Is the best option of treatment being offered to patients with multiple coronary artery diseases?

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oronary arterial disease (CAD) is currently the most prevalent condition requiring cardiologist care. In spite of the progress in the clinical treatment of this disease over the last few years, there has been a change from clinical or surgical treatment to interventionist treatment, using several forms of angioplasty (PTA) [1]. To decide what is the best treatment method, it is necessary to considerer the strong evidence that single- and two-arterial CAD have similar outcomes, when comparing the clinical, interventionist and surgical treatments, but this is not true in relation to multiple vessel disease. Some facts must be discussed by patients suffering from multiple vessel diseases and a multidisciplinary team formed by a physician, hemodynamicist and surgeon so that a decision can be taken, not based on "cosmetology", but on proof based on medical evidence. With this in mind, the following aspects must be considered: a) the ischemia-free survival rate without re-interventions based on scientific works and b) the procedure cost over short, medium and long terms.

Databases have shown a reduction in the number of coronary artery bypass grafting surgeries (CABG) performed over the last few years with a drop of almost 20% comparing the total number of coronary artery surgeries performed in the USA in 1996 (182,911) and in 2004 (149,680) [2]. Simultaneously, the number of PTAs with or without the use of stents has increased almost geometrically [3].

In spite of having already observed a trend [4,5] in which PTAs have had an increase with subsequent reduction in the number of CABG, in Brazil there has only been a rise in the number of PTAs with the number of CABG remaining stable [6]. The question is: is it possible that this increase is due to criteria based on evidence or simply a tendentious choice? Why are more and more patients with tri-arterial diseases referred for a less effective treatment at a greater cost?

CABG is the most studied surgical procedure in medical history and the results have been widely reported over the last 10 and 20 years [7]. The surgery has been successful in reducing ischemia and increasing the survival rate in most groups of patients with multiple vessel diseases. With the technical improvements and forms of myocardial protection, results have achieved hospital survival rates of 98%. PTA, however, is a relatively new procedure with an enormous potential of change over the next few years but without an effective follow-up thus far. After its initiation using a balloon, PTA first developed with the use of plain stents and subsequently with drug-eluting stents, an evolution in which the devices presented with varying degrees of restenosis and early thrombosis [8].

To prove the "new" effectiveness of treatments with PTA, several studies have been performed, trying to demonstrate the superiority of this treatment in relation to the surgical treatment, with 6 performed with simple PTA (GABI [9], EAST [10], RITA [11], ERACI [12], CABRI [13], BARI [14]), five with the use of stents (MASS-2 [15], AWESOME [16], ERACI-2 [17], SoS [18], ARTS [19]) and another five with the use of drug-eluting stents (CARDia [20], SYNTAX [21], FREEDOM [22], COMBAT [23], Le MANS [24]).

However, the study groups used were low-risk groups in which CABG has little benefit and even so the surgical group included patients with "more severe" conditions, considering the number of arteries involved and the left ventricular function. Taggart [25], in his works, demonstrated two fundamental points; a) that the surgical treatment, in patients with lesions of one and two vessels who present with normal left ventricular function, does not increase the survival rate [26]; b) clinical trials comparing PTA and CABG, in spite of being, in the majority of cases, for single and two-lesion patients, present a greater re-intervention free survival rate for CABG and that groups in which surgery is recommended are not compared, including for multiple vessel disease and when the left ventricular function is compromised. This conclusion is evident in subgroups in which there is proximal involvement of the three main branches, in diabetic patients, when there is myocardial dysfunction and in those patients with injury to the left coronary arterial trunk [8,27]. Studies that compared CABG and PTA in patients with

equivalent multiple arterial lesions [28], the event-free survival rate, including acute myocardial infarction, heart-related death and any type of revascularization, always favored surgical revascularization. This study reaffirms a previous study by the same group, MASS-2 [15], in which on comparing both the event-free survival rate and the necessity of new interventions, the outcomes of the Surgical Group were better than the PTA Group and even the results of the Clinical Treatment Group were better than the PTA Group (Figures 1 and 2). In the same work, we can see the evolution of the three randomized groups over the first year, in which a statistical difference was demonstrated in respect to angina and the necessity of future CABG (Table 1).

We will not discuss here the biases of these works, which involved a group of patients of less than 5% of the original sample (of a series of 15 trails considering 8826 patients, only 441 were eligible to participate). When the results that portray the real world are analysed, the difference is more evident [29]. Taking into consideration the work by Yusuf et al. [26], for patients selected for comparative studies between



Fig. 1 – Event-free survival rate at one year, comparing CABG, PTA and clinical management. Figured taken from Hueb et al. [15]. Reproduced with the authorization of the Journal

Table 1. Evolution after a year of the three randomized groups.

CABG and PTA, there must be a great proportion of patients for which treatment with CABG is superior to clinical management.

Moreover, there is a necessity to review the concept of treatment in relation to its form [30]; while the approach to the disease in PTA is related to the "culprit lesion", in CABG it is related to the epicardial tissue of the artery. This fact added to the placement of a foreign body inside the lumen of the artery, contributes to high restenosis rates. An analysis of works that demonstrate the induction of a systemic inflammatory response on using coronary stents with restenosis as its first manifestation, contributes to the discussion on the physiopathology of this "new disease". Gomes et al. [31] demonstrated that, in spite of the introduction of covered stents having reduced intraluminal restenosis, the inflammatory response and restenosis at the edges remain. The same authors [32] affirm that CAD submitted to post-PTA CABG is a different disease to CABG without prior PTA and that the results may be different because of arteritis and, probably, associated myocarditis.

Another factor to be considered, when any type of



Fig. 2 – Re-intervention free survival rate at one year, between CABG, PTA and clinical management. Figure taken from Hueb et al.[15]. Reproduced with the authorization of the Journal

	CT (n = 203)	PTA (n = 205)	CABG (n = 203)	p-value
Death	3 (1.5%)	9 (4.5%)	8 (4.0%)	0.23
AMI with Q	10 (5.0%)	16 (8.3%)	4 (2.0%)	0.01
II or III (CCS) Functional Class	126 (63.6)	87 (45.3%)	75 (39%)	< 0.0001
Stroke	3 (1.5%)	2 (1.0%)	3 (1.5%)	0.29
CABG	12 (6.0%)	7 (3.5%)	_	< 0.0001
PTA	4 (1.97%)	18 (8.78%)	1 (0.5%)	0.008

CCS - Canadian Cardiovascular Society. AMI – acute myocardial infarct. CT-clinical treatment PTA – Angioplasty. CABG – Coronary artery bypass grafting. Table taken from Hueb et al.[15]. Reproduced with the authorization of the Journal

treatment is analysed, is the cost mainly in a country such as Brazil, in which 75% of patients are cared for by the government healthcare system. Previous works demonstrated that in spite of the initial cost being higher in CABG, after a year of follow-up, the PTA group becomes more expensive, not only because of the number of re-interventions, but also for the use of drug-eluting stents [33-35]. When the costs to the government healthcare system per patient for all types of PTAs and CABGs are taken in consideration, it can be seen that the cost of PTAs is 75% greater than CABG only with hospitalizations [4]. With this, it can be seen that in the USA, the use of the covered stents is 80% of the total [36], giving higher costs to the healthcare system, so that Medicare had to reduce its payments to reference centers by a third thereby reducing the quotas for companies that produce covered stents [37].

With all these data, it is difficult to understand, why the best treatment [1] for the lowest cost is not offered to patients.

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