Endovascular stents in the management of coarctation of the aorta in the adolescent and adult: one year follow up

D A Harrison, P R McLaughlin, C Lazzam, M Connelly, L N Benson

Abstract

Objectives—To test the hypothesis that endovascular stents used with dilation of coarctation of the aorta (CoA) improve late outcomes. Balloon dilation for CoA has been limited by concerns over the risk for acute dissection, late restenosis, or aneurysm formation.

Design—All patients seen with CoA between November 1994 and September 1997 underwent attempted stent implantation. Follow up was obtained for all patients and a subgroup (n = 18) had repeat catheterisation at a mean (SD) of 1.3 (0.5) years to assess residual gradient and stent-CoA morphology.

Results—Stents were placed in 27 patients (15 male and 12 female patients, mean age 30.1 (13.1) years), of whom seven had prior surgical coarctectomy and one had a prior balloon dilation. Hypertension was present in 26 patients (mean pressure 164 (26)/86 (13) mm Hg), of whom 16 were on antihypertension drugs. CoA gradients were 46 (20) mm Hg (range 18–106 mm Hg) at baseline and 3 (5) mm Hg after the procedure. One patient had a stroke following the procedure; another patient had incomplete dilation and underwent a second procedure. At 1.8 (1) years after the procedure the mean pressure was 130 (14)/74 (11) mm Hg with seven patients on antihypertension treatment. The clinical gradient was 4 (8) mm Hg (range 0–32 mm Hg). At follow up angiography, the mean gradient was 4 (6) mm Hg, and two patients had a gradient over 10 mm Hg. Aneurysms formed in three patients at the dilation site; one patient was referred for surgery.

Conclusion—In this age group stent management for CoA appears to be an effective technique and results in sustained reduction in CoA gradients at early term follow up, but aortic aneurysm was detected in 17% of patients who had repeat angiography.

Keywords: angioplasty; coarctation of the aorta; congenital heart defects; stents

Surgical correction of coarctation of the aorta (CoA) improves a patient’s natural history and assists in control of hypertension.\(^1\) Residual problems of recurrent CoA include need for reoperation, persistent hypertension, and the uncertain significance of exercise induced hypertension.\(^2\)\(^3\)\(^4\)

Over the past two decades non-surgical catheter directed management for CoA has been attempted by using balloon angioplasty.\(^5\)\(^6\) Although initial procedural success has been well documented, balloon management has been complicated by aortic dissection,\(^7\)\(^8\)\(^9\)\(^10\)\(^11\)\(^12\)\(^13\)\(^14\)\(^15\)\(^16\)\(^17\)\(^18\) incomplete relief of the CoA gradient,\(^9\)\(^13\)\(^15\)\(^16\)\(^17\)\(^18\) cerebrovascular accidents,\(^19\)\(^20\)\(^21\)\(^22\) and aneurysm formation.\(^9\)\(^15\)\(^16\)\(^17\)\(^20\)\(^21\)\(^22\) Perhaps the most limiting feature of CoA angioplasty is significant residual gradient (> 20 mm Hg) at late follow up in 7%–36% of patients.\(^1\)\(^6\)\(^15\)\(^16\)\(^17\)\(^18\)\(^19\)\(^20\)\(^21\)\(^22\)\(^23\)\(^24\) Residual gradient has been attributed to an incomplete immediate result, recurrent stenosis,\(^23\)\(^24\) and associated transverse arch hypoplasia.

Endovascular stents have been applied in the management of CoA as an alternative to balloon dilation alone.\(^14\)\(^19\)\(^20\)\(^25\) Stent use has been limited by the maximal diameters that can be achieved without inducing strut fatigue; follow up of their use has been of limited duration. Recently, larger stents have become available (Johnson & Johnson Interventional, Warren, New Jersey, USA), which can be expanded to 25 mm in diameter, well within the arch diameter of an adult aorta. We hypothesised that the use of such implants would reduce the development of restenosis, would not compromise vascular wall integrity, and would avoid the need for reintervention. We report the immediate and early follow up of a group of consecutive adolescents and adults for whom stent angioplasty for CoA was the primary management strategy, including haemodynamic and angiographic assessment after implantation.

Methods

Patient population
Consecutive adult and adolescent patients (n = 27, weight > 50 kg) referred for management of CoA at either The Hospital for Sick Children or Toronto Congenital Cardiac Centre for Adults, Toronto, Canada, between November 1994 and September 1997 were offered and accepted participation in catheter directed management of their lesion. Because efficacy and safety issues have not been defined in terms of anatomy, no patient was excluded because of wall calcification, isthmal hypoplasia, CoA location (for example, near the left subclavian artery), or arch irregularity. Informed consent was obtained for all patients before the procedure.
The CoA was dilated with a 15 mm diameter balloon in six patients, an 18 mm balloon in six patients and the 5 cm stent in the remainder. Available the 4 cm stent was placed in 15 patients. In one patient in the proximal isthmus CoA minimum diameter was 6.6 (3.6) mm. The mean peak gradient was 46 (20) mm Hg. Preinterventional changes in blood pressure (mm Hg) were shown.

Baseline blood pressure proximal to the CoA was 149 (30)/80 (15) mm Hg (mean 103 (18) mm Hg), and distal to the CoA was 102 (18)/75 (12) mm Hg (mean 84 (13) mm Hg) (p = 0.001 systolic mean pressure difference). The mean peak gradient was 46 (20) mm Hg. CoA minimum diameter was 6.6 (3.6) mm (range 1–16 mm). Of the two stent lengths available the 4 cm stent was placed in 15 patients and the 5 cm stent in the remainder. The CoA was dilated with a 15 mm diameter balloon in six patients, an 18 mm balloon in six patients (one pre-dilated with a 15 mm balloon), and a 20 mm balloon in 15 patients. In one case the CoA was immediately distal to the left subclavian artery and the stent was deployed on the day following the procedure was planned for the majority of patients.

FOLLOW UP PROTOCOL
Patients were scheduled for clinical follow up at six weeks and one year after the procedure. If they were normotensive at the first follow up assessment, patients on antihypertension drugs were given a trial off treatment.

Patients (n = 18) underwent a haemodynamic reassessment of the stent gradient and stent-CoA morphology one year after the procedure. Angiography was performed to assess the residual CoA diameter, the possibility of intimal hyperplasia, the presence of an aneurysm (defined as > 2 mm dyskinesis of the aortic wall), and the position of the stent relative to the aortic wall. Two investigators (LB, DH) measured aortic diameters off line.

DATA ANALYSIS
All measurements were calculated as mean (1 SD). Groups were compared by a Student’s t test (paired analysis where appropriate) with the level of significance set at 0.05. Groups were also compared using χ² or Wilcoxon rank-sum tests where appropriate.

Results

POPULATION
Twenty seven patients (15 male patients) underwent stent implantation for CoA. Mean age at procedure was 30.1 (13.1) years (range 14–63 years). Seven patients had had previous surgery for CoA and balloon angioplasty had been performed in one patient. Hypertension was present in 26 patients, with a mean systemic blood pressure of 164 (26)/86 (13) mm Hg, and 16 patients were on antihypertension drugs. Seven patients were taking more than one drug; of the nine patients on monotherapy, five were on a β blocker, three were on an angiotensin converting enzyme inhibitor, and one was on a calcium channel blocker. Cardiac surgery for problems other than CoA was required for four patients (aortic valve replacement in three, closure of ventricular septal defect in one). One patient presented with myocardial infarction at age 34 years and was scheduled for coronary artery bypass. CoA was managed with stent implantation before surgery at the request of the cardiac surgeon.

PROCEDURE
Baseline blood pressure proximal to the CoA was 149 (30)/80 (15) mm Hg (mean 103 (18) mm Hg), and distal to the CoA was 102 (19)/75 (12) mm Hg (mean 84 (13) mm Hg) (p = 0.001 systolic mean pressure difference). The mean peak gradient was 46 (20) mm Hg. CoA minimum diameter was 6.6 (3.6) mm (range 1–16 mm). Of the two stent lengths available the 4 cm stent was placed in 15 patients and the 5 cm stent in the remainder. The CoA was dilated with a 15 mm diameter balloon in six patients, an 18 mm balloon in six patients (one pre-dilated with a 15 mm balloon), and a 20 mm balloon in 15 patients. In one case the CoA was immediately distal to the left subclavian artery and the stent was deployed on the day following the procedure was planned for the majority of patients.
Table 1  Haemodynamic measurements: immediate results and at follow up

<table>
<thead>
<tr>
<th></th>
<th>Ascending aortic pressure (mm Hg)</th>
<th>Descending aortic pressure (mm Hg)</th>
<th>Mean pressure ascending aorta (mm Hg)</th>
<th>Mean pressure descending aorta (mm Hg)</th>
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<tbody>
<tr>
<td>Before procedure</td>
<td>149 (30)/80 (15)</td>
<td>102 (19)/73 (12)</td>
<td>103 (18)</td>
<td>84 (13)</td>
</tr>
<tr>
<td>After procedure</td>
<td>132 (3)**/80 (19)</td>
<td>130 (1)***/80 (19)</td>
<td>98 (23)</td>
<td>97 (2)**</td>
</tr>
<tr>
<td>Follow up</td>
<td>125 (1)***/75 (11)</td>
<td>121 (18)***/73 (12)</td>
<td>91 (14**)</td>
<td>89 (14**)</td>
</tr>
</tbody>
</table>

Data are mean (SD). *p = 0.01, **p = 0.05, ***p = 0.005, ****p = 0.001, †not significant, paired t test compared with pre-procedure data.

intentionally placed across its origin without clinical effect to the patient. In another patient the catheter shifted position during balloon inflation, resulting in stent deployment distal to the CoA site, and was redilated within the descending aorta to stabilise its position. Two additional stents were then placed across the CoA site, one of which overlapped the origin of the left subclavian artery. One patient had minor distal migration of the stent during placement, although its final position adequately covered the CoA. No change in left arm pressure was noted in either patient. The final patient had recurrent CoA after a surgical coarctectomy. This patient had a substantially calcified scar at the restenosis site, which prevented initial low pressure dilation. A stent was placed during the first procedure with improvement in the gradient from 50 mm Hg before to 20 mm Hg after implantation. Unfortunately, after expansion of the balloon, and perhaps because of irregularities in the vessel wall, the balloon ruptured and required surgical removal. This patient returned for a second procedure in which a high pressure balloon was used. The gradient was 27 mm Hg before the second procedure, 2 mm Hg after the procedure, and 9 mm Hg at follow up, although a small residual waist within the stent contour was still present.

Blood pressures immediately following implantation were 132 (33)/80 (19) mm Hg (mean 98 (23) mm Hg) proximal and 130 (34)/80 (19) mm Hg (mean 97 (24) mm Hg) distal to the CoA (table 1). This was a slightly greater ascending systolic than descending systolic pressure (p = 0.04), with no difference in ascending or descending mean pressures. The mean peak gradient was 3 (5) mm Hg, significantly less than before the procedure (p = 0.001 (fig 2), and the minimum diameter within the stent increased to 16.9 (2.6) mm (range 9–20 mm, p = 0.001) (fig 3). The ratio of the CoA site to the transverse arch diameters rose from 0.35 (0.2) (range 0.18–1.00, n = 25) to 0.92 (0.15) (range 0.62–1.13, n = 18) (p < 0.001).

PROCEDURE COMPLICATIONS

One patient with previous aortic valve replacement and prior coarctectomy had a right middle cerebral artery embolism immediately following the procedure, resulting in a minor residual neurological deficit complicated by seizures. In one patient the dilation balloon ruptured during stent inflation, described above. Other procedural issues with stent placement are commented on above.

CLINICAL FOLLOW UP

One patient had coil embolisation for a persistent arterial duct at the time of stent implantation. Six months after stent insertion, a clinically important aneurysm (> 2 cm) was noted associated with a residual gradient of 25 mm Hg. Angiographically, the stent had not fully expanded the associated hypoplastic isthmus. This patient’s CoA was among the smallest, being 3 mm in diameter initially, and with the stent dilated to 15 mm. Surgical removal of the stent with resection of the circumferential aneurysm and placement of an interposition graft were accomplished uneventfully.

The remaining 26 patients had clinical follow up at a mean of 1.8 (1.0) years (range 0.1–3 years). Average blood pressure was 130 (14)/74 (11) mm Hg, and clinical gradient was 4 (8) mm Hg (range 0–32 mm Hg). Seven patients were still receiving antihypertension treatment at their last assessment.

HAEMODYNAMIC ASSESSMENT

Follow up haemodynamic assessment was obtained in 18 of 26 patients 1.3 (0.5) years after the procedure. Blood pressure was 125 (21)/75 (11) mm Hg (mean 91 (14) mm Hg) proximal and 121 (18)/73 (12) mm Hg (mean 89 (14) mm Hg) distal to the stent (ascending systolic pressure greater than descending, p = 0.02, mean pressures not significantly different). The mean CoA gradient was 4 (6) mm Hg (range 0–26 mm Hg) (fig 2).

Minimum stent diameter was 17.2 (2.2) mm (range 13–20 mm) (fig 3). Aneurysms were present at the CoA site in three patients, one of
whom required surgery (noted above); the others had areas of aortic wall dyskinesia and are under clinical follow up (fig 4). Thirteen of the 18 patients with angiographic follow up had minor intimal hyperplasia (<1 mm) without luminal compromise. Additionally, in 13 of the 18 patients the proximal, distal, or both ends of the stent were not in contact with the aortic wall. We have been unable to detect any consequence of stent non-adherence; specifically there have been no distal embolic events and no evidence for obstruction to flow through the stent struts.

The mean diameter of the transverse arch, the minimum stent diameter, and the descending aorta diameter were 21.4 (2.8), 17.2 (2.2) and 26.1 (6.6) mm, respectively. The proximal mid and distal stent diameters were within 1 mm of each other. The mean transverse arch was 1.15 (0.41) and the descending aorta 1.52 (0.35) times the minimum stent diameter. Of 16 patients for whom measurements of the descending aorta were available, three had a descending aorta more than double the size of the minimum stent diameter and four had a descending aorta between 1.5 and 2.0 times the minimum stent diameter.

MANAGEMENT OF PRIMARY COMPARED WITH RECURRENT COA

Seven of the 26 patients were managed for recurrent CoA after prior surgery. Blood pressure before angioplasty was similar in the two groups (surgical patients 166 (20)/83 (17) mm Hg v non-surgical 164 (27)/87 (12) mm Hg), although there was a trend for more of the surgical patients to be on drugs (surgical 71% v non-surgical 55%). Haemodynamic measurement before implantation revealed the surgical group to have a lower peak gradient (29 (12) mm Hg v 51 (20) mm Hg in the non-surgical group, p = 0.003), the minimum obstructive diameter being larger for the surgical group (9.9 (2.2) mm v 5.4 (3.3) mm, p = 0.001). Immediately after implantation the gradient was the same in both groups (surgical 3.3 (8.2) mm Hg v non-surgical 3.0 (4.1) mm Hg, p = 0.57) as was the minimum stent diameter (surgical 16.4 (3.8) mm v non-surgical 17.1 (2.2) mm, p = 0.91). Late haemodynamic follow up data were available for five surgical and 13 non-surgical patients. Gradient and minimal stent diameters remained similar for both groups (gradient 2.6 (4.0) mm Hg v 5.2 (7.3) mm Hg, p = 0.50; diameter 17.6 (2.3) mm v 17.1 (2.3) mm, p = 0.83, in surgical and non-surgical patients, respectively).

Discussion

In the present study CoA balloon dilation with endovascular stent implantation resulted in a significant reduction in the CoA site gradient. At clinical follow up only seven of 26 patients were taking antihypertension drugs. The average blood pressure in all patients was 130 (14)/74 (11) mm Hg. Haemodynamic and angiographic assessment 1.3 (0.5) years after the procedure showed a mean CoA gradient of only 4 (6) mm Hg; aneurysms were detected in three patients, one of whom required surgery. The other two had minor areas of dyskinesia in the aortic wall.

Previous studies of balloon angioplasty in adults consistently showed an improvement in the CoA gradient. However, studies with more than 10 patients reported a residual mean gradient ranging from 8 (10) mm Hg to 18 (15) mm Hg, and residual gradients of >20 mm Hg were found in 7%-26% of patients. At follow up, clinical hypertension or continued use of antihypertension drugs was reported in 27%-68% of patients, and late haemodynamic restenosis (when reported) was found in 7%-9%. Other late complications were aneurysm formation in 7-15% and aortic dissection requiring urgent surgery in three patients.

To address these concerns, endovascular stents have been proposed as a potential technique to improve upon balloon angioplasty alone. As such, they have been applied in the setting of congenital heart disease for a number of indications including pulmonary arterial stenosis, and conduit and systemic venous obstructions with good sustained results. Previous evaluations of stent management of CoA have included case reports, and small series (n < 10) in which patients were often selected because their conditions were judged to be inappropriate for balloon angioplasty alone. These studies used smaller stents than were available for our study (often 3 cm length and over stretched diameter 18 mm). Nevertheless, case series have reported encouraging results compared with those reported for angioplasty alone. For many patients, there was no residual gradient and mean residual gradients were 1 (1.6) mm Hg in 2 (3) mm Hg, 4 (1) mm Hg, and 13.3 (23.2) mm Hg in four studies. In three studies with clinical follow up and a combined population of 25 patients, five remained hypertensive at follow up. Uniquely in our study, our population comprised consecutive patients presenting for management of CoA, all of whom were scheduled for clinical follow up and the majority of whom agreed to haemodynamic re-evaluation. It is important that the stent device inserted was substantially larger and longer (maximal rated diameter) than those used by others and that the majority of patients were adults.

Excellent immediate haemodynamic results were obtained in this study, with a reduction in...
Stent management for coarctation

CoA gradient to 3 (5) mm Hg and only one of 26 patients with a residual gradient of 20 mm Hg. Further, haemodynamic reassessment in 18 patients showed that the CoA site gradients remained low (mean 4 (6) mm Hg) with persistent but dilatable stenosis documented in only one of these patients. Persistent hypertension at clinical follow up was present in seven of 26 patients, less than has been reported for angioplasty alone.13 14 17 19 22 25 An- eyrsm remain a potential issue of concern and were found in three of 18 patients studied, one of whom required surgery. The morphology of the lesion and potential for aneurysm development in this latter patient deserves further comment. In a subset of patients with CoA (as in this patient), the obstructive element is an invagination of the wall of the vessel (fig 4) rather than an endoluminal shelf. Therefore, there is little or no vessel circumference for either dilation or stenting. These patients should undergo either serial progressive stent enlargement or primary referral to surgery. Discordant motion during the cardiac cycle between the ascending and descending aorta, seen angiographically, is the clue to this anatomical situation. Alternatively, application of intravascular ultrasound may define this higher risk subgroup, by defining the margins of the aortic wall. In this regard, intravascular ultrasound may also be of value in addressing additional issues such as the potential (although not observed in this study) for stent thrombus formation, particularly between stent struts that are not opposed to vessel wall.

The mechanism of minor wall irregularities, which may lead to clinically significant aneurysms, may be tears in the thin non-fibrous shelf region of the vessel wall. As the stent struts maintain an enlarged circumference with exposure of the thinned wall to arterial pressure, a wall irregularity beyond the limits of the stent may develop. Taking care in not over-distending the vessel—that is, stent dilation of the fibrous shelf to the limits of normal aortic wall—and the possibility of staged, sequential dilations over a period of several months may avoid this problem. Further studies are required to address these approaches. Anatomical proximity of the CoA to the left subclavian artery makes stent placement near or up to the ostium unavoidable. Angiographic and haemodynamic observations in instances where placement was close to or impinging upon the left subclavian ostium revealed no evidence of flow compromise. That this is not a surprise rests with the unobstructive character of flow through the stent struts.

Angiographic follow up has shown minor intimal hyperplasia in a majority of patients (13 of 18 in the present series). At present this does not have a haemodynamic effect and we believe that the intima is unlikely to proliferate to an extent that would cause obstruction in the future. That this may be the case is not surprising, given the high flow through the region and the large diameter achieved.22 This possibility will need to be re-evaluated over a longer follow up time. Proximal or distal stent non-adherence to the aortic wall was observed in 13

of the 18 patients who were re-evaluated angiographically. The significance of this observation is uncertain because no clinical consequence has been detected. Our observations did not define any anatomical subgroup in whom stenting was technically unachievable or not acutely effective. The noted procedural complications are avoidable with experience. Whether patients with CoA in the absence of an obstructing shelf should have sequential stent expansion or should not have the procedure at all has not been determined.

There are potential problems in the long term follow up of endovascular stents, which were not observed in this study. Specifically, there is the possibility of stent migration into the descending aorta, although because no migration was detected in the first year we think that this will be unlikely. Metal fatigue has not been observed because the implants were expanded well within their rated diameters. If metal fatigue were to occur, it may not result in an embolic complication because the majority of the stent is covered with endothelium. The possibility for device erosion is not known.

LIMITATIONS

Although this is the largest reported group of patients managed with stent implantation for CoA, this study was limited by the relatively small number of patients, making accurate evaluation for risk difficult and infrequent complications less likely to be observed. Similarly, the follow up time was short in relation to the patients’ expected lifespan, thus preventing observation of late complications, which may have important future consequences. There were two major limitations to the study design. Firstly, we did not compare the study population with a surgical cohort. A randomised comparison with surgical patients would be the optimal assessment but would require multicentre involvement to accomplish within a meaningful time. Such a study would be limited to centres that can offer expertise in both techniques. Secondly, we did not perform an exercise test evaluation during follow up. Exercise induced hypertension is frequently observed, with uncertain clinical significance. It would be of interest in the future to compare the effectiveness of stent techniques with surgical techniques in preventing exercise induced hypertension.

CONCLUSION

In summary we have shown that stent implantation as primary management for CoA is an effective intervention for gradient reduction with acceptable procedural risk. Application of this technique has significantly improved upon the problem of restenosis observed following angioplasty alone. The possibility for late complications has not yet been evaluated but is expected to be low. The noted alterations in wall configurations (small aneurysms) require careful observation to determine their static or progressive nature. We believe that stent implantation can be recommended as an acceptable alternative to surgery for CoA.
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