</td>
<td>97 → 111</td>
<td>Complete study</td>
</tr>
<tr>
<td>JS</td>
<td>ESC, II LSM (79)</td>
<td>ESC, II MSM (109)</td>
<td>—</td>
<td>77 → 80</td>
<td>Incomplete study, asymptomatic ↓ BP</td>
</tr>
<tr>
<td>SM</td>
<td>II LSM (74)</td>
<td>III HSM (92)</td>
<td>—</td>
<td>66 → 81</td>
<td>Murmur too soft to record; asymptomatic ↓ BP</td>
</tr>
<tr>
<td>WW</td>
<td>I-II LSM (81)</td>
<td>III HSM (103)</td>
<td>85</td>
<td>73 → 77</td>
<td>Complete study</td>
</tr>
<tr>
<td>SS</td>
<td>I-II LSM (67)</td>
<td>II MSM (86)</td>
<td>—</td>
<td>61 → 72</td>
<td>Murmur too soft to record; Complete study</td>
</tr>
<tr>
<td>CP</td>
<td>II MSM (73)</td>
<td>III HSM (92)</td>
<td>—</td>
<td>59 → 74</td>
<td>Murmur too soft to record; Asymptomatic ↓ BP</td>
</tr>
<tr>
<td>ES</td>
<td>LSC (43)</td>
<td>LSC (49)†</td>
<td>65</td>
<td>36 → 41</td>
<td>Complete study</td>
</tr>
<tr>
<td>TEC</td>
<td>II LSM (72)</td>
<td>IV HSM (88)</td>
<td>HSM on Tilt</td>
<td>59 → 73</td>
<td>Complete study</td>
</tr>
<tr>
<td>RM</td>
<td>MSC, I-II LSM (85)</td>
<td>SC → S₁, III HSM (109)</td>
<td>120–170</td>
<td>83 → 95</td>
<td>Complete study</td>
</tr>
<tr>
<td>ST</td>
<td>II LSM (75)</td>
<td>III HSM (158)</td>
<td>—</td>
<td>71 → 92</td>
<td>Technically poor upright phon; Complete study</td>
</tr>
<tr>
<td>BJ</td>
<td>Normal (73)</td>
<td>ESC (122)</td>
<td>ESC upright only</td>
<td>77 → 98</td>
<td>Complete study</td>
</tr>
<tr>
<td>SR</td>
<td>Normal (95)</td>
<td>ESC (107)</td>
<td>ESC upright only</td>
<td>78 → 98</td>
<td>Complete study</td>
</tr>
</tbody>
</table>

*Shortening of Q-systolic click or murmur interval.
†Heart rate in parentheses.
‡Δ Q-SC = 40 msec.

Abbreviations: ESC = Early systolic click; MSC = Midsystolic click; LSC = Late systolic click; MSM = Midsystolic murmur; LSM = Late systolic murmur; S₁ = First heart sound; HSM = Holosystolic murmur; ND = Not done.
Figure 3

The phonocardiograms recorded during the catheterization of the same patient as in figure 2. On the left, supine, the systolic clicks are again shown, in a similar position in systole compared to the pre-cath phonocardiogram. The phonocardiogram recorded at 45° head-up tilt is shown on the right. The clicks are earlier in systole compared to supine. The murmur, although audible through a stethoscope, was not recordable. The heart rate on tilt is 85, compared to 124 on standing when the phonocardiogram in figure 2 was recorded.

Figure 4

The changes in cardiac index, stroke volume index, and heart rate are shown here. In each instance the dots on the left of the panel are the supine values and those on the right of each panel are the values at 45° head-up tilt. The majority of the supine values are normal. The patient with the high stroke volume index and slow heart rate had congenital complete heart block.
hypotension upon passive tilt which obviated complete upright studies. Asymptomatic reduction in systemic blood pressure occurred in five others, which tended to diminish the auscultatory changes. Some of the hemodynamic changes with upright tilt are shown in figure 4. The cardiac index was normal or elevated in all and decreased an average of 21%. Stroke volume decreased an average of 29%. The average heart rate increase was 12 beats/min, but several patients had an exaggerated heart rate response to upright tilt. These patients tended to have the most dramatic changes in auscultatory findings as well. A significant increase in the PEP/LVET ratio routinely occurred.

Left ventricular cineangiography demonstrated systolic prolapse of one or both mitral leaflets in all 22 patients. Eighteen had posterior leaflet prolapse only, and 4 had prolapse of both leaflets. The cineangiograms were technically satisfactory for volume determinations in 16 patients. The left ventricular end-diastolic volume varied from 48 to 100 ml/m² (normal = 70 ± 20 ml/m²) and decreased significantly (P < .001) at 45° head-up tilt, an average of 16.5% (fig. 5). The left ventricular end-systolic volume likewise decreased significantly (P < .005) in the upright position, an average of 18% (fig. 6).

The amount of mitral regurgitation was mild in all patients. Five patients had only systolic clicks at the time of study; no mitral regurgitation was seen in these patients although leaflet prolapse was evident. In two patients, the amount of mitral regurgitation was clearly greater in the upright position compared to supine. Differences in amount of regurgitation in the remainder of the cases could not be discerned primarily because the small amount of regurgitated contrast quickly dissipated into the nonopacified blood in the left atrium. No objective means of comparison sensitive enough to detect small differences in regurgitation is currently available. Our available timing devices were also inadequate to detect the precise timing of the onset of prolapse. Visual estimation of onset of prolapse was not reproducible by the observers with enough precision to give meaningful results. It has been shown by others, however, that the systolic click occurs at the point of maximum prolapse of the mitral leaflets, so the earlier clicks likely mean earlier prolapse. The amount of prolapse, determined by planimetry as previously described, was sig-

![Figure 5](image_url)

*The changes in LV end-diastolic volume occurring with upright tilt are shown. The supine values are represented by the black dots on the left and are connected to the upright values on the right for each patient.*

![Figure 6](image_url)

*The changes in LV end-systolic volume from supine, on the left, to upright, on the right, are shown. The net change in end-systolic volume was similar to that of the end-diastolic volume.*
phomomas are associated with significant increases in
the amounts of mitral valve prolapse which are
associated with significant reductions in left ven-
tricular end-diastolic and end-systolic volumes.

No data are available on the quantitation of the left
ventricular volume changes with upright tilt by the
angiographic method with which to compare our
results. Paley et al.\textsuperscript{21} noted approximately a 25% decrease in ventricular volumes by the thermodilution
technique at 60° tilt. Rushmer has also documented
the reduction of volume in the upright position in
dogs.\textsuperscript{17--18} The determination of left ventricular
volume by the area-length method from single plane
supine angiograms is a well established tech-
nique. There is no reason to believe that it would not
be equally appropriate for upright cineangiography
performed in the same projection. The twenty minute
interval between the upright and supine cineangiogra-
grams used for volume calculation has been shown by
others to be sufficient to allow the increase in plasma
volume induced by a contrast injection to dissi-
pate.\textsuperscript{30--33} The increased plasma volume produced by
the previous contrast injection could have exaggerated
the supine ventricular volume if the blood volume
change had not been taken into account. The changes
in ventricular volumes occurring with upright tilt in
the patients in this study are presumed to be normal
since the other hemodynamic and systolic time inter-
val changes agreed with previously obtained values in
normal individuals.\textsuperscript{17--20}

The greater prolapse demonstrated in the upright
position was reproducible in multiple beats in a given
patient, and by separate observations. The method
used quantitates only posterior leaflet prolapse, since
only 30° RAO cineangiograms were compared.
Greater prolapse in the upright position might have
been evident if left anterior oblique cineangiography
had also been performed in the upright tilt position,
since anterior leaflet prolapse is usually better seen in
this projection.

The possibility that the change in leaflet prolapse
occurring upright may have been related to a change
in the position and axis of the heart must be con-
sidered. The orientation appeared similar to supine
by visual inspection and it is doubtful that this is a signifi-
cant factor.

The demonstration of a reduction in left ventricular
volumes and an increase in the amount of mitral
prolapse does not necessarily mean that they are
related as cause and effect. That the volumes
decreased in every case in which the amount of
prolapse increased suggests that they are related. The
explanation for the greater prolapse of the mitral
leaflets in the upright position compared to the
supine, based on ventricular volume changes, is likely

![Figure 7](image-url)

\textit{The increases in area under the prolapsed leaflet from the supine
position, on the left, to the upright position, on the right, are shown.
The patient with the greatest change in the amount of prolapse had
an increase in heart rate of 40 beats/min when tilted to 45°.}
as follows: with a smaller end-diastolic volume, the supporting structures of the mitral valve, i.e., ventricular wall and papillary muscles, are physically closer to the mitral valve ring at end-diastole. There would be less tension on the chordae tendineae and the leaflets would, therefore, be closer to a position of

![Figure 8](image)

Figure 8

End-diastolic frames from 30° RAO cineangiograms done in the supine position, on the left and in the 45° head-up tilt position, on the right, from a patient in this series are shown. The end-diastolic volume (EDV) in the upright frame is clearly smaller (70 ml/m² vs 98 ml/m² in the supine position).

![Figure 9](image)

Figure 9

End-systolic cine frames with accompanying reference figures for the supine position, on the left, and the upright position, on the right, are shown. The characteristic appearance of contrast underlying the posterior leaflet is apparent. The end-systolic volume is significantly smaller in the upright study (27 ml/m² vs 34 ml/m² in supine position). The area under the prolapsed leaflet (AP) is larger in the upright cine (19.6 cm² vs 18.6 cm² in supine position).

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prolapse. With ventricular systole, which results in a smaller end-systolic volume, the leaflets then prolapse more and likely earlier, which would permit more prolonged and greater regurgitation. Considering the relative inelasticity of the mitral leaflets and chordae tendineae, changes in the relationship of the supporting structures of the valve, i.e., papillary muscles and ventricular wall, to the point of attachment of the leaflets (i.e., valve ring) would logically explain the alterations in mitral valve function seen in patients with mitral valve prolapse. Changes in ventricular volumes would clearly be the major determinant of this relationship.

Standing up resulted in a marked tachycardia in many of these patients (table 1). This was associated with the development of pansystolic murmurs in the majority of them (table 1). The greater changes in auscultatory findings with the greater increase in heart rate would be explained by the further reduction in ventricular volume because of marked reduction in the diastolic filling period, which would be additive to the peripheral pooling that occurs on assuming upright posture. The inotropic stimulus of the increased sympathetic activity that occurs with upright tilt, 25,26 which is likely greater in the standing position, and with increases in heart rate, may contribute to the reduction in both end-systolic and end-diastolic volumes seen in this study.

The major determinant of the greater prolapse seen in this study appears to be ventricular volume reduction resulting primarily from peripheral pooling. In four patients the heart rate during the filming of the cineangiograms was the same or slower during the upright injection as during the supine injection, presumably related to the deep, held inspiration. The ventricular volume still decreased and the amount of mitral valve prolapse did increase in these patients. The changes in volume and prolapse were greater in those with increases in heart rate, although not statistically significant, consistent with the additive effects of heart rate on ventricular volume reduction as previously discussed.

Further support for the volume theory can be deduced from the response to prompt squatting. Prompt squatting routinely results in movement of the clicks and murmurs back to a later systolic position, similar to supine (fig. 2). Prompt squatting increases venous return and slows the heart rate, which results in lengthening of the diastolic filling period. 27,28 An increase in ventricular volume would then result, 18 which would produce more tension on the mitral valve leaflets and chordae by increasing the distance between the ventricular wall and papillary muscles and the valve ring. Later and less leaflet prolapse would then result during systole.

In conclusion, the movement of the systolic clicks toward the first sound and the murmur prolongation and intensification that occur in the upright position in patients with mid-systolic clicks and late systolic murmurs are associated with greater prolapse of the mitral leaflets, which could allow greater and more prolonged regurgitation. A primary determinant of the greater prolapse appears to be the reduction in left ventricular volume that occurs in the upright position.

References

POSTURAL CHANGES IN MSC-LSM SYNDROME


Postural changes in left ventricular and mitral valvular dynamics in the systolic click - late systolic murmur syndrome.
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