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Non-surgical Septal Reduction Procedure in Down's Syndrome with Hypertrophic Obstructive Cardiomyopathy - A Case Report

By Shanthi Sivanandam, MD and James H. Moller, MD

Keywords: Hypertrophic Cardiomyopathy, Alcohol septal reduction, Down's syndrome
Running title: Alternative treatment for Hypertrophic Obstructive Cardiomyopathy

The authors have no potential conflicts of interest, real or perceived, to report with regards to this work.

Introduction

We report our experience of Non-surgical septal reduction procedure in Down's Syndrome with Hypertrophic Obstructive Cardiomyopathy.

A 21-year old male, with Down's Syndrome, mental retardation had been followed for a number of years for a small ventricular septal defect. When last seen five years ago he was asymptomatic. Echocardiogram obtained in 1997 demonstrated a small pressure restrictive perimembranous ventricular septal defect. Interventricular septal thickness (IVS) was 7mm; left ventricular posterior wall (LVPW) was 7mm. In 2003 he was noted to have shortness of breath, diaphoresis with activity, decreased exercise tolerance and was

becoming more sedentary with New York Heart Association (NYHA) Class III symptoms.

Physical examination showed typical features of Down's Syndrome. Cardiovascular examination revealed a grade 2/6 high-pitched midsystolic murmur best heard along the lower left sternal border. Electrocardiogram showed left ventricular hypertrophy with strain (Figure 1). Echocardiogram showed marked asymmetric septal hypertrophy measuring 25mm; Left ventricular posterior wall measuring 15mm, consistent with hypertrophic cardiomyopathy (HCM) (Figure 2). Continuous wave Doppler estimated a 70mm Hg sub-aortic gradient due to dynamic systolic anterior motion of the mitral valve with septal contact. Thus, the diagnosis hypertrophic obstructive cardiomyopathy was made. Medical management with beta blockers was ineffective in controlling symptoms. There was no change in his exercise capacity and he continued to be in NYHA Class III. Due to progressively worsening symptoms, Non-surgical septal reduction procedure (NSRP) was considered for this patient to reduce outflow obstructive symptoms and was successfully performed.

Methods and Result

A non-surgical septal reduction procedure with intracoronary (septal) ethanol injection to



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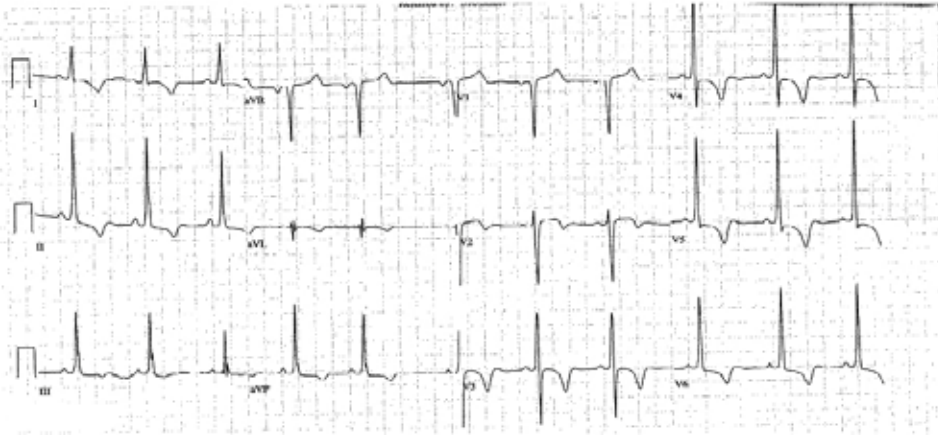


Figure 1: Pre-procedure ECG showing left ventricular hypertrophy with strain.

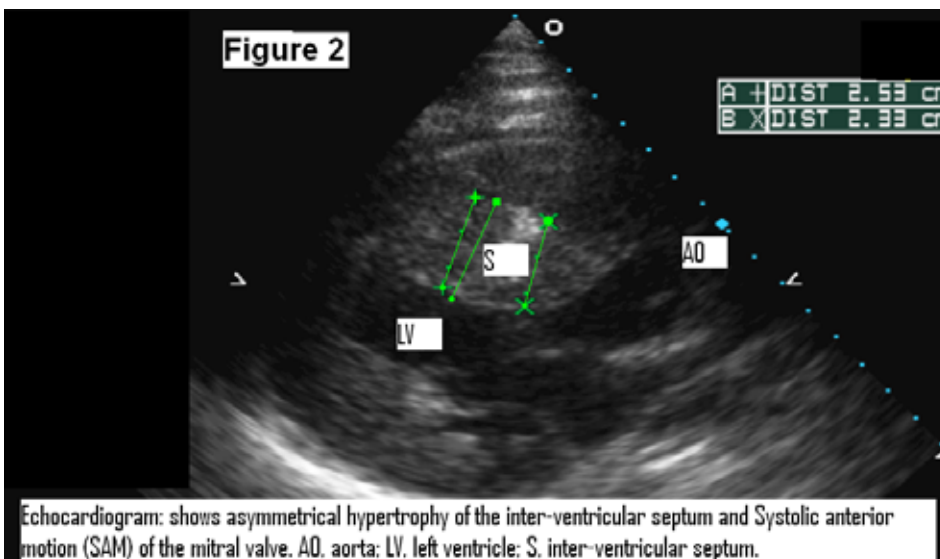


Figure 2: Echocardiogram shows asymmetrical hypertrophy of the inter-ventricular septum and Systolic anterior motion (SAM) of the mitral valve.

reduce the left ventricular obstruction was performed. Prior to NSRP he underwent a prophylactic pacemaker and defibrillator because of the high incidence of sudden death induced by ventricular tachycardia and ventricular fibrillation in hypertrophic obstructive cardiomyopathy. In brief, a high fidelity Radi pressure wire was placed into the LV through a 6 Fr catheter. The

catheter was withdrawn to the aorta and simultaneous pressures were continuously recorded in the left ventricle and aorta. Coronary angiography was performed to exclude atherosclerotic coronary arterial disease and identify the septal perforators. A 2.0/15 Crosssail balloon catheter was introduced into the first septal branch of the left anterior descending coronary artery.

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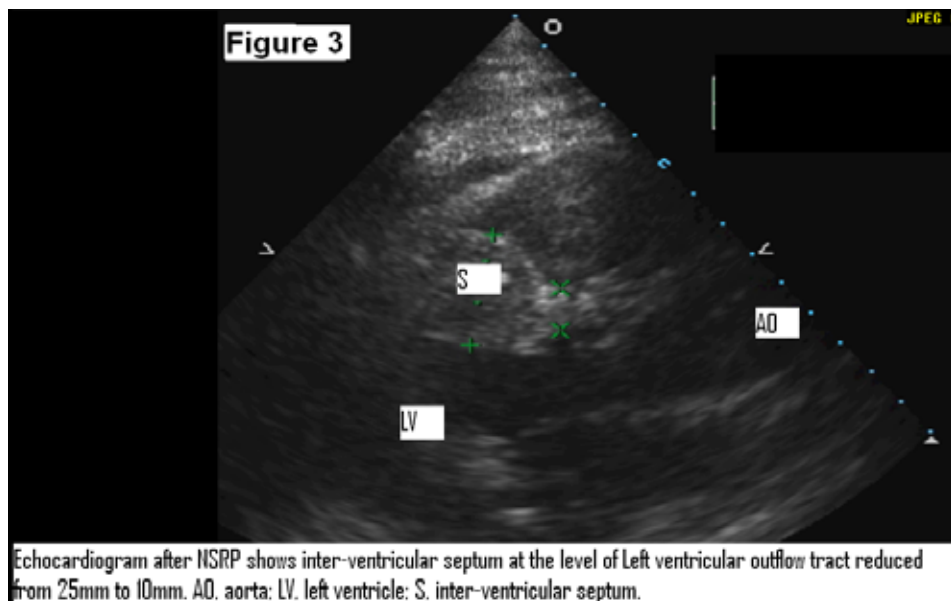


Figure 3: Echocardiogram after NSRP shows inter-ventricular septum at the level of left ventricular outflow tract reduced from 25mm to 10mm.

	Echo gradient	LVSP(cath)	Aorta(cath)	Cath gradient
2003	70mm Hg			
Resting 2004(cath)		142mm Hg	72mm Hg	70 mm Hg
1 st injection (cath)				60 mm Hg
2 nd injection (cath)		99mm Hg	83mm Hg	16 mm Hg
2 nd injection (cath)				
1 month later	4mm Hg			
1 year later	No gradient			
4 year later	No gradient			
Table: shows LVOT gradient pre and post Nonsurgical septal reduction procedure				

After the balloon was inflated, the distribution of the first septal branch was verified by contrast two-dimensional echocardiography after the injection of 2cc of myocardial echo contrast Definity. After confirming which territory of the basal septum contributed to the LVOT obstruction and no other myocardial territory was involved,

a total 1.6 ml of ethanol was injected slowly. The balloon occlusion was maintained for 10 minutes. There was an immediate reduction in LVOT peak systolic gradient to 20mm Hg. After balloon deflation the gradient slowly rose to 60mm Hg. The target septal perforator was again identified and an additional 1.6ml of



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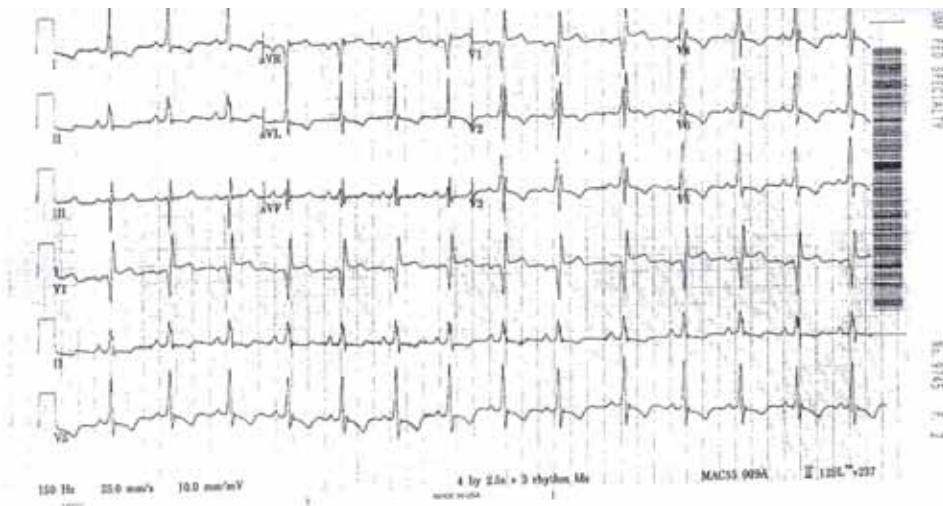


Figure 4: Post procedure ECG showing left ventricular hypertrophy with strain and right bundle branch block.

ethanol was injected through the inflated balloon resulting in 16mm Hg LVOT systolic peak gradient. The LVOT pressure gradients were measured before and immediately after the procedures by catheter and by two-dimensional echocardiography. LVOT catheterization gradient reduced from 70mm Hg to 16mm Hg. His hospital course following the procedure was uneventful, and he was discharged home in less than 2 days. One month after the NSRP echocardiogram revealed a mean gradient across the LVOT of 4mm Hg. One and four year follow-up echocardiograms revealed no LVOT gradient (Table 1). Interventricular septum measured 25mm prior to NSRP. One and four year follow-up revealed IVS measuring 10mm at the level of the left ventricular outflow tract (Figure 3).

Patient remains asymptomatic with no shortness of breath and improved exercise tolerance. He is NYHA class I with no limitation of activities. Since the placement of prophylactic pacemaker and defibrillator, there was no arrhythmia, no heart block, no ventricular tachycardia or ventricular fibrillation. Recent EKG showed normal sinus rhythm, Left ventricular hypertrophy with strain and Right bundle branch block (Figure 4).

Discussion

Hypertrophic cardiomyopathy (HCM) is a primary myocardial disorder characterized by asymmetric left ventricular hypertrophy and dynamic outflow obstruction and is inherited as a Mendelian autosomal dominant trait. HCM can be caused by a mutation in any 1 of 5 genes that encode proteins of the cardiac sarcomere: beta-myosin heavy chain (on chromosome 14), cardiac troponin T (chromosome 1), troponin I (chromosome 19), alpha-tropomyosin (chromosome 15), and cardiac myosin-binding protein C (chromosome 11). In addition, mutations in 2 genes encoding essential and regulatory myosin light chains have been reported in what may be an extremely rare form of HCM. This genetic diversity is further compounded by intragenic heterogeneity, with a total of more than 100 individual disease-causing mutations identified for these genes; the majority represents missense mutations in which a single amino acid residue is substituted with a different amino acid in the globular head or head-rod junction regions of the myosin molecule. The prevalence of HCM in the general population has been estimated as 1:500, higher than was previously postulated.



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Annual mortality for HCM in an unselected population has been reported to be about 1% to 2%, and sudden death represents the most common cause. Sudden death is assumed to be due to idiopathic ventricular arrhythmias, but hemodynamic factors and myocardial ischemia may be involved as well. Long-term consequences of HCM attributable to outflow obstruction have been emphasized, particularly progression of disabling symptoms and death related to heart failure.

Surgical myectomy has been the preferred method of treatment for symptomatic patients with significant hemodynamic outflow tract obstruction (systolic pressure gradient at rest ≥ 50 mm Hg, after provocation ≥ 100 mm Hg) for more than 20 years. Recently introduced non-surgical septal reduction procedure with intracoronary ethanol, however, is a new promising treatment for patients with hypertrophic obstructive cardiomyopathy. This technique avoids a thoracotomy and cardio-pulmonary bypass to remove the excessive myocardium in the left ventricular outflow tract. Ethanol is infused into one or more septal perforator branches of the left anterior descending coronary artery to cause necrosis and the resultant shrinkage of the proximal hypertrophied interventricular septum. The result is akinesis and enlargement of the narrowed LVOT. Myocardial contrast echocardiography (MCE) is used to guide the targeted delivery of ethanol during nonsurgical septal reduction procedure. MCE can provide quantitative assessment of the extent of myocardium supplied by each septal perforator. MCE definitely is selectively injected into the septal perforator artery can potentially provide an excellent definition of the vascular bed perfused by this vessel and can delineate the area at risk before induced infarction. MCE localized the septal territory and ensured that balloon inflation prevented retrograde spillage into the left anterior descending coronary artery because the septum was the only opacified wall. Coronary angiography was performed to exclude significant atherosclerotic coronary artery disease.

Non-surgical septal reduction therapy was first performed and reported by Dr. Ulrich Sigwart in 1995. His initial series reported on the first three patients in the world to undergo this new procedure. The procedures were all successful, and there was sustained clinical improvement at 1 year. These excellent results in a small series of patients led the way for more studies of this new procedure.

In the US, the first procedure was performed at the Baylor College of Medicine. In their 1-year follow-up study of 50 patients, there was a mean decrease in the outflow tract gradient from 74 mm Hg at baseline to 6 mm Hg at 1 year.

Eleven patients (22%) in this group required permanent pacemaker implantation secondary to refractory heart block after the procedure. Two patients (4%) died during the follow-up period. Data from 2001 showed regression of left ventricular hypertrophy, septal thickness at the infarction site decreased from 20 mm before to 12 mm at 1 year and 10 mm at 2 years. Wall thickness throughout the left ventricular circumference was significantly reduced after NSRP, along with a preserved EF. Regression of LVH may also

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contribute to symptomatic improvement and is probably related to the fact that the increasing hypertrophy in patients with obstruction contributes to a decrease in LV compliance and impaired exercise tolerance. NSRP essentially eliminated the LVOT gradient and was associated with marked reduction in symptoms and improved exercise tolerance. NSRP also resulted in regression of LVH which may be another beneficial effect, given that the frequency of sudden death in HOCM patients increases with increased LVH. Part of this enthusiasm for NSRP derives, understandably, from the relative ease with which NSRP can be performed in experienced hands compared with surgery, involving shortened postoperative recovery, less discomfort and avoidance of cardiopulmonary bypass.

Although, early studies demonstrated an increased incidence of complete heart block, as the technique has been modified with slower rate of injection ethanol (1 to 1.5ml/min instead of bolus) using intracoronary myocardial contrast echocardiography to help target ethanol to the culprit septal segments, the incidence of complete heart block has reduced to 6.7%. The data by Fernandes et al in 2005 on long-term outcome (follow-up of 5 years with mean follow-up 3.6 ± 1.4 years) of alcohol septal ablation had shown resting left ventricular outflow tract gradient at baseline versus last follow-up visit showed a decrease from 74 ± 30 to 4 ± 13 and the dobutamine-provoked gradient of 88 ± 29 decreased to 21 ± 21 mm Hg. Complications of procedures included death 1.5% (2/130), heart block requiring permanent pacemaker 13% (17/130), and coronary dissection 4.4% (6/130). Alcohol septal ablation decreased symptoms and improved exercise performance, indicating that it is an effective procedure for symptomatic HOCM.

Nielsen et al reported in 2003, 50 alcohol septal ablation procedure on 46 patient using echocardiographic contrast localization, slow alcohol injection, and shorter balloon catheters. There was a decrease in the LVOT gradient from 84.2

(± 30.8) mm Hg at baseline, to $18.5 (\pm 14.8)$ mm Hg immediately after alcohol septal ablation. The septal thickness decreased from 2.21cm at baseline, to 1.67cm at 3 months. Three patients (6.7%) of the 45 developed complete heart blocks, requiring permanent pacing.

Even in experienced hands alcohol septal ablation may incur morbidity and mortality similar to that of septal myectomy.

Conclusion

NSRP is a promising alternative procedure for selected patients and an important addition to the therapeutic management of hypertrophic obstructive cardiomyopathy. There is a learning curve in the performance of the procedure which affects the success and complication rate. NSRP is efficacious in providing symptom relief and improving exercise tolerance.

CCT

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ISHAC MILANO: From the Joint Meeting of Workshop IPC & ISHAC - March 22nd – 25th, 2009

By John P. Cheatham, MD

For the first time in four years, the International Symposium on the Hybrid Approach to Congenital Heart Disease (ISHAC) left the friendly confines of Nationwide Children's Hospital and Columbus, Ohio, to join the 7th International Workshop on Interventional Pediatric Cardiology (IPC) in beautiful Milano, Italy. On the last day of the joint meeting, March 25th, over 620 attendees participated in lectures, panel discussions, and live cases highlighting the importance of teamwork between interventional cardiology and cardiothoracic surgery in the hybrid therapies for CHD.

The morning began with a welcome and introduction from the Program Co-Directors, Drs. Mario Carminati, John Cheatham, and Mark Galantowicz. Dr. Cheatham then gave a lecture on how to establish a Hybrid program by overcoming obstacles, suite design, and the teamwork required. The morning session was highlighted by Dr. Ina Michel-Behnke and Dr. Mark Galantowicz updating the audience on the intermediate results of the Hybrid Approach to Hypoplastic Left Heart Syndrome from Giessen and Columbus, respectively. The results were outstanding from these two leading institutions and compared favorably to the Norwood/Sano results of the leading heart centers worldwide. Next, alternative pulmonary artery banding techniques using adjustable external bands and internal flow restrictors were discussed by Dr. Renato Assad from São Paulo and Dr. Cheatham. Finally, the choices and techniques for PDA stenting were discussed by Dr. Carlos Pedra, also from Brazil.



A beautiful venue for a meeting.



Welcome - (left to right) John P. Cheatham, Mark Galantowicz and Mario Carminati.



Sharon Hill lecturing on transcranial doppler in HLHS.



John P. Cheatham lecturing on how to build a Hybrid Program.



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Live case from London of Hybrid Stage I HLHS.



Zhen Xu lecturing on histotripsy.

Dr. Shakeel Qureshi showed the tape-recorded live case broadcast of Hybrid PA banding and PDA stent implant in a baby with HLHS variant and aortic valve stenosis from Evelina Children's Hospital in London. With his surgical colleague, Mr. David Anderson, Shakeel then proceeded to perform successful balloon aortic valvuloplasty using a trans-aortic approach through a purse-string sutured sheath. It was a wonderful demonstration of Hybrid techniques and collaborative thinking to achieve the best result for our patients.

The next live case was broadcast from Giessen where Dr. Dietmar Schranz treated the attendees by demonstrating the techniques for PDA stenting in a baby with pulmonary atresia and ventricular septal defect. A brief discussion between the moderators, panelists, operator and attendees followed and resulted in an excellent clinical result. The morning session continued with a long distance lecture by Dr. Ziyad Hijazi from Chicago, on how to deal with the atrial septum in babies with HLHS, while Dr. Pedra provided insight on how to treat retrograde aortic arch obstruction when it occurs after Hybrid



PEDIATRIC CARDIOLOGIST Tucson, Arizona

Due to expansion we are seeking a third BC/BE pediatric cardiologist to join our Tucson practice. Our practice is part of a 17-member group with offices in the Phoenix and Tucson metropolitan areas. For the Tucson practice we are recruiting a generalist with experience in echocardiography, including trans-esophageal and fetal echo. And, it would be helpful but not essential if one is able to do simple diagnostic catheterizations. In the spring of 2009 we will be moving into a new state-of-art office located a half mile from the main hospital. In addition to our main office, we also see patients in several satellite offices. We cover two main private hospitals and one university hospital.

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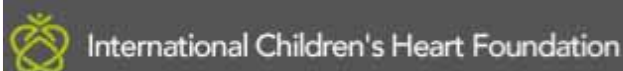
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Stage I palliation. The final morning lecture, given by Dr. Pedro Del Nido from Boston, focused on "rescuing" the borderline left ventricle by surgical removal of the endocardial fibroelastosis and returning to 2-ventricular physiology...sort of the LV "overhaul" procedure!

Lunch time was jam-packed with the announcement of the recipients of the 2nd Fung-Wexner Award for international scientific collaboration within The Heart Center at NCH. Dr. Zhen from Fuwai Cardiovascular Institute, Beijing, China, and Dr. Wold from Nationwide Children's Hospital received the award for their continued work in establishing an International Tissue Bank for CHD. After a delicious and healthy Italian cuisine was served, the attendees prepared for the Perventricular Approach Session by sipping Italian coffee and eating scrumptious pastries. Dr. Gianfranco Butera from San Donato, Milano began the afternoon session by describing the challenges of percutaneous closure of muscular VSD, while Dr. Hijazi compared the technique to perventricular closure of those defects in newborns and small infants. Dr. Del Nido then educated the interventional cardiologists that surgeons have a few tricks up their sleeves by demonstrating live 3-D echo guided closure of muscular VSD using perventricular approach. The session ended with Dr. Shengshou Hu from Fuwai Hospital in Beijing describing their innovative work in the perventricular approach to R-sided obstructive CHD, avoiding CP bypass.

The third live broadcast of the day from Giessen demonstrated the pre-Comprehensive Stage II cardiac catheterization after successful PA bands, PDA stent, and balloon atrial septostomy. This nicely led into the 3rd major topic at ISHAC where intraoperative stents were discussed. Dr. Galantowicz lead off

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ISHAC faculty and program directors at the Gala.

by explaining that surgical removal of endovascular stents is just a matter of attitude...a thought not shared by many cardiac surgeons! Next, Dr. Evan Zahn described the techniques of implanting intraoperative pulmonary artery and aortic stents using both endoscopic and fluoroscopic guidance, once again emphasizing the need for collaboration! Following this lecture, Drs. Ralf Holzer and Alistair Phillips performed the fourth live case broadcast, demonstrating periventricular closure of a muscular VSD, as well as endoscopic-guided intraoperative PA stent implantation from the Cardiovascular Research Surgical Suites in Columbus.

The final session of ISHAC and the Grande Finale of the joint meeting centered on future materials, techniques, and research. Dr. Felix Berger from Berlin gave a preview of new biodegradable stents and polymers. Then Dr. Zhen Xu from the University of Michigan Biomedical Engineering Department dazzled the audience with histotripsy...using therapeutic ultrasound bubble clouds that act as microscopic scalpels. She showed both the surgeons and interventional cardiologists why we may be out of jobs someday!

Dr. Loren Wold from the Center for Cardiovascular and Pulmonary Research Center, Nationwide Children's Hospital, reported the importance of future tissue banking and genetic biomarkers in unraveling some of the mysteries of congenital heart disease. The final lecture from Sharon L. Hill, ACNP, at NCH stressed the importance of determining neurodevelopmental outcomes after HLHS palliation and the future use of transcranial Doppler analysis to determine cerebral blood flow...before, during, and after each staged-procedure. This may provide insight into an important aspect of our patients' care.



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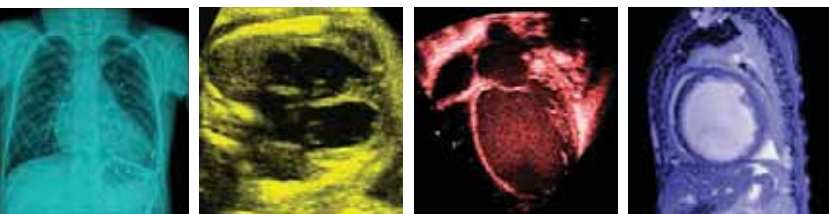
CONTEMPORARY QUESTIONS IN CONGENITAL HEART DISEASE

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DEAR PARTICIPANT

We are delighted to welcome you to the 4th Toronto Symposium, Contemporary Questions in Congenital Heart Disease.

After three extraordinarily successful meetings in 2005, 2006 and 2007, the topic of our 2009 meeting will be Heart Failure and Transplantation.

Once again, we have invited a world-class faculty of scientists, physicians, surgeons, and allied professionals to participate with the Toronto team in a "state of the art" conference.

The Toronto Symposium aims to be a little different from the usual medical meeting. The title of each lecture, no matter whether addressing issues of basic science or clinical management, is framed as a topical question. Consequently we expect that the answers will be of direct relevance to your practice. This meeting will be suitable for anyone working in the field of congenital heart disease, but please note that we are limited to just 250 places, and have been sold-out prior to previous meetings.

So register early to avoid disappointment!

While there are some concurrent sessions, be assured there is no need for you to miss anything. Each of the lectures will be recorded, and each participant will receive a DVD shortly after the meeting. Again, this is a little out of the ordinary, showing both a video of the lecturer in real time, and the simultaneous Power-Point presentation. An example of the format can be seen on our symposium website at www.sickkids.ca/Centres/heart-centre/Cardiac-symposium. Copies of the DVD's from previous symposia can be purchased by e-mailing the Symposium organizer at cardiac.symposium@sickkids.ca.

We are looking forward to a focused, detailed, and rewarding meeting, located on Toronto's beautiful waterfront. We hope you will be able to join us.

Sincerely,
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The Toronto Congenital Cardiac Centre for Adults will
celebrate its 50th anniversary on October 7, 2009.



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Pediatric Cardiology of New Mexico, the state's busiest and most comprehensive pediatric cardiology practice, is recruiting a 5th BC cardiologist with 2-3 years of post-fellowship experience in interventional catheterization. We are a private practice with comprehensive surgical and interventional services and a very robust state-wide clinical outreach system. We are affiliated with New Mexico's largest tertiary care hospital which boasts a new wing containing a 58-bed NICU and 21-bed PICU with around the clock in-house neonatologists, pediatric intensivists and pediatric hospitalists.

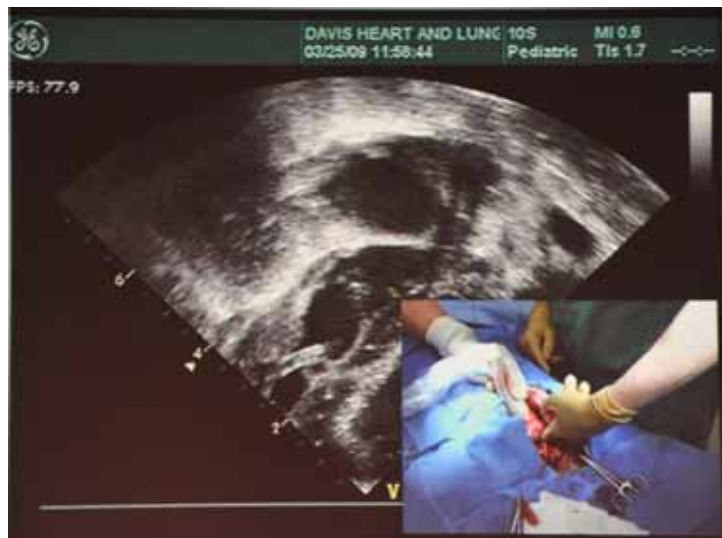
New Mexico, the "Land of Enchantment," is conducive to an active and outdoors lifestyle with abundant opportunities for skiing, hiking, fly-fishing and mountain biking.

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Live case from Columbus - Ralf Holzer and Alistair Phillips showing perventricular MVSD closure.

ISHAC and Workshop IPC came to a close and Program Co-Directors, John Cheatham and Mark Galantowicz reminded everyone that ISHAC 2010 returns to Columbus, Ohio, September 1st - 3rd with a 2-day Symposium and 1-day special hands-on Skills Workshop. Our Keynote Speakers will be Professor Philipp Bonhoeffer from Great Ormond Street and Mr. Marc de Leval from the International Congenital Cardiac Centre, London, UK. Rumor even has it that an Ohio State University Buckeye football game may be a part of the meeting...stay tuned! Then, Drs. Gianfranco Butera and Massima Chessa, on behalf of Dr. Carminati and the entire program staff from Ab Medica, thanked everyone for attending, the faculty, and the sponsors and welcomed them to the 8th Workshop IPC scheduled for 2011 in Milano, Italy.

CCT

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Experience with a Novel Miniaturized Multi-plane Transesophageal Echocardiographic Transducer

By Eric M. Graham, MD and
Girish S. Shirali, MBBS

With the advances in cardiac surgery, anesthesia and post operative care and the manifest advantages of correcting abnormal cardiac physiology early, there continues to be a growing trend towards complete repair of complex intracardiac operations undertaken in smaller and smaller patients. Transesophageal echocardiography (TEE) has been shown to be safe and cost effective in the intraoperative period and has become standard of care in many institutions.¹⁻³ Prior to starting the operation, TEE can confirm or further delineate the cardiac anatomy. Prior to separating from cardiopulmonary bypass TEE can assure adequate intracardiac de-airing, lowering the risk of air emboli. After separation from cardiopulmonary bypass TEE can confirm the adequacy of repair, assess atrioventricular and arterial valvar competency and assess ventricular filling and function prior to leaving the operating room. Confirming the adequacy of a repair prior to leaving the operating room is important because inadequate repairs can be immediately corrected. Whereas, patients leaving the operating room with significant residual defects have experienced increased morbidity, mortality and financial cost.³⁻⁶ Despite the advantages of TEE there are some potential disadvantages. These include esophageal or gastric trauma or perforation, left atrial, aortic or airway compression resulting in hemodynamic or ventilation compromise, and inadvertent tracheal extubation. Despite these potential concerns, several studies have shown a low rate of complications, predominantly airway compression and inability to pass the transducer.² Neonates and small infants are particularly at risk for the

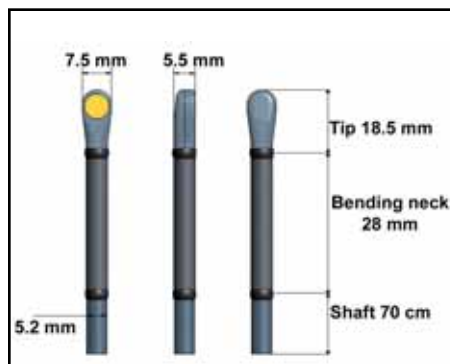


Figure 1: This demonstrates the dimensions of the micro-multiplane TEE probe.

potential complications given their small size compared to the transducer size.^{1, 2} Despite current reports that clinically significant hemodynamic and ventilation compromise is infrequent even in children as small as 2-5 kg, concerns remain.^{7, 8} These have led to reluctance to use TEE in small infants and neonates. Paradoxically, it is the small neonate undergoing intracardiac repair that probably has lower tolerance for residual defects, and thus more to gain from intraoperative TEE.

Pediatric TEE probes that are in general usage are equipped with either a biplane or multiplane transducer. The biplane probe has 64 elements in each transducer, an output frequency of 5.5 or 7.5 MHz and a tip dimension of 9.1 x 8.8 mm. The multiplane probe has a 48 element transducer with a center frequency of 6 MHz (range 4 to 7 MHz) and a tip dimension of 10.7 x 8.0 mm.⁹ Improving technology has led to a new miniaturized multiplane TEE. We report our experience with the world's smallest multiplane TEE probe, the micro-TEE transducer (Philips Medical, Andover,

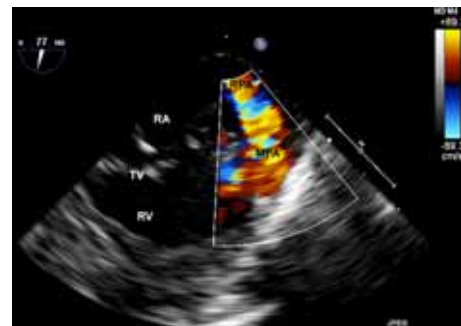


Figure 2: Color flow Doppler imaging of the right ventricular inflow – outflow view in a 2.5 kg patient who was undergoing surgery for a large membranous ventricular septal defect. MPA, main pulmonary artery; RA, right atrium; RPA, right pulmonary artery; RV, right ventricle.

MA, USA). The shaft width is 5.2 mm with a transducer tip width of 7.5 mm and height of 5.5 mm (Figure 1, Table 1). The micro-TEE is a 32 element phased array multiplane TEE transducer complete with 2D, color, pulse wave, Hi PRF and continuous wave Doppler, MMode and CF MMode. It has a center frequency of 6 MHz (range 3.2 MHz to 7.4 MHz). The controls on this probe are identical to those on the minimultiplane TEE that is in general usage. Specifically, there is a 180 degree manual image plane control with angular display, anterior and posterior articulation and an articulation brake. The transducer has tip temperature sensing and display for added safety.

Patients are currently being enrolled at the Medical University of South Carolina's Institutional Review Board approved study evaluating image quality of this probe in neonatal and pediatric TEE. To date, over 20 patients have been studied, primarily in the operating room but also in the catheterization suite under general anesthesia. Probe insertion has been attempted in patients 1.7 kg and above. All insertions have been successful and well-tolerated. High-quality diagnostic images have been obtained consistently (Figure 2-4).

This advance enables us to provide the small neonate and infant with intra-operative TEE imaging, thus optimizing repairs and outcomes.

Table 1 Transducer comparison

	Tip Length (mm)	Tip Width (mm)	Tip Height (mm)	Shaft (mm)	Pt Wt (kg)
X7-2t	40	16.6	13.1	10	30
Mini-mTTE	27	10.6	8	7.5	3.5
Micro-mTEE	18.5	7.5	5.5	5.2	?



CHICAGO Rush University Medical Center Electrophysiologist

The Department of Pediatrics, the Electrophysiology, Arrhythmia, and Pacemaker Service, and the Center for Congenital and Structural Heart Disease at Rush University Medical Center, located in downtown Chicago, seek an electrophysiologist.

We are in quest of a cardiologist with fellowship training in pediatric and congenital/structural electrophysiology. The candidate should have expertise in invasive and non-invasive electrophysiology and skills and expertise in diagnosis and management of complex arrhythmias. Willingness to perform routine adult EP procedures is highly desirable. Conjoint appointment in the Department of Internal Medicine will be considered based on the candidate's qualification and level of interest.

This recruitment is part of a key strategic growth initiative in a multidisciplinary advanced congenital/structural cardiology program with state of the art mechanical support and clinical trials. Experience in clinical research is desirable. Candidates should be eligible for faculty appointment at the Assistant Professor or Associate Professor level. Rush is home to one of the first medical colleges in the Midwest and one of the nation's top-ranked nursing colleges, as well as graduate programs in allied health, health systems management and biomedical research.

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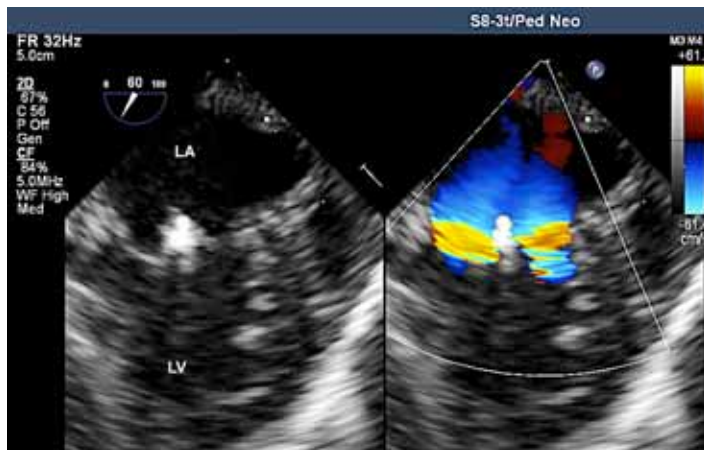


Figure 3: This image was obtained in a 3 month old, 5 kg infant with atrioventricular septal defect and severe left-sided AV valve regurgitation. In addition to closing the 'cleft', the surgeon created a double-orifice left AV valve. This is a 60-degree image of the surgically-created double-orifice mitral valve, demonstrating prograde flow through both orifices. LA, left atrium; LV, left ventricle.



Figure 4: Simultaneous 2D and color flow Doppler in a 4 month old, 5.4 kg infant undergoing repair of tetralogy of Fallot. The asterisk marks the severely anteriorly malaligned outlet septum. There is right to left flow across the VSD. Note the long-segment stenosis of the right ventricular outflow tract, pulmonary valve and main pulmonary artery. MPA, main pulmonary artery; RA, right atrium; RV, right ventricle; RVOT, right ventricular outflow tract; VSD, ventricular septal defect.



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Acknowledgments: The authors thank the faculty and fellows of the Divisions of Pediatric Cardiology, Cardiothoracic Surgery, and Cardiothoracic Anesthesia, and the Pediatric Cardiology sonographers for their assistance with this project.

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Director of Interventional Cardiology and Director of Non-Invasive Imaging

The University of Maryland Hospital for Children is developing a comprehensive Children's Heart Center to meet the cardiovascular healthcare needs of the children of Maryland. We are recruiting for two full time positions: Director of Interventional Cardiology and Director of Non-Invasive Imaging. Sub-specialty board certification with advanced sub-specialty training or equivalent work experience is required for each position. The ideal candidates will have proven leadership and program development experience. Clinical duties will focus primarily within the domain of each position, although all Children's Heart Center physicians participate to varying degrees in the general pediatric cardiology and outreach practices. The Children's Heart Center supports integrated quality enhancement and clinical research practices to improve patient outcomes and patient/family experience.

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University of Maryland Department of
Pediatrics
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Medical News, Products & Information

Toshiba Launches InTouch Flex Service Agreement

To further its commitment to providing innovative, quality customer solutions, Toshiba America Medical Systems, Inc. has launched the new InTouch™ Flex Service Agreement, offering customers unparalleled flexibility, security and value. Considered the first of its kind in the diagnostic imaging industry, the InTouch Flex Service Agreement allows customers to secure fixed price points for both the full service and partnership agreements at the point of purchase and after the warranty, convert the agreement back and forth between a full service security agreement and a partnership agreement, as needed. This flexible service approach enables customers to adapt service plans throughout the lifetime of the agreement to match real-time needs.

"Toshiba's InTouch Flex Service Agreement provides us a flexible, cost-effective service arrangement to meet our changing needs today and in the future," stated George Morley, Director of Biomedical Engineering at PinnacleHealth, a premier nonprofit healthcare system with in-house service staff already under the new InTouch Flex Service Agreement. "The flexibility offered by Toshiba is unmatched in the service marketplace. It allows us to utilize our in-house staff effectively, but also rely on Toshiba Service when additional support or expertise is required."

PinnacleHealth, serving Central Pennsylvania, is the first healthcare system to take advantage of the InTouch Flex Service Agreement. Under the InTouch Flex Service Agreement, service problems are first handled by PinnacleHealth's in-house service team through diagnosis remotely via phone or by a service team member on site. Receiving the same training as Toshiba engineers, PinnacleHealth service staff can troubleshoot issues quickly and order product equipment for repairs directly from Toshiba. If PinnacleHealth's service team needs additional assistance, a service engineer from Toshiba arrives within two hours to address the customer's needs. This one-of-a-kind service partnership results in a situation where the customer's interests are at the forefront for both PinnacleHealth service staff and Toshiba.

PinnacleHealth purchased more than \$14.5 million of Toshiba medical imaging equipment as part of a five-year strategic business agreement, most of which is serviced under the new InTouch Flex Service Agreement. PinnacleHealth sources 85% of its new medical imaging systems from Toshiba's leading product portfolio including CT, MR, Vascular X-ray, X-ray and Ultrasound.

As the service needs of hospitals, IDNs and imaging centers change over time, Toshiba's InTouch Flex Service Agreement enables these facilities to adjust their service agreement to meet these needs. The InTouch Flex Service Agreement allows customers to switch between the following existing full service and partnership agreements:

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"We pride ourselves on viewing service as a collaboration, listening to our customers' needs and providing unique customized solutions," said Ted Nemetz, Vice President, Service Business Unit, Toshiba. "Whether a customer requires a partnership agreement with the majority of service handled by their in-house staff or a full service agreement with Toshiba service specialists handling the workload, the service agreement pricing never exceeds the original point of purchase pricing."

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These fellowships will provide salary support for a 1-3 year program of clinical, translational, or basic research. Applicants should have completed their core clinical training in a relevant sub-specialty (paediatric cardiology, paediatric cardiac surgery, paediatric cardiac critical care, adult congenital heart disease, etc), and ideally will have some preexisting research experience. These posts are open to MD, and non-MD trainees, and registration for a postgraduate degree will be possible for appropriately trained individuals. Small amounts of start-up funding may be available from the Heart Centre but projects will be expected to garner additional peer-reviewed funding where necessary.

Applicants are encouraged to communicate with potential research supervisors within the Heart Centre (<http://www.sickkids.ca/Centres/heart-centre/Our-members/index.html>), prior to or coincident with their application. Alternatively, de-novo applications can be made but should include a brief research plan with clearly defined aims and objectives so that they can be directed to potential supervisors.

Please include a full curriculum vitae and research plan with all applications. The closing date for applications is July 22 2009. Remuneration - commensurate with status, to be negotiated.

All qualified candidates are encouraged to apply; however Canadian citizens and permanent residents will be given priority. The Hospital for Sick Children hires on the basis of merit and is committed to equity in employment.

Inquiries and applications should be made to:
Ms. Sue Morton, Administrative Assistant
Labatt Family Heart Centre
The Hospital for Sick Children
555 University Avenue, room 1725
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e-mail: sue.morton@sickkids.ca

Philip Witchel Research Fellowship in Paediatric Heart Failure

The Cardiomyopathy and Heart Function Service of The Labatt Family Heart Centre, The Hospital for Sick Children, and the University of Toronto, are pleased to offer a funded research position in the field of paediatric heart failure. The fellowship has been active since 2005, and an appointment is usually held for 12 to 24 months. The position is open to applicants to begin in January 2010.

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Candidates interested in pursuing this unique opportunity should forward their CV, and letter of Intent to:

Paul F. Kantor MD
Head, Cardiomyopathy and Heart Function Service
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2009 Request for Barth Syndrome Research Proposals

The Barth Syndrome Foundation, Inc. (BSF) and its affiliates are pleased to announce the availability of funding for research internationally on the natural history, biochemical basis, and treatment of Barth Syndrome.

Barth syndrome is a serious X-linked genetic condition associated with cardiomyopathy, neutropenia, skeletal muscle weakness, exercise intolerance, growth delay, and diverse biochemical abnormalities (including defects in mitochondrial metabolism and phospholipid biosynthesis). Because many clinical and biochemical abnormalities of Barth Syndrome remain poorly understood, we are seeking proposals for research that may shed light on any aspect of the syndrome. We are determined to find improved treatments and ultimately a cure for this rare and under-diagnosed disorder.

We are most interested in providing "seed money" to be used by experienced investigators for the testing of initial hypotheses and collection of preliminary data leading to successful long-term funding by the National Institutes of Health (NIH) and other major granting institutions around the world. In addition, we are especially interested in attracting new investigators to the very interesting field of Barth Syndrome research.

We anticipate awarding several one- or two-year grants of up to \$40,000 each. Funds will be available as soon as the successful grant applicants have been notified.

We have a simple grant process. Applications should be of 10-15 pages in length and must follow the instructions listed on the BSF website. In general terms, detailed information about the specific aims, significance, research design and methods, personnel, and budget will be required, along with evidence of application to the relevant Institutional Review Board for any work involving human subjects.

Completed proposals will be forwarded to the BSF Scientific and Medical Advisory Board (as well as outside reviewers, in certain cases) for evaluation. Based on the recommendations of the Scientific and Medical Advisory Board, the BSF Board of Directors and those of our three international affiliates will make the final funding decisions for the grant applications. Please review our "Grants Awarded" webpage for a listing of grants that BSF has awarded to date.



Director of Electrophysiology & Pacing

The Heart Center at Nationwide Children's Hospital (NCH) and the Ohio State University is seeking a second board-certified pediatric and/or adult electrophysiologist to join our 20 member multidisciplinary Section as Director of Electrophysiology and Pacing. The candidate must be established in the electrophysiology community, command an innovative vision and prepared to participate in programmatic planning that encompasses all aspects of our Heart Center's mission (excellence in clinical service, education and research). In addition to Electrophysiology our Heart Center has established programs in Adult CHD & Transition, Interventional Catheterization, Non-Invasive Imaging, Heart/Heart-Lung/Lung Transplantation, innovative neonatal surgery & hybrid approach and Advanced Practice Nursing. We have a well-supporting out-patient network. In 2008 our Heart Center was recognized by both USNWR and Parent Magazine and we are an Optum Center of Excellence. We are partnered with the Center for Cardiovascular and Pulmonary Research at NCH-Research Institute. NCH is a free standing children's hospital with a dedicated 10 bed CT-ICU and 15 bed Cardiac Step-down Unit, two state-of-the-art catheterization laboratories including an EP laboratory. Our Heart Center has a dedicated service line administration including a VP and Director of Nursing. Our cardiology fellowship is recognized by both the American Boards of Pediatrics and Internal Medicine.

Candidates may submit their curriculum vitae to Timothy F. Feltes, MD, Nationwide Children's Hospital, 700 Children's Drive ED 617, Columbus, Ohio 43205; Call: 614 772-2565 or e-mail to: timothy.feltes@nationwidechildrens.org



The Ohio State University is an Equal Opportunity, Affirmative Action Employer. Women, minorities, veterans, and individuals with disabilities are encouraged to apply.



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A Cause for Today.... A Cure for Tomorrow

The deadline for submission of completed research grant applications from interested researchers is October 31, 2009. Grants will be awarded early March 2010.

Contact Information:
Matthew J. Toth, PhD
Science Director
Barth Syndrome Foundation, Inc.
mtoth@barthsyndrome.org or
www.barthsyndrome.org

First-Degree Relatives of Patients with the Most Common Cardiac Birth Defect Should be Screened for Larger-Than-Normal Aortas

Bicuspid Aortic Valve (BAV), a condition in which patients' aortic valves have just two leaflets instead of the normal three, is the most common cardiac anomaly, affecting up to two percent of the general population. The defect can result in calcification deposits on the heart valve, leakage of the valve and may result in a feeling of tightness in the chest, as well as shortness of breath. The condition is easily diagnosed; often physicians can hear a "click" or a murmur when they listen to a BAV patient's heart with a stethoscope.

Studies have shown that BAV is likely genetic, although the gene has not been identified, and in some families, incidence of this defect could run as high as 20%.

A new study, published in the *Journal of the American College of Cardiology*, suggests that nearly a third of first-degree relatives (siblings, children or parents) of BAV patients are likely to have enlarged aortas, a potentially serious condition that can only be detected by undergoing transthoracic echocardiograms. This was found even in the absence of any abnormalities of the heart valve itself.

According to the study, 32% of first-degree relatives with no heart valve abnormality had significantly larger aortas that expected for age, gender and body size as compared to no enlargement seen in control patients. Also, the study found that the aortas of the first-degree relatives had abnormal stiffness similar to the patients with congenital bicuspid valve. Generally, when aortas are 50 millimeters in diameter, surgery is recommended in order to prevent a rupture of the aorta.

"If you know that a relative does have bicuspid aortic valve, then you know that you should be screened," said study author Kirsten Tolstrup, MD, Assistant Director of the Cardiac Noninvasive Laboratory at the Cedars-Sinai Heart Institute. "BAV appears to be a genetic condition that has many different manifestations, so we will be studying the genes."

Kirsten Tolstrup, MD, Assistant Director of the Cardiac Noninvasive Laboratory at the Cedars-Sinai Heart Institute, is available to discuss the study's findings and provide additional details.

This study, conducted among 54 patients with bicuspid aortic valve and 48 first-degree relatives of those patients as well as 45 matched controls found:

- 32% of apparently healthy first-degree relatives have enlarged aortas
- 53% of BAV patients had enlarged aortas
- 9.4% of first-degree relatives had BAV

The findings suggest that patients with bicuspid aortic valve and their first-degree relatives should have a screening echocardiogram to be evaluated for dilated aorta and bicuspid aortic valve.

The study abstract can be accessed at:
<http://content.onlinejacc.org>

For additional information, call Sandy Van at 800-880-2397 or Sally Stewart at 310-248-6566.

Citation: *Journal of the American College of Cardiology*, June 8, 2009, "Aortopathy is Prevalent in Relatives of Bicuspid Aortic Valve Patients"

Comprehensive Cardiogenetic Testing for Families of Sudden Unexplained Death Victims Can Save Lives

Relatives of a young person who dies suddenly should always be referred for cardiological and genetic examination in order to identify if they too are at risk of sudden death, a scientist told the annual conference of the European Society of Human Genetics in May. Dr. Christian van der Werf, a research fellow at the Department of Cardiogenetics, Academic Medical Centre, Amsterdam, The Netherlands said that, although his team's research showed that inherited heart disease was present in over 30% of the families of sudden unexplained death (SUD) victims, the majority of such relatives were currently not being referred for examination.

When an individual aged 1-50 years dies suddenly, autopsy reveals an inheritable heart disease in the majority of the victims. But in approximately 20%, autopsy does not reveal cause of death. "We thought that cardiological and genetic examination of surviving first degree relatives of these SUD patients might reveal an inherited heart disease," said Dr. van der Werf.

In the largest such study to date, the team looked at the outcome of first degree relative screening in 127 families who



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Chief of the Division of Cardiology

The Department of Pediatrics of the University of Texas Health Science Center at San Antonio is seeking a physician leader for the position of Chief of the Division of Cardiology. The candidate must be fellowship trained and board certified in pediatric cardiology and either possess or be able to easily obtain an unrestricted Texas medical license. The tenure track faculty appointment will be at the associate or professor level.

The Division Chief will be responsible for developing the Children's Heart Network, a joint program of the Departments of Pediatrics and Surgery and CHRISTUS Santa Rosa Children's Hospital. The Division Chief, along with the Chief of Pediatric Congenital Heart Surgery, will be charged with developing a regional pediatric cardiology program including research, state-of-the-art clinical services, and strong educational programs for both medical students and pediatric residents. The Division Chief has the opportunity to join an established clinical practice with 4 pediatric cardiologists, including an echocardiographer and an interventionalist. The Division Chief will have the ability to recruit additional faculty members and initiate a research portfolio for the Division. CHRISTUS Santa Rosa Children's Hospital is a 200+ bed facility providing care to more than 150,000 children each year.

The School of Medicine has approximately 230 medical students at each level, and Division faculty is engaged in the training of these medical students and 45 pediatric residents. Candidates with interests in epidemiology, interventional cardiology, cardiac intensive care, or heart failure/transplant are encouraged to apply.

Please submit a letter of interest and curriculum vitae to Thomas C. Mayes, M.D., M.B.A., Professor and Chairman, Department of Pediatrics, The University of Texas Health Science Center at San Antonio, 7703 Floyd Curl Drive, San Antonio, Texas 78229-3900.

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had suffered an SUD and where either there had been no autopsy (53.8%), or the autopsy did not reveal a cause of death. The average age at death of the SUD victims was only 29.8 years old.

The initial examination of the relatives consisted of taking personal and family medical history and a resting ECG. A second cardiac autopsy of the SUD victim was undertaken if tissue had been stored and was available. Additional cardiological examinations of the relatives were performed where necessary. Genetic analysis of the associated candidate gene(s) was performed in material obtained from the deceased person or in those relatives who showed clinical abnormalities.

The researchers found inherited heart disease in 36, or 32% of the families. These results meant that doctors were able to treat affected relatives and try to prevent their succumbing to sudden cardiac death. "The scale of heart disease that we found in such families underlines the necessity for general practitioners to refer first degree relatives of SUD victims to a specialised cardiogenetics department as soon as possible", said Dr. van der Werf. "Currently we estimate that only 10% of SUD families are being examined for inherited heart conditions.

The study is the second report from the registry of families who attended the Amsterdam centre's cardiogenetics department because of unexplained sudden death of a relative aged 1-50 years. The scientists intend to continue to report the yield of family screening in an increasing number of families.

"At present we are conducting a study to stimulate general practitioners and other involved physicians to request autopsy and DNA-storage for SUD patients and to refer relatives to a cardiogenetics department after a case of sudden death at young age. We hope this will lead to identification of more families at risk of sudden cardiac death, in which preventive measures then can be taken" said Dr. van der Werf.

"Relatives of young sudden death victims are often referred to cardiologists for cardiological examination. We believe relatives should instead be referred to cardiogenetics departments, where clinical geneticists, cardiologists and psychosocial workers cooperate. These professionals specialise in inherited heart diseases and their clinical and psychosocial implications, and can provide a better quality of care. Additionally, cardiologists should receive more education in inherited heart diseases. By taking these measures we can save lives and avoid further distress for families who have already suffered enough," he said.



Barth Syndrome
Foundation

The Barth Syndrome Foundation

P.O. Box 974, Perry, FL 32348

Tel: 850.223.1128

info@barthsyndrome.org

www.barthsyndrome.org

Symptoms: Cardiomyopathy, Neutropenia, Muscle Weakness, Exercise Intolerance, Growth Retardation

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PEDIATRIC CARDIOLOGISTS

The **Division of Pediatric Cardiology at the University of Utah School of Medicine** and Primary Children's Medical Center is recruiting additional pediatric cardiologists with major interests in:

- **noninvasive imaging,**
- **transplant/heart failure, or**
- **cardiac intensive care.**

The candidate should be BC/BE in pediatric cardiology and have a strong clinical background with expertise and interest in at least one of the areas listed above. The division currently consists of 21 division members and has a very active, growing clinical program, an active, thriving fellowship training program, and a very active clinical research program; it is one of the participating centers in the Pediatric Heart Disease Clinical Research Network funded by the NIH. Protected time and mentoring for clinical research will be available within the Division for clinical research studies.

The successful candidate will receive a faculty appointment at the University of Utah. The Pediatric Cardiology Division is based at Primary Children's Medical Center, a tertiary referral center for a three-state area located on the hills overlooking Salt Lake City. The area offers an excellent quality of life with immense cultural and recreational opportunities readily available. The University of Utah is an Equal Opportunity Employer and welcomes applications from minorities and women and provides reasonable accommodations to the known disabilities of applicants and employees.

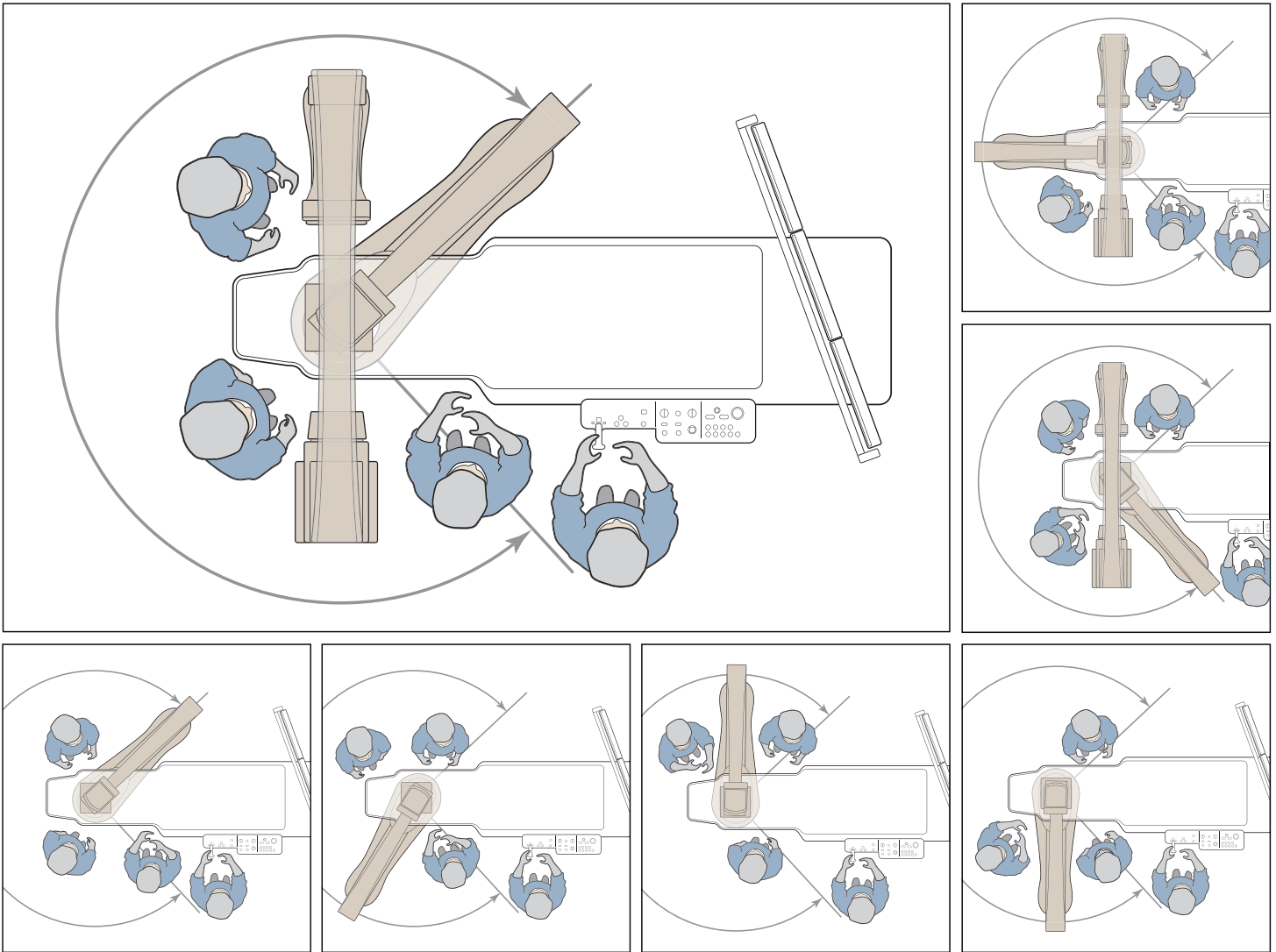
Interested individuals should contact:

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Do you or your colleagues have interesting research results, observations, human interest stories, reports of meetings, etc. that you would like to share with the congenital cardiology community?

Submit a summary of your proposed article to Congenital Cardiology Today at: RichardK@CCT.bz



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