CASE REPORT

Myocardial ischaemia in a case of a solitary coronary ostium in the right aortic sinus with retroaortic course of the left coronary artery: documentation of the underlying pathophysiological mechanisms of ischaemia by intracoronary Doppler and pressure measurements

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Abstract

Only a few cases of a single coronary ostium and retroaortic course of the coronary artery have been described. Almost all cases reported so far had additional coronary artery or valvar disease. However, myocardial ischaemia may be caused by the coronary malformation alone. A 40 year old woman with severe myocardial ischaemia in the absence of clinically relevant coronary atherosclerosis is described. To clarify the origin and mechanisms of ischaemia, intracoronary Doppler, pressure and ultrasound studies were performed using microtransducers. In its outer portion along the course behind the ascending aorta, coronary blood flow velocities were increased, there was an external elliptical compression, and distal coronary flow reserve was reduced. Furthermore, an overshoot in diastolic pressure above aortic pressure was detectable within this portion. Dobutamine stimulation exaggerated the observed intracoronary haemodynamics and induced myocardial ischaemia. The intracoronary diagnostic procedures performed were helpful in clarifying the pathophysiological mechanisms of func-

tional coronary obstruction and ischaemia in this malformation. Bypass surgery was successfully performed with symptomatic improvement.

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A solitary coronary ostium in or immediately cephalad to the right aortic sinus and the course of the left coronary artery dorsal to the ascending aorta is a very rare congenital anomaly. This coronary malformation has been classified as the type IV.A.2a according to the classification of Roberts.¹ At present, only a few cases with this abnormality have been described in the literature. In most of these patients, the diagnosis was incidental either during angiography or necropsy. Most patients had significant concomitant coronary atherosclerosis causing symptoms of angina, or myocardial ischaemia and infarction.²⁻⁴ Even in the absence of severe coronary atherosclerosis, myocardial ischaemia can occur in cases of anomalous origin of the left coronary artery from the right sinus of Valsalva, if the left coronary artery passes posterior to the aorta.⁵ The pathophysiological mechanisms leading to myocardial ischaemia in the absence of coronary artery disease are incompletely understood. This is in contrast to other coronary anomalies where the left coronary artery arises from the right coronary artery and courses between the aorta and pulmonary artery (interarterial or intertruncal course). This has been associated with myocardial ischaemia and sudden death.⁶⁻¹⁰ The mechanism in this scenario is most likely a transient occlusion of the left coronary artery caused by an increase in blood flow through the aorta and the pulmonary artery, resulting in kinking or pinching of the artery.^{5,6} In contrast, in the abnormality presented here, coronary obstruction is usually absent on angiography.¹¹

Figure 1 (A) 12 lead ECG showing ST segment elevation in leads I, aVL, V2-V5 with ST segment depression in II and aVF directly after exercise testing, associated with severe symptoms of angina. (B) At rest, ST segment changes diminished and symptoms were alleviated.

Figure 2 Coronary angiogram ((A) left anterior oblique projection; (B) right anterior oblique projection) showing a solitary coronary ostium in the right aortic sinus and retroaortic course of the left coronary artery. Note the aorta (Ao) and the catheter in the pulmonary artery (PA). The arrow indicates the location of altered intracoronary haemodynamics compared with the proximal and distal segments, as confirmed by the position of the intravascular transducers angiographically. RCA, right coronary artery; LAD, left anterior descending coronary artery; CX, left circumflex coronary artery.

We present the first report describing myocardial ischaemia, and its potential pathophysiological mechanisms, in a patient with a left coronary artery originating from the right sinus of Valsalva coursing posterior to the aorta in the absence of clinically relevant coronary atherosclerosis.

Case report

A 40 year old woman reported a four week history of exercise induced angina pectoris; ST segment elevation was documented during bicycle ergometry in the anterior electrocardiographic leads (V2–V5) (fig 1). Immediately after exercise testing, torsade de pointes occurred. There was no history of coronary artery disease or hypertension. The patient had a history of cigarette smoking for a few years before admission. Physical examination revealed an otherwise healthy woman. Coronary angiography revealed a solitary ostium in the right aortic sinus associated with an aberrant retroaortic course of the left coronary artery (figs 2 and 3). There was a 30% atherosclerotic stenosis in the mid-portion of the right coronary artery. Despite several additional angiographic projections, it was impossible to demonstrate a significant phasic luminal narrowing of the left main within its retroaortic course.

Intracoronary Doppler flow velocity and pressure measurements using miniaturised 0.014 inch guidewire transducers and intravascular ultrasound were performed at rest and during dobutamine stimulation.^{12–16} Intravascular Doppler showed only slightly increased blood flow velocities along the curved retroaortic segment of the left main stem (arrow, fig 3), which were enhanced during intravenous infusion of dobutamine (5–20 µg/kg) (fig 4A), compared with measurements at rest (fig 4B) and with measurements at the proximal or distal portions of the left coronary artery (not shown). Distal coronary flow reserve, meas-

ured as the ratio of maximal flow velocity after stimulation with 12 mg of intracoronary papaverin to baseline flow velocity, was significantly reduced at 1.7.¹⁷

Intracoronary pressure measurements demonstrated a local overshoot in diastolic pressure (fig 5) at the site of accelerated flow velocity within the course of the left descending coronary artery. In vitro validation of pressure measurements within squeezed coronary segments using the 0.014 inch pressure microtransducer has recently been demonstrated.¹⁸ Intravascular ultrasound demonstrated only a slight systolic and diastolic reduction of cross sectional area at this portion (fig 4A) compared with a normal distal portion (fig 4B), without signs of atherosclerosis. Measurements during intravenous infusion of dobutamine demonstrated augmentation of blood flow velocities and of intravascular pressure (fig 5), and a further reduction in cross sectional area. Moreover, angina symptoms with ST segment elevation in the anterior leads occurred similar to those during exercise testing. In view of the severity of symptoms and objective evidence of

Figure 3 Schematic drawing of the coronary anomaly with the course of the left coronary artery posterior to the ascending aorta. AO, aorta; PA, pulmonary artery; LCA, left coronary artery; RCA, right coronary artery; CX, left circumflex coronary artery; LAD, left anterior descending coronary artery.

Figure 4 Intracoronary blood flow velocities (left) and intravascular ultrasound (right) within the left coronary artery at its sharp course dorsal the ascending aorta during dobutamine stimulation (A) and at rest (B). ADPV, average diastolic flow velocity; ASPV, average systolic flow velocity; APV, average peak flow velocity; DSVR, diastolic/systolic velocity ratio; BAPV, basic average peak flow velocity; PAPV, peak average peak flow velocity. RATIO, coronary flow reserve.

Figure 5 Intracoronary pressure measurements, obtained using a fiberoptic microtransducer incorporated in the distal segment of a 0.014 inch guidewire, show the overshoot in pressure within the retroaortic course of the left coronary artery compared with aortic pressure. To avoid artificial measurements, the guidewire was repeatedly rotated and its position angiographically confirmed. There was no visible external compression causing catheter entrapment. LCA, left coronary artery.

ischaemia and ischaemia induced malignant arrhythmias, the patient had aortocoronary bypass surgery with the left internal mammary artery to the left anterior descending coronary artery. Intraoperatively, the course of the left coronary artery behind the aorta was confirmed. No relevant stenosis of the left coronary artery along its course was evident. At eight weeks' follow up the patient was free from angina, and arrhythmias did not occur during or after exercise testing.

Discussion

Congenital coronary malformations are rare causes of angina, myocardial infarction or malignant arrhythmias.¹⁹ Usually, the diagnosis is incidental and clinical symptoms are attributed to other associated disorders, most commonly coronary artery disease. Five possible anatomical anomalies with a single coronary orifice in the right sinus of Valsalva with different clinical implications are known.

- The left coronary artery crosses between aorta and pulmonary artery (interarterial, intertruncal). This anomaly has been associated with sudden death from coronary obstruction during or shortly after exercise in otherwise healthy subjects⁵⁻¹⁰
- The left coronary artery crosses anterior to the pulmonary infundibulum, in which the left coronary artery is a branch of the conus branch of the right coronary artery. Myocardial ischaemia can occur, but is rare and its cause unclear⁵
- The left coronary artery has a septal (intermuscular, subpulmonic) course within the crista supraventricularis and the ventricular septum. Several case reports have been reported with this anomaly but without any clinical significance²⁰⁻²¹
- The left coronary artery crosses posterior to the atrioventricular valves, which is known as the most common course without clinical relevance
- The left coronary artery crosses posterior to the aorta (retroaortic), which is the case here, and in which myocardial ischaemia may occur but its mechanisms are unknown.

None of the mechanisms for myocardial ischaemia suggested above can be invoked in the present anomaly, as the retroaortic course of the left main stem would not explain external vessel compression or obstruction at its origin. In addition, none of the previously reported patients with this coronary anomaly had overt ischaemic symptoms or myocardial infarction (or scar tissue at necropsy) that could be related to the malformation. However, this report demonstrates the early onset of severe angina with anterior ST segment elevation and ischaemia induced malignant arrhythmias in a patient without significant coronary artery or other heart disease. Coronary angiography was unrevealing in terms of the cause of ischaemia. Intracoronary Doppler flow velocities at rest were only slightly increased; however, coronary flow reserve during hyperaemic provocation was significantly reduced indicating the presence of a functional obstruction. Dobutamine stimulation resulted in a further increase in flow velocities and ECG changes with significant ST segment depression and angina symptoms. Furthermore, use of an intracoronary pressure transducer made it possible to identify a localised pressure phenomenon that occurred at the most "external" point of the retroaortic course of the left main stem as illustrated in fig 3. The pressure curves were characterised by an overshoot in diastolic pressure above the aortic diastolic pressure. The existence of higher intramural than aortic or intraventricular pressures has been described before, and a similar pressure phenomenon with higher intracoronary than aortic pressures was recently validated in vitro and in patients with myocardial bridges.¹⁸⁻²²⁻²³ In these intramural courses of coronary artery segments, the overshoot in systolic pressure is most likely caused by a central high pressure segment connected to more compressed proximal and distal segments.

Intravascular ultrasound demonstrated an external phasic diastolic vessel compression producing an elliptical shaped vessel lumen that was most evident during dobutamine stimulation. This phenomenon did not occur in any other portion of the coronary artery.

The intracoronary haemodynamic alterations were most likely caused by the diastolic extension of the aortic bulbus and sinus of Valsalva compressing the left main coronary artery at the outer most portion of its retroaortic course. Most likely, the left coronary artery was unable to compensate for the expansion of the aortic bulbus by its own motion as the vessel was more or less fixed at the aortic origin and during its distal course. Intraoperatively, it appeared that the left main stem was compressed at the above described location, while the other segments of the left coronary artery were completely normal. It remains unclear why the patient was asymptomatic until age 40 and why symptom onset was relatively sudden; reduced flexibility of the elastic properties of the vessel wall by age might in part serve as an explanation.

In summary, we demonstrate a case of severe myocardial ischaemia in a patient with a single coronary orifice in the right sinus of Valsalva and a retroaortic course of the left coronary artery. Intravascular diagnostic procedures at rest and during pharmacological stimulation demonstrated the mechanisms leading to myocardial ischaemia in this rare coronary anomaly. Intracoronary Doppler, pressure and ultrasound measurements were clearly of benefit in this patient in diagnosis and hence in guiding treatment.

- 1 Roberts WC. Major anomalies of coronary arterial origin seen in adulthood. *Am Heart J* 1986;111:941-63.
- 2 Shirani J, Roberts WC. Solitary coronary ostium in the aorta in the absence of other major congenital cardiovascular anomalies. *J Am Coll Cardiol* 1993;21:137-43.
- 3 Lipton MJ, Barry WH, Obrez I, et al. Isolated single coronary artery: diagnosis, angiographic classification, and clinical significance. *Radiology* 1979; 130:39-47.
- 4 Kimbiris D, Iskandrian AS, Segal BL, et al. Anomalous origin of coronary arteries. *Circulation* 1978;58:606-15.
- 5 Bittl JA, Levin DC. Coronary arteriography. In: Braunwald E, ed. *Heart disease. A textbook of cardiovascular medicine*. 5th ed. Philadelphia: WB Saunders Company, 1997:235-75.
- 6 Cheitlin MD, De Castro CM, McAllister HA. Sudden death as a complication of anomalous left coronary origin from the anterior sinus of Valsalva. A not so minor congenital anomaly. *Circulation* 1974;50:780-7.
- 7 Joswig BC, Warren SE, Vieweg WV, et al. Transmural myocardial infarction in the absence of coronary arterial luminal narrowing in a young man with single coronary artery anomaly. *Cathet Cardiovasc Diagn* 1978;4:297-304.
- 8 Liberthson RR, Dinsmore RE, Fallon JT. Aberrant coronary artery origin from the aorta: report of 18, review of literature and delineation of natural history and management. *Circulation* 1979;59:748-54.
- 9 Kragel AH, Roberts WC. Anomalous origin of either the right or left main coronary artery from the aorta with subsequent coursing between aorta and pulmonary trunk: analysis of 32 necropsy cases. *Am J Cardiol* 1988;62:771-7.
- 10 Barth CW III, Roberts WC. Left main coronary artery originating from the right sinus of Valsalva and coursing between aorta and pulmonary trunk. *J Am Coll Cardiol* 1986;7:366-73.
- 11 Angelini P. Normal and anomalous coronary arteries: definitions and classification. *Am Heart J* 1989;117:418-34.
- 12 Doucette JW, Corl PD, Payne HM, et al. Validation of a Doppler guide wire for intravascular measurements of coronary artery flow velocity. *Circulation* 1992;85:1899-911.
- 13 Di Mario C, Meneveau N, Gil R, et al. Maximal blood flow velocity in severe coronary stenoses measured with a Doppler guidewire. *Am J Cardiol* 1993;71:54D-61D.
- 14 Ofili EO, Labovitz J, Kern MJ. Coronary flow velocity dynamics in normal and diseased arteries. *Am J Cardiol* 1993;71:3D-9D.

- 15 Emanuelsson H, Dohnal M, Lamm C, *et al.* Initial experiences with a miniaturized pressure transducer during coronary angioplasty. *Cathet Cardiovasc Diagn* 1991;24:137–43.
- 16 Abildgaard A, Oystein J, Klow NE. A pressure-recording guide wire for measurements of transstenotic gradients in arteries. In vitro validation. *Invest Radiol* 1994;29:178–83.
- 17 Gould KL, Kirkeeide RL, Buchi M. Coronary flow reserve as a physiologic measure of stenosis severity. *J Am Coll Cardiol* 1990;15:459–74.
- 18 Klues HG, Schwarz ER, vom Dahl J, *et al.* Disturbed intracoronary hemodynamics in myocardial bridging—early normalization by intracoronary stent placement. *Circulation* 1997;96:2905–13.
- 19 Chaitman BR, Lesperance J, Saltiel J, *et al.* Clinical, angiographic, and hemodynamic findings in patients with anomalous origin of the coronary arteries. *Circulation* 1976;53:122–31.
- 20 Schulte MA, Waller BF, Hull MT, *et al.* Origin of the left anterior descending coronary artery from the right aortic sinus with intramyocardial tunneling to the left side of the heart via the ventricular septum: a case against clinical and morphologic significance of myocardial bridging. *Am Heart J* 1985;110:499–501.
- 21 Roberts WC, Diccio BS, Waller BF, *et al.* Origin of the left main from the right coronary artery or from the right aortic sinus with intramyocardial tunneling to the left side of the heart via the ventricular septum: the case against clinical significance of myocardial bridge or coronary tunnel. *Am Heart J* 1982;104:303–5.
- 22 Stein PD, Marzilli M, Sabbah HN, *et al.* Systolic and diastolic pressure gradients within the left ventricular wall. *Am J Physiol* 1980;238:H625–30.
- 23 Sabbah HN, Stein PD. Effect of acute regional ischemia on pressure in the subepicardium and subendocardium. *Am J Physiol* 1982;242:H240–4.

SHORT CASES IN CARDIOLOGY

Cardiac valve invasion in chronic adult T cell leukaemia

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The cause of adult T cell leukaemia (ATL)¹ is human T cell leukaemia virus type I (HTLV-I), which was discovered in 1980.² This virus is endemic in southwestern Japan, the Caribbean, Africa, and South America. However, cardiac valve involvement by ATL cells is extremely rare³ and there is no previous echocardiographic evidence for this involvement. We present serial echocardiographic observations in a patient with chronic ATL in whom leukaemic involvement of the mitral valve showed pronounced improvement after chemotherapy.

A 58 year old woman was admitted because of two previous episodes of acute heart failure. On examination, generalised skin eruptions and lymphadenopathies were noted. Third and a fourth heart sounds (gallop rhythms) and a grade 3/6 pansystolic murmur were present at the apex. Transthoracic echocardiography

showed extensively thickened mitral and tricuspid valves with moderate regurgitations. The left ventricle was mildly dilated with reduced wall motion. Transoesophageal echocardiography showed an echogenic endocardial lesion at the left atrial wall and appendage, which extended continuously to the bizarre mitral valve abnormality (fig 1). Gallium scintigraphy showed a significant cardiac accumulation.

Serological tests were positive for anti-HTLV-I antibodies by particle agglutination and immunofluorescence. ATL cell infiltration was found in lymph node and skin biopsy specimens. Leukaemic cells from the lymph node expressed T cell markers (CD2+, CD3+, CD4+, CD7+, CD8-, and HLA-DR+). Together with echocardiographic and gallium scintigraphic findings, valvar invasion of tumour cells was suspected, possibly with myocardial involvement.

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Figure 1 Transoesophageal echocardiogram showing the bizarre mitral valve abnormality. (A) Echogenic endocardial lesion within the left atrium (arrows). (B) Pronounced improvement after chemotherapy.

Combination chemotherapy with cyclophosphamide, doxorubicin, vincristine, and prednisolone was started. After three chemotherapy sessions, the lesions in the mitral and tricuspid valves responded favourably with obvious reduction in thickness and improved flexibility in leaflet motion (fig 1). Repeated gallium scintigraphy demonstrated disappearance of the abnormal cardiac accumulation.

Despite these improvement the patient had an another episode of acute heart failure 10 months later and underwent mitral and aortic valve replacements. At surgery, the mitral valve was extremely thickened. Similar whitish thickenings were found in the left atrial wall and in the basal left ventricular wall. The diagnosis of valvar ATL cell infiltration was confirmed by histopathological examination of a surgically excised mitral valve.

This is the first echocardiographic documentation of cardiac valve involvement in ATL. The serial morphological changes of valvar lesions in response to chemotherapy have not been reported previously. This case has two important messages: first, it may at times be difficult to differentiate echocardiographically leukaemic valvar invasion from myxomatous valvar changes seen in patients with mitral valve prolapse, not only because of the similar-

ity of echocardiographic valvar features but also because of the rarity of leukemic valvar invasion; second, it is important to raise the possibility of involvement of the myocardium by ATL cells when the echocardiographic valvar abnormality is detected in the presence of positive HTLV-I serology, as myocardial involvement is probably more common as suggested by this case as well as by previous pathological reports.^{4 5} We propose ATL associated cardiac valvulopathy as one of the important signs of cardiac invasion by ATL cells that may be detectable echocardiographically.

- 1 Uchiyama T, Yodoi J, Sagawa K, *et al.* Adult T-cell leukemia: clinical and hematologic features of 16 cases. *Blood* 1977;50:481-92.
- 2 Poiesz BJ, Ruscetti FW, Gazdar AF, *et al.* Detection and isolation of type C retrovirus particles from fresh and cultured lymphocytes of a patient with cutaneous T-cell lymphoma. *Proc Natl Acad Sci USA* 1980;77:7415-19.
- 3 Gabarre J, Gessain A, Raphael M, *et al.* Adult T-cell leukemia/lymphoma revealed by a surgically cured cardiac valve lymphomatous involvement in an Iranian woman: clinical, immunopathological and viromolecular studies. *Leukemia* 1993;7:1904-9.
- 4 Roberts WC, Bodey GP, Wertlake PT. The heart in acute leukemia. A study of 420 autopsy cases. *Am J Cardiol* 1968; 21:388-411.
- 5 Roberts WC, Glancy DL, DeVita VT. Heart in malignant lymphoma (Hodgkin's disease, lymphosarcoma, reticulum cell sarcoma and mycosis fungoides). A study of 196 autopsy cases. *Am J Cardiol* 1968;22:85-107.