Coronary Artery Perforation

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Abstract

Percutaneous coronary intervention (PCI) is considered as the standard treatment of obstructive coronary artery disease in indicated patients. Even though PCI gives symptomatic angina improvement, but associated with serious complications like coronary artery perforation (CAP), the incidence is quite low. With the more complex lesions for successful angioplasty, different devices are required, which in turn increase the incidence of CAP in these patients. Here we review the classification, incidence, pathogenesis, clinical sequela, risk factors, predictors, and management of CAP in the current era due to PCI.

Introduction

Percutaneous coronary intervention (PCI) for obstructive coronary artery diseases is accepted and has standardized procedure with minimal complication rates, including iatrogenic coronary artery perforation (CAP). Although angiographically significant coronary artery dissection is known to occur in up to 30% of all conventional balloon angioplasties,1,2 coronary perforation has been reported to occur in 0.3 to 0.6% of all patients undergoing PCI.3-6

Previous studies mentioned that predilatation before stenting predisposes for these complications, but subsequent studies disprove it. Increased incidence of CAP was reported with different coronary devices like atherectomy or rotablation, as more complex coronary lesions are being stented now. Here we review the incidence, causes, clinical sequela, and management of coronary perforation in the current era.7

Definition of Coronary Artery Perforation

CAP is defined as an anatomical breach in the wall of a coronary vessel due to the penetration of the three layers of the vessel wall, resulting in extravasation of blood or dye into the pericardium, myocardium, or adjacent cardiac chamber or vein.8

Consequences of Coronary Artery Perforation

Consequences of CAP depend on the location and severity. Location wise, if CAP occurs to the right or left ventricle, if not massive, then usually no immediate clinical consequences occur. If CAP occurs into the myocardium, myocardial hematoma occurs. If CAP occurs into the pericardium, then cardiac tamponade may occur. The severity of CAP was classified by Ellis et al (mentioned subsequently).

The mechanism of balloon angioplasty is by producing localized microdissections in the media and plaque fracture, not extending into the deeper layers of the arterial wall. CAP occurs when these dissections become extensive and penetrate through the vessel wall.

Incidence of Coronary Perforation

With standard simple PCI, the incidence of CAP is 0.1%,9,10 CAP incidence increase with the usage of GP IIb/IIIa inhibitors.11-14 Different incidences in different studies may be due to the difference in the definition of CAP, CTO intervention, and more aggressive debulking strategies for complex PCI success. Ajluni et al, in their large retrospective analysis, CAP reported in 0.4% of the PCIs.3

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Predictors and Causes of CAP

Mainly the factors which lead to CAP are:

1. Patient-related factors
2. Procedure-related factors
3. Adjuvant therapy-related factors
4. Lesion-related factors
5. Stent-related factors
6. Balloon-related factors (Details included in the Procedure-related factors)

1. Patient-Related Factors
Risk factors for CAP include female gender, old age, non-ST (stent thrombosis) MI (myocardial infarction), lesion complexity, chronic total occlusion (CTO) intervention, no. of stents, and hypertension. CAP reported was 46% in female versus 26% in males, which was statistically significant ($p = 0.001$). Increased incidence in women may be due to the old age group and small size of the coronary arteries. Additional risk factors include the lower baseline creatinine clearance, previous coronary artery bypass grafting (CABG),15 history of congestive heart failure, and multivessel coronary artery disease.16-19

2. Procedure-Related Factors
CAP can occur at different stages of the procedure, along with the gadgets used.

A. Type of guidewire and guidewire advancement.
B. Balloon/stent advancement.
C. Balloon/stent inflation.
D. Oversizing of the stent or ruptured balloon.
E. Improper position of the stent or balloon.
F. Type of balloon or stent.

A. Type of Guidewire and Guidewire Advancement
Heavy-weight and hydrophilic guidewires usage increases CAP incidence.20 Pressure wires used for fractional flow reserve (FFR) estimation are stiffer and less flexible wires than routine coronary wires, which require careful manipulation in complex or tortuous vessels. In Fig. 1, CAP occurred in proximal LCX when pressure wire was kept in LCX for equalization, before testing the FFR of left anterior descending artery (LAD) (Fig. 1).

During PCI, we have to pay attention to the distal tip of the guidewire position. Inadvertent advancement of guidewire more distally can lead to CAP. This type of excess free movement of guidewires occurs more frequently with hydrophilic wires.

B. Balloon/Stent Advancement
In tortuous and calcific lesions forcible advancement of the stent or the balloon can lead to CAP.

C. Balloon/Stent Inflation
Usually, either with the balloon or stent, dilation up to 1:1 ratio of the balloon or stent to an artery is considered ideal. Increase in this ratio is an important risk factor for CAP. The same thing was demonstrated by Ajluni et al in their study. According to this study, CAP occurred more frequently with the balloon to artery ratio of 1.3 ± 0.3 ($p < 0.001$). Similarly, in a registry by Ellis et al, the balloon to artery ratio of those patients undergoing percutaneous transluminal coronary angioplasty (PTCA) complicated by perforation was 1.19 ± 0.17 versus 0.92 ± 0.16 for those without perforation ($p = 0.03$). This observation has been confirmed in another large randomized study. There was a two to threefold increase in severe dissection leading to vessel occlusion when the ratio was more than 1.1. Also, balloon rupture, particularly those associated with pinhole leaks (as opposed to longitudinal tears), may create high-pressure jets that increase the risk of dissection or perforation.

Another situation where CAP can happen is during high-pressure inflation of the balloon in resistant coronary lesions. To prevent this complication, new noncompliant balloons, where we can dilate up to 35 atm (for example OPN NC balloons), are available.

D. Oversizing of the Stent or Ruptured Balloon
During oversized balloon inflation, if a rupture of the balloon occurs, then chances of CAP are more. In Fig. 2, the oversized proximal implanted stent balloon was used to dilate the distal lesion, which produced CAP.

E. Improper Position of the Balloon or Stent
If a stent or balloon is passed over the unrecognized subintimal passage of the wire, which can happen especially in CTO lesions, this may cause severe dissection or even
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The incidence of CAP is ~0.5 to 3% with the debulking devices like directional coronary atherectomy (DCA), excimer laser angioplasty, and rotational or extraction atherectomy. Even IVUS usage is also mentioned as one of the predictors of CAP, especially when intravascular ultrasound-guided PCI optimization was tried. In this case, Yukon stent of 3 × 18 mm is deployed in mid LAD lesion, and IVUS imaging was done to see the proper expansion of the stent. As there is under-expansion in the distal component of the stent on IVUS 3.25 NC balloon was used to dilate; immediate angiogram showed CAP which was controlled with prolonged balloon inflation and anticoagulation reversal (Fig. 3).

5. Stent-Related Factors

Stiff stents require high-pressure inflation for proper expansion, which can predispose for CAP. Another problem with stiff stents is that they are less traceable, so if used in the tortuous vessel again perforation chances increase.

Sites of CAP

Coronary perforations can be made into:

- Main vessel.
- Distal vessel.
- Branch vessel.
- Collateral vessel.

Management of main and distal vessel perforations is discussed in the below section. Usually, septal collateral perforation produces myocardial hematoma. If recognized early and the procedure abandoned, then there may not be any consequences, but this requires observation like other site perforations to see for the increase in hematoma, then compression effects may occur. If epicardial collateral perforation occurs, then blood may seep into the pericardium.

Types of Perforation

Ellis et al. evaluated a novel angiographic classification scheme for CAPs as a predictor of outcome. In a multicenter registry of 12,900 PCIs, 62 (0.5%) perforations were reported and categorized as:

- Type I: Extraluminal crater without extravasation.
- Type II: Epicardial fat or myocardial blush without contrast jet extravasation.
- Type III: Extravasation through frank (>1 mm) perforation or
Type III: “Cavity spilling” (CS) referring to Type III perforations with contrast spilling directly into either the left ventricle, coronary sinus, or another anatomic circulatory chamber.

The hypothesis that CAP may be the extension of the dissection is substantiated angiographically as the Ellis Type I perforation is identical to the previously described NHBLI (National Heart, Lung, and Blood Institute) Type C dissection. Angiographically we can classify CAP as:

- Free perforation—free contrast extravasation into the pericardium (Ellis Type III); or
- Contained perforation—when contrast staining is seen around the vessel without free contrast leak.

Clinical Outcome after Perforation

Clinical outcome mainly depends on the severity of perforation. CAP with cardiac tamponade is closely associated with mortality.28,29 Even though aggressive management was done at the time of perforation, the complication rates are high. MI occurred in 16.7 to 50%, emergency surgery in 50%, and death in 9 to 19%.28,29 Late complications like pseudoaneurysm are more frequently associated with DCA or cutting balloon usage.28

According to Ajluni et al, contained perforation (tamponade 6%, CABG 24%, death 6%) had lesser event rate than free perforations (tamponade 20%, CABG 60%, death 20%). The incidence of MI or death, tamponade in Type I, II, III, III CS were 0, 8%; 14%, 13%; 19%, 63%; 0, 0, respectively. Type III CS was associated with no event rate. Twenty percent of the CAP was due to guidewire, and 80% was during or after stent implantation.30

Management of Coronary Perforation

1. Balloon Inflation
First step, immediately after CAP is, appropriate size balloon to be inflated, either proximal to or at the CAP site, to occlude the vessel and thus prevents the further leakage of blood into the pericardium.

2. Reversal of Anticoagulation
Second, the reversal of the anticoagulation (especially heparin) with protamine is done. Our aim is to achieve an activated clotting time of less than 150 seconds.

A concern previously arose for anticoagulation reversal due to artery or stent thrombosis, which was disproved subsequently.31 In diabetic patients who were on protamine...
insulin injection, protamine administration should be avoided. If the patient is on GPI with abciximab before CAP, then it is better to give platelet concentrate. As tirofiban and eptifibatide GPIs have shorter half-lives, stoppage of those drugs is sufficient. There was no increased incidence of cardiac tamponade in those who received GPI. After acute CAP management, it is advisable to continue antiplatelet therapy, as this has resulted in rebleeding. Even in other studies, also GPI and bivalirudin were not shown as significant risk factors.

3. Pericardiocentesis
A. Early cardiac tamponade: If early cardiac tamponade is there, then pericardiocentesis is mandatory.
B. Delayed tamponade: Guidewire-associated CAPs and complex lesions are more likely present with delayed rather than early cardiac tamponade. Still, the need for surgical assistance is less (5%).

4. Prolonged Balloon Inflation
A little bit longer than CAP neck and equal size of the perforated artery balloon should be inflated at the CAP site for 10 minutes. The ischemic tolerance of the patient decides the duration of inflation and repeated inflations also need to be done till the perforation seals off or the ischemic duration tolerated by the patient. Many times, patients adapt to the more extended time of artery occlusion in subsequent dilations than in the first time occlusion time due to postischemic adaption.

To decrease the myocardial ischemia during long balloon inflation time, autoperfusion balloons can be used. This type of balloon allows blood to flow from the proximal segment of the inflated balloon from the side holes and the blood travels through the balloon, then perfuses through the distal segment of the coronary bed. Another method is microcatheter distal perfusion technique, in which microcatheter is used to perfuse the distal bed on another coronary wire.

Most of the time, Type I or II perforations can be managed with prolonged balloon inflation. However, we have to observe the patient even after initial stabilization for progression to tamponade or delayed tamponade, especially in guidewire-related perforations.

Longer duration balloon inflations were required, if the severity of the perforation is the higher grade (Type I vs. Type II: 44 ± 37 minutes vs. 21 ± 13 minutes, respectively; \( p < 0.05 \) and Type II vs. Type III: 48 ± 37 minutes vs. 20 ± 13 minutes; \( p < 0.05 \)). If the perforation is not sealed, then proceed to covered stent.

5. Covered Stents for Proximal to Mid Coronary Perforations
Covered stents are preferred modality of treatment when CAP is in proximal or mid coronary arteries. Covered stents which were introduced initially for coronary aneurysms, are very useful to treat the CAP. Two requisites for the covered stent usage are perforation should be in proximal or mid of the vessel, and distal wire should be in the true lumen.

The primary requisites for the usage of covered stents are the appropriate size of the perforated vessel, accessibility of the perforation site (in tortious and calcific vessels covered stent trackability becomes difficult), there should not be important big side branches, and the site of perforation should be very clear.

Covered stents from different companies are available with different materials. Symbiot stent by Boston scientific is made of double-layered polytetrafluoroethylene (PTFE) on a modified self-expanding nitinol stent. Jostent by Abbott company (covered stent) is made up of single PTFE layer in-between two coaxial stainless steel stents. Nuvasc stent-graft from Cardiovasc is made up of a single layer of PTFE coated with synthetic material P-15 on a stainless steel stent. P-15 is a cell adhesion protein, promotes the endothelialization.

The major drawback of the above-covered stents is the trackability. To improve the traceability, newly pericardial covered stents are designed. Venous covered stents were reported to be used in SVG perforations. Even though autologous vein graft stents are available, but to prepare them in an emergency situation is not practical. Papyrus stent has easy trackability as electrospun polyurethane membrane is used. Thin layers of polyethylene terephthalate of mesh stent is also another trackable stent.

Another problem with the covered stents is stent thrombosis in 5.7% and restenosis in 31.6% (angiographic) of the patients. So, it is advisable to give a longer duration of dual anti platelet therapy (DAPT)."
• So, now you have a “partially expanded” stent with a layer of “balloon” loaded on a wire. This newly prepared stent O has two layers: (1) metal outer layer and (2) balloon inner layer (Fig. 4d).

v. Now load stent I (3 × 28) on the newly prepared stent O with the help of the loading wire and position it so that partial part of stent I projects from the proximal and distal ends of the stent O (This is why stent I should be slightly larger than stent O) (Fig. 4e).

vi. Crimp stent O on stent I, thereby preparing a “sandwich stent,” which is ready for use. This sandwich stent will have three layers ([1] Metal inner layer, [2] Balloon middle layer, [3] Outer metal layer—as illustrated below) (Fig. 4f).

vii. Now load this newly prepared “covered stent” on the coronary guidewire and implant it with higher than nominal pressure across the perforated site. If there is still some leakage go to higher pressures, so as to get proper apposition of the newly prepared “covered stent.” (The purpose of taking a longer inner stent is to prevent “seepage of blood” at the edges of the outer layer. Moreover, the edges of the outer stent can be “post dilated” with another new balloon with a slightly higher pressure than nominal, for better apposition in case there is any leakage.)

(Important corollary: A similar “peripheral covered stent” can be done using two renal stents for peripheral artery perforations, in case peripheral covered stents are not available on the shelf. This has been successfully done in a case by the author also.)

7. Methods to Treat Distal CAP
For distal perforations we can use gel foam or metal coils. Other embolization materials described in the literature are coagulated blood from the patient, thrombin, two-component fibrin-glue, collagen, transcatheter subcutaneous tissue delivery, cyanoacrylate liquid glue, denatured alcohol, tris-acryl gelatin microsphere, or polyvinyl alcohol particles and use of a local drug delivery catheter.

a. Coils: Coils are a metallic wire with Dacron or wool as thrombogenic materials. We have to select the size of the coil, which should be bigger than the vessel perforated. The too big coil may dislodge in the proximal segment of the artery or too small one may embolize distally. These coils may be delivered through the guide catheter or more precisely exactly to the distal segment through microcatheter. Microcoils can be used for sealing of perforation without reversal of anti-coagulation. If required, they can be coated with fiber or gel for thrombogenicity. The microcoils are made of platinum, so they are more radiopaque with less risk of thrombosis.

b. Microspheres: These are hydrophilic nonabsorbable spherical particles. The size of these microspheres may range from 1 to 1,500μm and delivered to the site of perforation through microcatheter. These were tried mainly in collateral perforations. Safety of this method requires validation.

c. Thrombin injection: Thrombin promotes fibrin formation due to its platelet activator property. Solutions or glue with thrombin or fibrinogen has to be delivered to the perforation site with microcatheter carefully or over the balloon, to prevent the spillage of the material proximally.

d. Autologous blood clots: Major advantages of using autologous blood clots are easy availability, no cost, biocompatibility, and will be lysed automatically later. These blood clots are usually mixed with contrast media or saline, and then injected to the particular site.

e. Fat embolization: Autologous subcutaneous fat advantage is the same as an autologous blood clot. The mechanism of thrombus generation by this fat is, it causes a physical barrier and prevents the blood leakage, in addition to its thrombogenic property. This fat is usually mixed with contrast for radiopacity.

f. Miscellaneous: Other materials that have been used for embolization include synthetic glues, two-component adhesives made of fibrinogen and thrombin, collagen, polyvinyl alcohol particles, and protamine. Experience of the operator and availability of the embolic materials are the limitations in their usage for distal perforations.

8. Surgery
Surgical ligation of the vessel at perforation site and bypass graft to the distal vessel are required for severe CAP patients. When multiple stents are used along with subepicardial
hematoma then it is better to seal the perforation site with additional Teflon or pericardial patch.60

The algorithm to follow in CAP is mentioned in – Figs. 5 and 6.

Fig. 5 First algorithm for the management of coronary perfusion.

**Diagnosis of Coronary Perforation**

Coronary perforation can be easily diagnosed by coronary angiography and echocardiography, and it is usually...
accompanied by new episodes of chest pain, hemodynamic deterioration, and electrocardiographic changes.

Angiographic evidence of perforation is the presence of blush, jet, coronary sinus compression, and contrast in the pericardium.

When there is delayed post PCI hypotension then delayed tamponade should be suspected. This is more common in case of perforations induced by a guidewire or GP IIb/IIIa.37, 61 Repeat echo after 24 hours is mandatory to detect the delayed pericardial collection, especially in the causes of distal perforation and covered stent usage.62

**Contained Perforation and Pseudoaneurysm**

Another important complication of CAP of contained variety is pseudoaneurysm (local vessel dilation >1.5 times when compared with the normal adjacent artery) formation. This can happen as early as 10 minutes to as late as 2 to 3 weeks. Even though there is no literature on the incidence of these pseudoaneurysms rupture, this needs arfeful follow-up, and may require surgery.63 This is the case of pseudoaneurysm after the CAP, subsequently treated with a covered stent (►Fig. 7).

**Prognosis after Treatment of CAP**

Depending on the severity of vessel wall injury mortality also increases, it can be high (21.2%), and may result in periprocedural myocardial infarction in 34.0%. High all-cause mortality does not only occur in-hospital but also at 30 days (10.7%) and 1 year (17.8%).62,64 In a study, the in-hospital mortality was significantly high in tamponade group (7.7%) than without (4.3%). This tamponade group had a threefold increase in death on long-term follow-up.65 Mortality was found to be significantly higher in acute tamponade compared with delayed tamponade (59% vs. 21%, respectively; p = 0.04).66

This excess mortality may be related to underlying ischemia due to untreated coronary stenosis, side-branch loss with periprocedural MI, access site complications, major bleeding, and transfusion and risk of stent thrombosis, and restenosis with covered stent usage.15

**Conclusion**

CAP even though rare, is a dreaded complication of PCI. Best modality of treatment is prevention. This requires
early detection and immediate attention on the cath laboratory table to treat CAP immediately. The preferred sequence of steps to be taken in the management of the CAP depend on the type of CAP.

Contained perforation may have a relatively benign course immediately in hospital than free perforations. Immediately after CAP, prolonged balloon inflation at CAP site, anticoagulation reversal, and pericardiocentesis should be done in case of proximal or mid coronary vessel perforation. If no improvement, then covered stent should be placed. In distal perforations, it is better to use coils or above said alternatives. Two-dimensional echocardiogram plays an important role not only in the early tamponade but also requires repetition after 24 hours. If all these measures fail, then plan for CABG. CAP patients require meticulous long-term follow-up as there are chances of pseudoaneurysm formation, covered stent thrombosis or stenosis, and more cardiovascular event rates.

Conflict of Interest
None declared

References


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54 Goel PK. Delayed and repeated cardiac tamponade following microleak in RCA successfully treated with intra arterial sterile glue injection. Catheter Cardiovasc Interv 2009;73(6):797–800


