

# Assessment of Left Ventricular Outflow in Hypertrophic Cardiomyopathy Using Anyplane and Paraplane Analysis of Three-Dimensional Echocardiography

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This study analyzes the alterations in size and geometry of the left ventricular (LV) outflow tract that occur in hypertrophic cardiomyopathy (HC) using transthoracic 3-dimensional echocardiography. Transthoracic 3-dimensional echocardiography was performed in 17 patients with HC (4 after myectomy) and in 10 normal subjects. Images were acquired with the rotational approach, with electrocardiographic and respiratory gating. From the 3-dimensional datasets, short-axis parallel slicing of the LV outflow tract at a 1 mm distance was performed at the onset of systole. For each slice, cross-sectional area and maximal and minimal diameter were calculated. Reconstruction of the LV outflow tract could be displayed in 3 dimensions in all patients, allowing orientation and clear definition of the irregular geometry. In patients with HC, the minimal LV outflow tract cross-sectional area was smaller than in normal subjects ( $2.3$

$\pm 1.0$  vs  $5.0 \pm 0.9$  cm<sup>2</sup>,  $p < 0.0001$ ). The ratio between maximal and minimal cross-sectional areas was higher in patients with HC than in normal subjects ( $2.6 \pm 0.9$  vs  $1.4 \pm 0.2$ ,  $p < 0.0001$ ). The ratio between maximal and minimal diameter of the smallest cross section of the LV outflow tract was also significantly higher in patients with HC than in normal subjects ( $1.6 \pm 0.3$  vs  $1.2 \pm 0.1$ ,  $p < 0.001$ ); a value of 1.36 separated normal subjects from HC patients without previous myectomy. In conclusion, precordial 3-dimensional echocardiography allows detailed qualitative and quantitative information on the LV outflow tract. Patients with HC are characterized by a highly eccentric and asymmetric shape of the LV outflow tract, and by a smaller minimal cross-sectional area than that seen in normal subjects.

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Postmortem studies and intraoperative findings indicate that 2-dimensional echocardiography may fail to give the full picture of the left ventricular (LV) outflow tract in patients with hypertrophic cardiomyopathy (HC).<sup>1-6</sup> Three-dimensional echocardiography is a new imaging modality which provides unique information on the spatial geometry of a given structure.<sup>7</sup> Based on our experience in an unselected population,<sup>8,9</sup> we believe that the alterations in size and geometry of the LV outflow tract in patients with HC could be more accurately analyzed with precordial 3-dimensional echocardiography. With these concepts in mind, we designed this study to gain further insight into the complex geometry of the LV outflow tract in patients with HC. With this aim, analysis of the LV outflow tract was performed using 3-dimensional datasets obtained after acquisition of precordial echocardiographic images.

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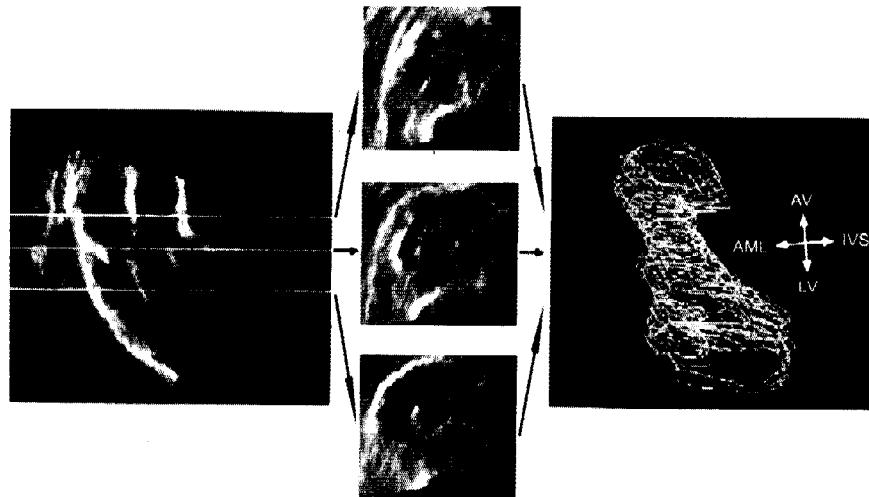
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## METHODS

**Study patients:** We prospectively selected 17 patients (13 men and 4 women; mean  $\pm$  SD age  $39 \pm 15$  years, range 19 to 65) with HC referred to the outpatient clinic of our institution for routine transthoracic echocardiographic follow-up. High-quality images were a prerequisite for inclusion in this study. The diagnosis of HC was based on M-mode and 2-dimensional echocardiographic demonstration of a nondilated hypertrophic left ventricle in the absence of other cardiac or systemic disease that could produce LV hypertrophy.<sup>10</sup> According to a previously established classification,<sup>11</sup> the patterns of distribution of LV hypertrophy were: type I, 1 patient; type II, 3 patients; type III, 12 patients; and type IV, 1 patient. Systolic anterior motion of the mitral valve was present in 12 patients and its severity was evaluated semiquantitatively from 0 = absence, to 3+ = contact with the interventricular septum during systole.<sup>5</sup> At the time of the echocardiographic study, a pressure difference was calculated from Doppler LV outflow tract velocity recordings, and obstruction (gradient  $> 30$  mm Hg under basal conditions) was detected in 4 patients. A septal myectomy had been performed in 4 patients. Ten asymptomatic subjects without evidence of LV hypertrophy were also studied for comparison. These controls were aged 20 to 49 years (mean  $28 \pm 8$ ) and 8 were men.

**Examination procedure:** Two-dimensional echocardiographic studies were performed with a commer-

**FIGURE 1.** Analysis of left ventricular (LV) outflow tract in a patient with hypertrophic cardiomyopathy. From the 3-dimensional dataset, parallel slicing of the LV outflow tract perpendicular to the long axis (paraplane echocardiography) is performed at 1 mm intervals, at the onset of ventricular systole. In the *left panel*, 3 representative cut-planes are indicated. The corresponding cross-sectional 2-dimensional images with the manually traced endocardial contours are shown in the *middle panels*. The final wire frame mode display represented in the *right panel* can be rotated on the screen along the 3 main axes for versatile qualitative 3-dimensional evaluation. AML = anterior mitral leaflet; AV = aortic valve; IVS = interventricular septum;



cially available system (Vingmed CFM 750 [Horten, Norway] or Toshiba Sonolayer SSH-140A [Tokyo, Japan]) equipped with a 3.5 MHz transducer, while the patient was lying in the 45° left recumbent position. After follow-up 2-dimensional echocardiographic study, the probe was positioned either at the left parasternal or apical window for acquisition of tomographic images of the LV outflow tract for 3-dimensional reconstruction. Patient movement during the image acquisition can be prevented by thoroughly explaining the procedure before the study. The operator has to find the central axis of rotation so that the conical datasets encompass the LV outflow tract. During the acquisition, movements of the transducer holder must be avoided diligently. Mirror images of the first and the final cut-plane indicate a correct 180° rotation. All subjects gave informed consent.

**Three-dimensional echocardiography:** IMAGE ACQUISITION: After the standard 2-dimensional examination, the video output of the echocardiographic system is interfaced to the acquisition system (Echo-scan, TomTec GmbH, Munich, Germany) and the transducer is fixed into a cylindrical holder. A step-motor mounted on this holder can rotate the probe around its longitudinal axis via a wheel-work interface. The step-motor is connected to the acquisition system which commands the rotation of the probe at 2° intervals over a span of 180°. Respiratory and electrocardiographic gating are performed after the operator has selected the end-expiratory phase by thoracic impedance measurement and an adequate RR interval. Thus, only beats falling in the predetermined RR interval and at the end-expiratory phase are selected by the steering logic of the system and acquired. Cycles that do not meet the preset ranges are rejected. Ninety sequential cross-sections are acquired, each during a complete heart cycle, encompassing a 3-dimensional conical volume. After distance calibration, images are stored in the computer memory for subsequent analysis.

**IMAGE PROCESSING:** The raw data are resampled off-line according to their temporal and spatial lo-

cation. The coordinates of each point are converted from a polar to a rectangular system, and the space between contiguous points is electronically filled with a trilinear cylindrical interpolation. Several algorithms are used to reduce noise and artifacts which can be created by patient and probe movements.

**IMAGE DISPLAY ANALYSIS:** Images were analyzed as follows:

1. After thorough examination of the 3-dimensional datasets with analysis of cardiac cross-sections in any desired plane ("anyplane echocardiography"), a cut-plane is selected where the LV outflow tract is visualized along its longitudinal axis.
2. The onset of ventricular systole is selected, as the first frame during the cardiac cycle in which the mitral valve appears closed.
3. Electronic parallel slicing ("paraplane echocardiography") of the LV outflow tract perpendicular to the vertical axis is performed at 1 mm intervals, from the hinge point of the anterior mitral leaflet to the point of coaptation of the mitral leaflets, and the corresponding 2-dimensional images (both the stop frame at onset of systole and the dynamic sequence) are displayed.
4. On the stop frame image at the onset of systole, the endocardial contour of the cross-section is manually traced for automatic area measurement, and maximal and minimal diameters are also measured.
5. Finally, reconstruction of the LV outflow tract at the onset of ventricular systole is displayed in wire frame mode and the representative image can be rotated on the screen for versatile 3-dimensional evaluation (Figure 1).

From analysis of the 3-dimensional datasets, the following measurements of the LV outflow tract at the onset of systole were considered: (1) minimal cross-sectional area; (2) ratio between maximal and minimal cross-sectional area (max/min cross-sectional area), as an index of the "eccentricity" of the LV outflow tract (the higher the value, the greater the variations of the cross-sectional area throughout the length of the LV outflow tract); (3) ratio between maximal (lateromedial) and minimal (anteroposter-

**TABLE I** Demographics, Patients Characteristics, and Measurements of the Left Ventricular Outflow Tract in Patients With Hypertrophic Cardiomyopathy

Patient Number	Age (yr) & Sex	Type LVH	Therapy	Gradient	SAM	CSA (cm <sup>2</sup> )	Max./Min CSA	Max./Min Diameter
1	19M	III	V	58	3+	2.9	2.0	1.9
2	21M	III	—	<10	2+	2.4	2.5	1.7
3	22M	I	—	<10	1+	3.0	1.8	1.4
4	24M	IV	—	<10	2+	2.9	2.4	1.6
5	25M	III	V	<10	2+	2.7	2.4	1.6
6	31F	III	S, V	50	3+	1.3	4.2	2.0
7	38F	II	V	<10	0	2.4	2.7	1.7
8	39F	III	V	<10	0	3.2	2.1	1.6
9	41M	III	V	20	1+	1.9	1.7	1.8
10	46F	II	A, V	<10	2+	0.7	4.1	1.4
11	53M	III	—	<10	1+	1.5	3.8	1.9
12	63M	II	M	100	3+	1.5	3.0	2.6
13	65M	III	V	35	3+	2.0	3.0	1.6
Mean ± SD	37 ± 15					2.2 ± 0.7	2.7 ± 0.8	1.75 ± 0.3
14	31M*	III	—	<10	0	2.1	2.4	1.3
15	34M*	III	S	<10	1+	0.9	3.9	1.2
16	44M*	III	S, V	<10	0	4.7	1.5	1.3
17	59M*	III	V	<10	0	3.0	1.6	1.3
Mean ± SD	42 ± 12					2.7 ± 1.6	2.3 ± 1.1	1.27 ± 0.05
Mean ± SD (overall)	39 ± 15					2.3 ± 1.0	2.6 ± 0.9	1.6 ± 0.3

\* Patients evaluated after myectomy.

A = amiodarone; CSA = cross-sectional area; LVH = left ventricular hypertrophy; M = metoprolol; Max./Min. = maximal/minimal; S = sotalol; SAM = systolic anterior movement; V = verapamil.

ior) diameter (max/min diameter) at the level of the cross-section with the minimal area, as an index of the "asymmetry" of the LV outflow tract (a ratio of 1 indicates a circular shape, with the highest values corresponding to the more elliptical shape of the cross sections).

Inter- and intraobserver reproducibility for the measurements of LV outflow tract with 3-dimensional echocardiography was assessed in all 27 patients. To assess interobserver variability, two observers (A.S. and Y.N.) independently measured the outflow tract area from the 3-dimensional datasets without prior knowledge of clinical data and without preselection of cut-planes. In addition, LV outflow tract cross-sectional area measurements were performed by 1 observer (A.S.) on 2 occasions (3 months apart, without preselection of cut-planes from the 3-dimensional datasets) to assess intraobserver variability.

**Statistical analysis:** Values are given as mean ± SD. Student's unpaired *t* test was used to compare the differences between HC and control subjects. Values of *p* < 0.05 were considered to be significant. Reproducibility of the LV outflow tract measurements was expressed in terms of mean differences and 95% confidence intervals.<sup>12</sup>

## RESULTS

Three-dimensional acquisition could be performed successfully in all patients. Echocardiographic acquisition of the image of the LV outflow tract for 3-dimensional reconstruction was performed either from the parasternal (*n* = 20) or apical (*n* = 7) windows, according to the image quality. The examination including the calibration procedure, selection of the optimal axis of rotation, a num-

ber of test runs, and the actual image acquisition required approximately 10 minutes in addition to the time required for the standard 2-dimensional echocardiogram. Three-dimensional reconstruction of the images was possible and of good quality in all patients. The time required for post-processing the raw data, and reconstruction and analysis of the images was approximately 20 minutes. Demographics, echocardiographic characteristics, and measurements of the LV outflow tract in each patient with HC are reported in Table I.

**Left ventricular outflow tract in patients with hypertrophic cardiomyopathy versus normal subjects:** CROSS-SECTIONAL AREA: The values of minimal cross-sectional area of the individual subjects are plotted in Figure 2. The minimal LV outflow tract cross-sectional area calculated with 3-dimensional echocardiography was significantly smaller in patients with HC than in the control group ( $2.3 \pm 1.0$  vs  $5.0 \pm 0.9$  cm<sup>2</sup>, *p* < 0.0001). Thirteen of the 17 patients with HC had a value smaller than controls, and 2 of the 4 HC patients (nos. 6 and 17) with higher values were evaluated after myectomy. After correction for body surface area the values were  $1.3 \pm 0.5$  and  $2.7 \pm 0.6$  cm<sup>2</sup>, respectively (*p* < 0.0001).

SHAPE: From the 3-dimensional datasets, the reconstructed LV outflow tract could be displayed as observed from different viewpoints. This simplifies visualization of the geometry and shape as well as the localization of the narrowing of the LV outflow tract in patients with HC (Figure 3). A similar display in a normal patient is shown in Figure 4. Some examples of 3-dimensional reconstruction of the LV outflow tract in normal subjects and in patients with HC are shown in Figure 5.

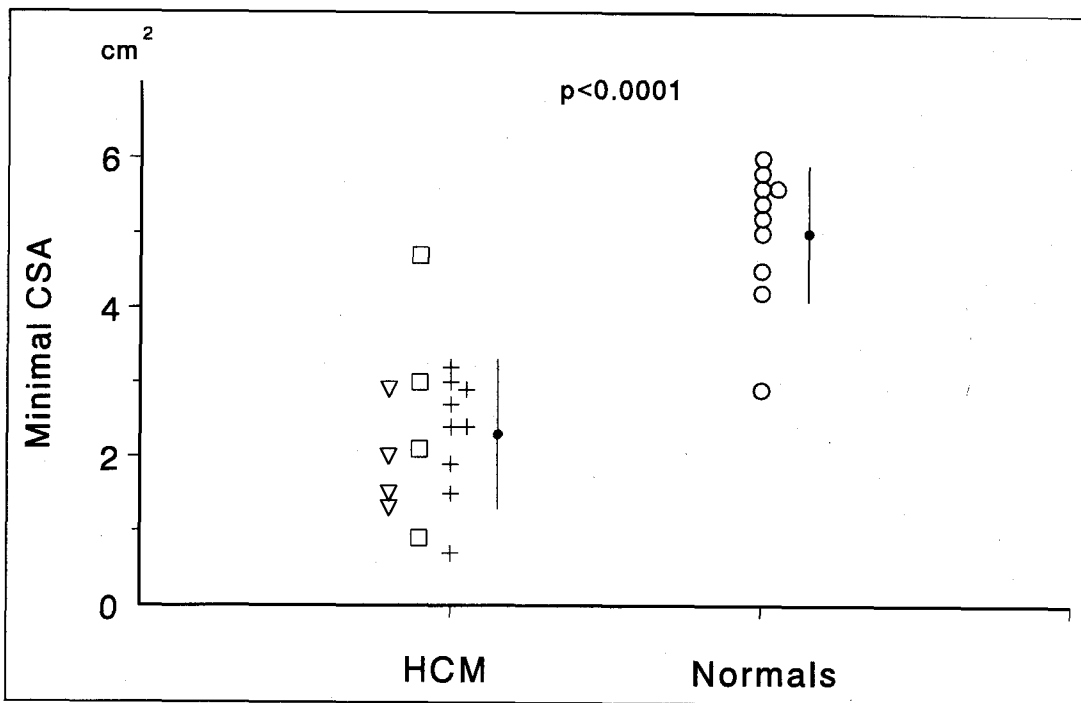


FIGURE 2. Minimal cross-sectional area (CSA) of the left ventricular outflow tract in patients with hypertrophic cardiomyopathy (HCM) and in normal subjects (○). + = patients without obstruction; □ = patients after myectomy; ▽ = patients with obstruction.

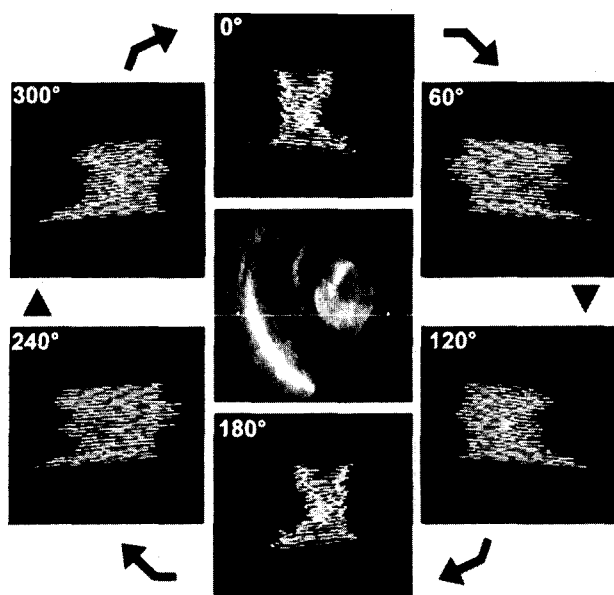


FIGURE 3. In this patients with hypertrophic cardiomyopathy, the wire frame display of the reconstructed left ventricular (LV) outflow tract is represented as observed from different viewpoints. The view at 0° corresponds to the cut-plane represented in the central panel. The images obtained after incremental 60° clockwise rotation are displayed in the corresponding panels. There is an eccentric and asymmetric shape of the LV outflow tract. From these images it is also apparent that the narrowing is localized at the middle-caudal part of the LV outflow tract and is mainly related to a reduction in the anteroposterior diameter.

MAX/MIN CROSS-SECTIONAL AREA (ECCENTRICITY INDEX): The max/min cross-sectional area of the LV outflow tract derived from the 3-dimensional datasets are displayed in Figure 6. From this figure, it is

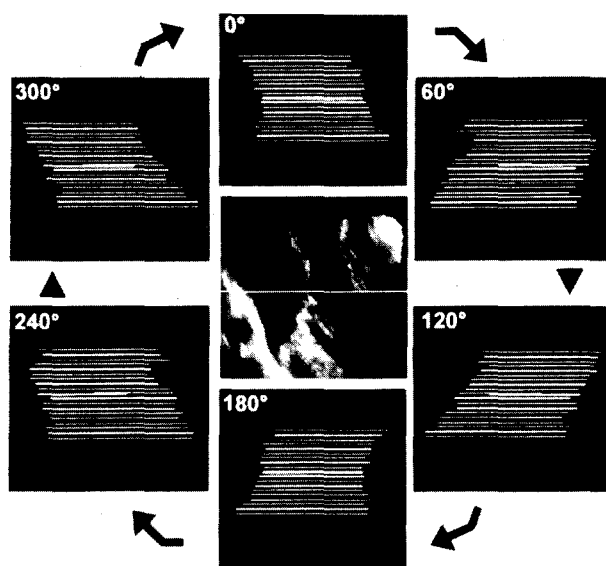
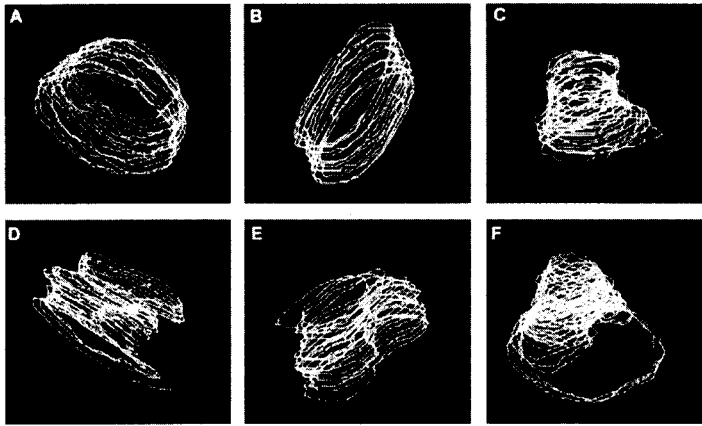


FIGURE 4. Three-dimensional reconstruction of the left ventricular (LV) outflow tract in a normal subject, with the same display as in Figure 3. Note the uniformity (indicated by the minimal variations of the diameters throughout the length of the LV outflow tract) as well as the symmetry (indicated by the similar diameters of individual cross-sections from different viewpoints) of the LV outflow tract.

apparent that patients with HC had higher ratios ( $2.6 \pm 0.9$ ), with a broad range of values (from 1.5 to 4.2) indicative of many irregular different shapes of the LV outflow tracts. In contrast, normal subjects had smaller ratios ( $1.4 \pm 0.2$ ), with a narrow range of values (from 1.1 to 1.6). In particular, each of the controls had a ratio of  $\leq 1.6$ , whereas 15 of 17 patients with HC had a ratio of  $> 1.6$ . The 2 patients



**FIGURE 5.** Three-dimensional reconstruction of the left ventricular outflow tract with wire frame display in normal subjects (A to C) and in patients with hypertrophic cardiomyopathy (D to F). The different irregular configuration of the left ventricular outflow tract in patients with hypertrophic cardiomyopathy can be evaluated.

with max/min cross-sectional area of  $\leq 1.6$  (patients 16 and 17) were evaluated after myectomy. Thus, an outflow tract area ratio of 1.6 appeared to separate patients with from subjects without HC.

**MAX/MIN DIAMETER (ASYMMETRY INDEX):** The individual values of max/min diameter of the LV outflow tract cross-section are displayed in Figure 7. This ratio was significantly higher in patients with HC than in normal subjects ( $1.6 \pm 0.3$  vs  $1.2 \pm 0.1$ ,  $p = 0.001$ ). Of interest, patients evaluated after myectomy had the lowest values; conversely, the highest values were found in patients with HC and LV outflow tract obstruction. An index of 1.36 appeared to separate

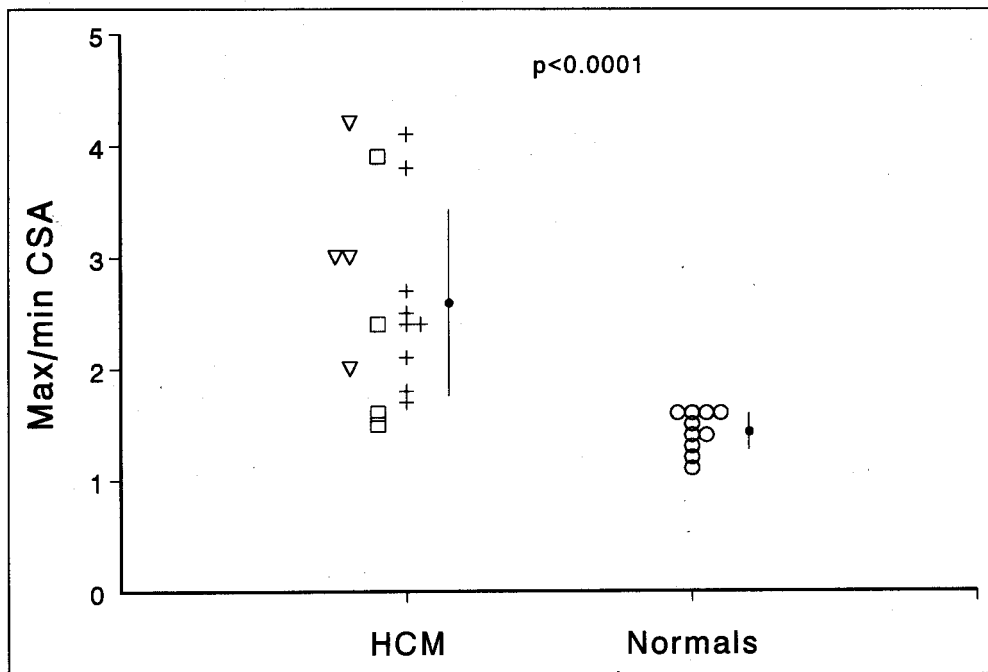
normal subjects from patients with HC who did not undergo operation.

**Reproducibility analysis of three-dimensional echocardiography:** **INTEROBSERVER VARIABILITY:** The difference between the 2 observers for measurements of the LV outflow tract was compared with the average of the 2 measurements for each patient. The mean difference between the measurements of the 2 observers was  $0.04 \text{ cm}^2$  (95% confidence interval  $-0.08$  to  $0.12 \text{ cm}^2$ ) for cross-sectional area.

**INTRAOBSERVER VARIABILITY:** The difference between the 2 measurements made by the same observer was compared with the average of the 2 measurements for each patient. The mean difference between the 2 measurements was  $0.13 \text{ cm}^2$  (95% confidence interval  $-0.01$  to  $0.25 \text{ cm}^2$ ) for cross-sectional area.

## DISCUSSION

Hypertrophic cardiomyopathy is a disease with a great individual variability and "no two hearts are alike."<sup>13</sup> The results of the present study indicate that 3-dimensional echocardiography allows visualization of the varied complex geometry of the LV outflow tract in patients with HC. With quantitative analysis of the 3-dimensional datasets, we demonstrated that in patients with HC the minimal cross-sectional area is smaller than that in normal subjects. In addition, most patients with HC have an irregular shape of the LV outflow tract as demonstrated by an eccentricity index of  $\geq 1.5$ . In normal subjects, this



**FIGURE 6.** Ratio between maximal and minimal cross-sectional area (max/min CSA) of the left ventricular outflow tract in patients with hypertrophic cardiomyopathy (HCM) and in normal subjects. A high ratio indicates an eccentric shape of the left ventricular outflow tract. Symbols as in Figure 2.

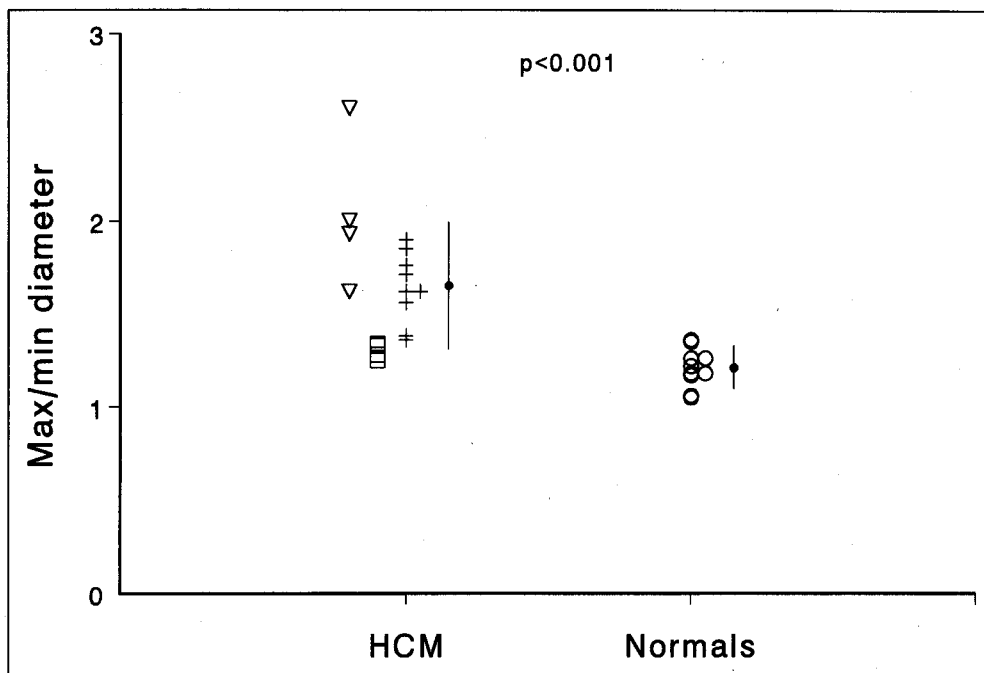


FIGURE 7. Ratio between maximal and minimal diameter (max/min diameter) measured at the cross-section of the left ventricular outflow tract with the minimal area in patients with hypertrophic cardiomyopathy (HCM) and in normal subjects. A high ratio indicates an asymmetric shape of the left ventricular outflow tract. Symbols as in Figure 2.

index is always  $\leq 1.6$ , indicating a uniform shape of the LV outflow tract, without significant variation of the cross-sectional area throughout its length. We have also demonstrated that the cross-sectional shape of the LV outflow tract is more elliptical in patients with HC than in normal subjects, as indicated by a higher ratio of max/min diameter measured at the plane of the minimal cross-sectional area. This finding is in agreement with the concept that in HC the hypertrophic ventricular septum narrows the LV outflow tract mainly along its anteroposterior diameter. Of interest, this asymmetry index was highest in patients with HC and obstruction of the LV outflow tract at rest. In contrast, in HC neither the minimal cross-sectional area nor the eccentricity index of the LV outflow tract separated this subgroup. This finding indicates that for similar cross-sectional area the asymmetry of the LV outflow tract plays an important role in determining the presence of significant obstruction at rest.

Patients who had undergone myectomy had a minimal cross-sectional area similar to other patients with HC, including those with obstruction (Figure 2). However, from Figure 7 it is clear that after myectomy the asymmetry index was lowest, indicating that the surgical remodeling of the LV outflow tract was adequate for the relief of the obstruction despite the finding that the cross-sectional area remained small compared with that in normal controls, and remained in the same range as that of the other HC patients without previous myectomy. Thus, in patients with HC, precordial 3-dimensional echocardiography has the potential to play a major role in tailoring the standard surgical resection of the interventricular septum to the individual patient's anat-

omy, which is crucial for safe and efficacious performance of myectomy, and also for evaluation of the results of surgery.

**Evaluation of left ventricular outflow tract using three-dimensional echocardiography:** Previous experience with 3-dimensional echocardiography for the evaluation of the LV outflow tract was based on morphologic analysis with volume-rendered display.<sup>14</sup> In contrast, for quantitative analysis of the LV outflow tract performed in the present study, we selected a multitude of cut-planes ("anyplane echocardiography") and performed a parallel scanning of the LV outflow tract at 1 mm intervals ("paraplane echocardiography"). This rate of sampling of the dataset is similar to the analysis done with magnetic resonance imaging or computed tomography, and allows detailed spatial information.<sup>15</sup> While some display modalities, such as volume rendering, are indicated more for representation of anatomic details,<sup>16-18</sup> the wire frame display format appears particularly suited to 3-dimensional reconstruction of the cardiac cavities, where areas, volumes, size, and shape can be adequately evaluated.

**Limitations of three-dimensional echocardiography:** In this study, patients were selected on the basis of high-quality images at 2-dimensional echocardiography, which yielded a success rate of 3-dimensional reconstruction of 100%. The same results cannot be expected from an unselected population where poor quality precordial images may prevent adequate quality of the reconstruction.

Echocardiographic images were acquired by an experienced technician and reconstruction was performed by a cardiologist after a learning period of >50 studies. This previous experience of 3-dimen-

sional echocardiography prevented artifacts in the 3-dimensional datasets, limited the time required both for acquisition and reconstruction, and resulted in a minimal variability for the measurements.

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