Peripheral venoarterial extracorporeal membrane oxygenation for periprocedural Cardiogenic shock during interventional cardiology

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ABSTRACT

Background: Cardiogenic shock (CS) is uncommon in the cardiac catheterization laboratory (CCL) among patients undergoing coronary angiography. Periprocedural CS is more frequent in high-risk patients and in technically demanding procedures. Aim: To describe the clinical outcomes of patients who underwent peripheral venoarterial extracorporeal membrane oxygenation (pVA-ECMO) for CS associated with interventional cardiology procedures. Material and Methods: Review of clinical records of seven patients treated between January 2014 and October 2018. Results: pVA-ECMO was implanted within 6 hours of the interventional cardiology procedure. All patients had coronary artery disease and one of them also had symptomatic severe aortic stenosis. One patient entered the CCL in cardiac arrest. Percutaneous coronary intervention (PCI) was performed in all patients; four patients underwent an emergency procedure and five patients experienced PCI complications. One patient undergoing transcatheter aortic valve replacement suffered acute severe aortic regurgitation. An intra-aortic balloon pump was inserted at the CCL in five patients. Six patients experienced cardiac arrest. Mean SAVE score was -4.3 and baseline lactate 55 mg/dl. pVA-ECMO mean duration was 5 ± 4 days. Survival after both hospital discharge and 12 months of follow-up was 85.7% Regarding vascular access complications, we observed one access site hematoma and one episode of cannulation site bleeding requiring surgical repair. Conclusions: pVA-ECMO should be considered in patients with periprocedural CS as a bridge to recovery. Its use was associated with improved clinical outcomes in this series.

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Key words: Cardiac Catheterization; Extracorporeal Membrane Oxygenation; Shock, Cardiogenic.

Oxigenación con membrana extracorpórea veno-arterial periférica para *shock* cardiogénico peri-procedimiento en cardiología intervencional

Antecedentes: El shock cardiogénico (SC) es infrecuente en el laboratorio de cateterismo cardíaco (LCC) entre pacientes que son sometidos a coronariografía. El SC peri-procedimiento es más frecuente en pacientes de alto riesgo y ¹Department of Cardiology, Instituto Nacional del Tórax. Santiago, Chile. ²Internal Medicine Department, Universidad de Chile. Santiago, Chile. ³Department of Cardiology, Hospital Militar de Santiago.

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en procedimientos técnicamente complejos. Objetivos: Describir los resultados clínicos de pacientes que fueron conectados a oxigenación con membrana extracorpórea veno-arterial periférica (ECMO-VAp) por SC peri-procedimiento de cardiología intervencional. Material y Métodos: Revisión de fichas clínicas de siete pacientes tratados en nuestro centro desde enero de 2014 a octubre de 2018. Resultados: ECMO-VAp fue utilizado dentro de las primeras 6 horas del procedimiento. Todos los pacientes tenían enfermedad coronaria y uno de ellos tenía además estenosis aórtica severa. Un paciente ingresó al LCC en paro cardíaco. Una intervención coronaria percutánea (ICP) fue realizada en todos los pacientes; 4 se realizaron procedimientos de emergencia y 5 pacientes tuvieron complicaciones de la ICP. A un paciente se le realizó un reemplazo valvular aórtico percutáneo y desarrolló una insuficiencia valvular aórtica aguda severa. Se instaló un balón de contrapulsación en el LCC en 5 pacientes. Seis pacientes tuvieron un paro cardiorrespiratorio. El valor del score de SAVE fue de -4,3 y el lactato basal 55 mg/dL. La duración media del ECMO-VAp fue 5 ± 4 días. La sobrevida al alta y a los 12 meses fue 85,7%. Como complicaciones del sitio de acceso vascular se observaron 1 hematoma y un episodio de sangrado que requirió reparación quirúrgica. Conclusiones: ECMO-VAp debería ser considerado en pacientes con SC peri-procedimiento como un puente a recuperación; su utilización estuvo asociada con mejoría de resultados clínicos en esta serie.

Palabras clave: Cateterismo; Choque Cardiogénico; Oxigenación por membrana extracorpórea.

Introduction

ardiogenic shock (CS) is uncommon in the cardiac catheterization laboratory (CCL), with an incidence of 0.24%-0.47% among patients undergoing coronary angiography^{1,2}. Periprocedural CS is more frequent in high-risk patients due to baseline cardiovascular disease and the technically-demanding procedures performed. Observational studies reveal an incidence of 3.5%-4.3% in patients with acute coronary syndromes^{3,4}, and a periprocedural CS incidence of 1.6% has been reported in intermediate-risk patients undergoing transcatheter aortic valve replacement (TAVR)⁵. Despite advances in percutaneous interventions and hemodynamic support, the prognosis for CS remains poor, with short-term mortality rates of 40%-60%^{2,6,7}.

Short-term mechanical circulatory support (MCS) aims to improve hemodynamic status and tissue perfusion to achieve recovery or provide time to establish a prognosis and definitive treatment. Consensus statements suggest that short-term MCS may be considered in patients experiencing periprocedural CS^{8,9}. Among MCS devices, peripheral venoarterial extracorporeal membrane oxygenation (VA-ECMO) has several

advantages, including easy installation, highly effective hemodynamic and respiratory support, compatibility with therapeutic hypothermia and low cost⁸.

This report describes the clinical outcomes of patients undergoing peripheral VA-ECMO for CS linked to interventional cardiology procedures at our center.

Patients and Methods

We conducted a retrospective observational study at a tertiary care center. The study included adult patients (\geq 18 years) undergoing interventional cardiology procedures who developed periprocedural CS and underwent peripheral VA-ECMO from January 2014 to October 2018. The study was approved by the Chilean Eastern Metropolitan Health Service ethics committee.

CS was defined according to the Society for Cardiovascular Angiography and Interventions (SCAI) consensus statement^{6,7}. Patients were retrospectively categorized into one of five CS stages at CCL admission, prior to VA-ECMO: At risk (A), Beginning (B), Classic (C), Deteriorating (D), Extremis or Refractory (E). The subscript "A" at any stage signifies cardiac arrest. Indications for peripheral VA-ECMO were at the discretion of the HEART team. Peripheral VA-ECMO was implanted by cardiac surgeons in the CCL or operating room (OR) within 6 hours of the interventional cardiology procedure in the case of deteriorating or refractory CS. Cannulation of femoral vessels was performed using a modified Seldinger technique, with 15-22 Fr arterial and 21-29 Fr venous cannulae, according to surgeon preference. Patients were postoperatively admitted to the ICU and treated per intensive care standards. VA-ECMO was removed in the OR by cardiac surgeons.

Clinical records were reviewed to record demographics, vital signs, laboratory data, clinical outcomes, and procedures performed during the hospital stay. Estimated in-hospital survival was assessed with the modified survival after VA-EC-MO (SAVE) score¹⁰. The Chilean Civil Registry database was reviewed for survival statistics.

Statistics

Continuous data are expressed as mean \pm standard deviation or median and categorical data as absolute number and percentage.

Results

Peripheral VA-ECMO was implanted in seven patients. Baseline characteristics are shown in Table 1. Mean age was 56.9 ± 16.9 years; 71.4% were male; 57.1% had hypertension; 42.9% had dyslipidemia; 42.9% had prior tobacco consumption. All patients had coronary artery disease. One patient had symptomatic severe aortic valve stenosis and severe stenosis in the middle segment of the left descending coronary artery and proximal segment of the circumflex artery, considered a candidate for elective TAVR and PCI after assessment by the HEART team. One patient entered the CCL in cardiac arrest (SCAI E_{a}); six patients (85.7%) were at risk for CS (SCAI A). Percutaneous coronary intervention (PCI) was performed in all patients. Four patients (57.1%) had emergency PCI due to ST-elevation myocardial infarction. Two patients (28.6%) with chronic coronary syndrome were scheduled for elective PCI for left main coronary artery bifurcation disease. Five patients (71.4%) experienced PCI complications: one dissection at the distal stent edge; one abrupt

left main coronary artery closure; one coronary perforation managed with a covered stent with later intraprocedural stent thrombosis; and two intraprocedural stent thromboses. One patient undergoing TAVR suffered acute severe aortic regurgitation. All complications were associated with hemodynamic instability and managed percutaneously during the procedure, achieving revascularization of target vessels or TAVR implant accordingly.

Upon hemodynamic deterioration, patients received vasopressor/inotropic medications, with a median of two drugs. An intra-aortic balloon pump (IABP) was inserted in the CCL in five patients (71.4%). Six patients (85.7%) experienced cardiac arrest, with a cardiopulmonary resuscitation time of 23 ± 12 minutes. Peripheral VA-EC-MO was implanted in the CCL in five patients (71.4%); in two cases as extracorporeal life support and in the others due to CS. Two patients (28.6%) completed PCI under VA-ECMO. Two patients (28.6%) had peripheral VA-ECMO implanted in the OR. Mean SAVE score was -4.3 ± 4.9 (median -5); baseline lactate 55 ± 36 mg/dl (median 36 mg/ dl); and modified SAVE score 6.4 ± 10.5 (median 7). VA-ECMO pump flow was 3.7 ± 0.6 L/min.

Table 2 summarizes clinical outcomes and complications. Peripheral VA-ECMO duration was 5 ± 4 days. Concomitant IABP was used in three patients (42.8%). At initial echocardiographic assessment, mean left ventricular ejection fraction was $38\% \pm 11\%$. Lactate clearance returned to normal values within 48 hours in six patients (85.7%) (Figure 1). Median invasive mechanical ventilation time, ICU and hospital stays were 14, 18 and 26 days, respectively. Survival to both hospital discharge and 12 months of follow-up was 85.7%. The deceased patient developed multi-organ failure and severe hypoxic-ischemic encephalopathy and was declared brain dead at 48 hours from VA-ECMO implant. One patient developed pulmonary edema refractory to adjustments in therapy, requiring bridging to central MCS (CentriMagTM) and transplant (performed after 86 days). The other patient with pulmonary edema was successfully weaned. Five patients (71.4%) had ventilator-associated pneumonia. Regarding vascular access complications, we observed one access site hematoma and one episode of cannulation site bleeding requiring surgical repair. Two patients required hemodialysis (28.6%).

Patient	1	2	3	4	5	6	7
Age (years)	51	68	37	40	86	59	59
Gender	Male	Male	Male	Female	Male	Female	Male
BMI (kg/m2)	27.9	31.9	24.2	24.2	25.5	21.2	25.7
Hypertension	Yes	Yes	No	No	Yes	No	Yes
Diabetes	No	No	Yes	No	No	No	No
Dyslipidemia	Yes	Yes	No	No	No	No	Yes
Tobacco use	No	No	No	Yes	No	Yes	Yes
Diagnosis pre-CCL	STEMI	CCS	STEMI	STEMI	SAVS	STEMI	CCS
CS stage pre-CCL	А	А	EA	А	А	А	А
PCI performed	Yes	Yes	Yes	Yes	Yes	Yes	Yes
PCI-treated vessel	LAD	LMCA	LAD	RCA	LAD/CX	LAD/CX	LMCA
PCI complication	Yes	Yes	No	Yes	No	Yes	Yes
TAVR performed	No	No	No	No	Yes	No	No
IABP pre-ECMO	Yes	Yes	Yes	No	No	Yes	Yes
CS stage pre-ECMO	D _A	E _A	EA	E _A	D _A	D	D _A
CPR time (minutes)	10	40	20	30	30	0	10
ECMO implant unit	CCL	CCL	OR	CCL	CCL	CCL	OR
ECLS	No	Yes	No	Yes	No	No	No
SAVE score	-5	-8	-10	0	-9	2	0
Lactate (mg/dl)	123	36	78	20	33	59	36
Modified SAVE score	-5	7	-10	15	6	17	15
ECMO flow (lpm)	4.2	2.9	3.7	4.3	2.8	3.7	4.0

Table 1. Baseline characteristics and procedural details of patients with periprocedural cardiogenic shock undergoing emergency peripheral VA-ECMO implant

BMI: Body mass index. CCL: Cardiac catheterization laboratory. STEMI: ST-elevation myocardial infarction. CCS: Chronic coronary syndrome. SAVS: Severe aortic valve stenosis. CS: Cardiogenic shock. PCI: Percutaneous coronary intervention. LAD: Left anterior descending artery. LMCA: Left main coronary artery. RCA: Right coronary artery. CX: Circumflex artery. TAVR: Transcatheter aortic valve replacement. IABP: Intra-aortic balloon pump. ECMO: Extracorporeal membrane oxigenation. CPR: Cardiopulmonary resuscitation. OR: Operating room. ECLS: Extracorporeal life support. Ipm: Liters per minute (at initiation).

Discussion

We describe an initial experience with peripheral VA-ECMO for periprocedural CS in a heterogeneous population undergoing interventional cardiology procedures. The use of peripheral VA-ECMO was effective in restoring systemic perfusion, allowing for completion of procedures; bridge to recovery, transplantation or decision and overall improved clinical outcomes.

Patients with periprocedural CS subjected to peripheral VA-ECMO had in-hospital and 12-month survival rates of 85.7% in our series. Prior studies in patients undergoing interventional cardiology procedures have described lower in-hospital survival rates despite implementation of VA-ECMO programs in the CCL. In patients with acute myocardial infarction experiencing refractory CS, several cases series reported survival rates of 22%-65% after peripheral VA-ECMO¹¹⁻¹⁷. In patients suffering hemodynamic complications during TAVR, rescue VA-ECMO support produced in-hospital survival rates of 44%-75%¹⁷⁻²⁰. Differences in survival might be partially explained by variability in definitions of refractory CS and associated multi-organic failure. This problem

Patient	1	2	3	4	5	6	7
ECMO duration (days)	7	2	2	12	2	4	5
IABP duration (days)	0	3	2	0	0	2	0
LVEF (%)	40	56	35	40	46	25	27
IMV duration (days)	36	3	2	15	4	14	30
ICU stay (days)	40	5	2	15	4	14	98
Hospital stay (days)	56	8	2	31	11	26	98
Pulmonary edema	Yes	No	No	No	No	No	Yes
Hospital-acquired infection	Yes	No	No	Yes	Yes	Yes	Yes
Neurologic complication	No	No	Yes	No	No	No	No
Vascular complication	Yes	No	No	Yes	No	No	No
Acute kidney injury	Yes	No	Yes	No	No	No	Yes
Hemodialysis	Yes	No	Yes	No	No	No	No
Outcome	CR	CR	Death	CR	CR	CR	HT
In-hospital survival	Yes	Yes	No	Yes	Yes	Yes	Yes
12-month survival	Yes	Yes	No	Yes	Yes	Yes	Yes

 Table 2. Evolution, clinical outcomes and complications of patients with periprocedural cardiogenic

 shock undergoing peripheral VA-ECMO implant

ECMO: Extracorporeal membrane oxigenation. IABP: Intra-aortic balloon pump. LVEF: Left ventricular ejection fraction (by Simpson's biplane). IMV: Invasive mechanical ventilation. ICU: Intensive care unit. CR: Cardiac recovery. HT: Heart transplant.

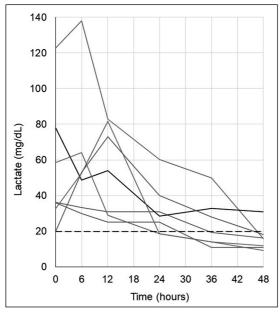


Figure 1. Evolution of lactate levels during the first 48 hours of hemodynamic support with peripheral VA-ECMO. Continuous grey lines show patients achieving survival to discharge (n = 6). Continuous black line marks the deceased patient (n = 1). Dashed black line marks the normal upper limit of lactate at 19.8 mg/dl.

could be solved by implementing the SCAI CS classification⁶. Jentzer et al. validated the risk stratification capacity of SCAI CS stages in patients admitted to a cardiac ICU (n = 10,004), showing good prognostic discrimination for in-hospital mortality (AUC 0.78)⁷. Extrapolating their results, our expected in-hospital mortality for a median SCAI CS stage D would have been approximately 60%.

Lactate prior to VA-ECMO seems to be valuable for clinical outcome prediction. Series of patients with acute myocardial infarction experiencing refractory CS with lactate levels over 70 mg/dl describe in-hospital survival rates of 35%-50% after VA-ECMO^{10,14-17}. The modified SAVE score, which incorporates lactate level at a cutoff of > 75 mg/dl, has enhanced prediction of adverse outcomes in patients with refractory CS undergoing VA-ECMO (AUC = 0.84)¹⁰. Our expected survival according to this score was 52%, lower than observed results. The modified SAVE score was validated in patients with lactate between 56 and 125 mg/dl, perhaps loosing accuracy in deteriorating CS. A study by Hryniewicz et al. in patients with refractory CS reported a mean of 3.4

vasoactive agents, median lactate of 39.6 mg/dl and in-hospital survival of 65% after VA-ECMO¹². Our series was similar in terms of baseline characteristics and lactate levels, but with a mean of only 2.3 vasoactive medications and more favorable outcomes. We concur with Hryniewicz et al. that early VA-ECMO implantation in patients with refractory CS, prior to higher lactate increases, could improve outcomes.

All patients experienced at least one complication associated with peripheral VA-ECMO. In terms of peripheral VA-ECMO complications in the context of periprocedural CS, previous studies report vascular complications requiring surgery in 11%-37%, bleeding related to ECMO in 8%-34%, neurologic complications in 9%-32% and dialysis requirement in 30%-46% of patients. Our results were similar for the aforementioned complications. Interestingly, installation by experienced operators in the CCL does not seem to produce a higher rate of vascular complications as compared to surgical implantation in the OR¹¹.

Limitations

Study limitations include the small number of patients and retrospective design, which preclude further analysis. There was inconsistent availability of some variables, including time to pump installation and transfusions. Comparisons with other types of MCS were not feasible, given the concomitant IABP and lack of availability of other devices.

Conclusion

Peripheral VA-ECMO was associated with improved clinical outcomes in this series. This procedure should be considered in patients with periprocedural CS as a bridge to recovery or decision. Prospective multicenter registries evaluating the safety, efficacy or futility of MCS in different settings are of significant interest.

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