

Catheter directed therapies: an option for elderly frail patients with pulmonary embolism requiring reperfusion

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Current guidelines of the European Society of Cardiology indicate that the optimal management of acute pulmonary embolism (PE) depends on the risk assessment of PE-related early mortality¹. Reperfusion therapy, preferably systemic thrombolysis, is still recommended as the first-line treatment in haemodynamically unstable patients. It should also be considered as a rescue therapy when initially normotensive subjects deteriorate despite adequate anticoagulation¹. However, real life shows that more than half of high-risk PE patients do not receive systemic thrombolysis because of increased risk of bleeding². Since the rate of major extracranial bleeding reached 11.1% in patients above 75 years of age in the PEITHO study, advanced age and frailty are often regarded as relative contraindications for systemic thrombolysis³. In such cases, surgical embolectomy or catheter-directed therapy should be considered as a primary or rescue reperfusion¹. However, surgical embolectomy is not usually immediately available. Thus, with growing clinical experience, catheter-directed therapies have become a real therapeutic option for PE patients⁴.

In this issue of EuroIntervention, Farmakis et al present the results of the US Nationwide Inpatient Sample from 2016 to 2020, including over 980,000 hospitalisations of PE patients aged ≥ 65 years (28.0% were frail)⁵. Reperfusion therapies were used in 4.9% of them, with an increase from 4.0% in 2016 up to 5.6% in 2020. Interestingly, the use of systemic thrombolysis and surgical embolectomy remained stable, while the application of percutaneous techniques almost doubled (from 1.7% to 3.2%). Although

high-risk PE is a strong indication for urgent reperfusion therapy, such treatment was used only in 1 out of 6 haemodynamically unstable patients, with no significant increase over a 5-year period. Moreover, frailty additionally limited the utilisation of reperfusion therapy in elderly patients with high-risk PE. These observations indicate that the treatment of elderly high-risk PE patients definitively remains an Achilles' heel in PE management and should be improved. Importantly, Farmakis et al give us some hints. Among frail high-risk patients, catheter-directed thrombolysis when compared to systemic thrombolysis was associated with reduced major bleeding (odds ratio [OR] 0.51, 95% confidence interval [CI]: 0.41-0.63) and even with a lower in-hospital mortality rate (OR 0.36; 95% CI: 0.28-0.46). Moreover, intracranial haemorrhage, which is the most feared and devastating complication of lytic treatment, occurred in only 1% of patients treated with percutaneous techniques, while in the group treated with systemic thrombolysis, it reached 3%. This indicates that catheter-directed therapies may be a safer option for elderly, frail PE patients requiring reperfusion and might even suggest that such therapies may be even more effective than systemic thrombolysis or surgical embolectomy.

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Although the clinical evidence for the use of percutaneous techniques is rapidly increasing^{6,7}, we need more high-quality evidence on catheter-based therapies performed with different devices in patients with different clinical characteristics including

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different PE risk categories. Moreover, local pulmonary embolism response teams should be established to guide the therapy for PE patients^{1,4,8}. Luckily, the results of currently ongoing randomised trials on percutaneous therapies for PE are expected in 2024, helping to determine the specific groups that could benefit from the various techniques^{9,10}.

Conflict of interest statement

The authors have no conflicts of interest to declare.

References

1. Konstantinides SV, Meyer G, Becattini C, Bueno H, Geersing GJ, Harjola VP, Huisman MV, Humbert M, Jennings CS, Jimenez D, Kucher N, Lang IM, Lankeit M, Lorusso R, Mazzolai L, Meneveau N, Ni Ainle F, Prandoni P, Pruszczyk P, Righini M, Torbicki A, Van Belle E, Zamorano JL; ESC Scientific Document Group. 2019 ESC Guidelines for the diagnosis and management of acute pulmonary embolism developed in collaboration with the European Respiratory Society (ERS). *Eur Heart J*. 2020;41:543-603.
2. Keller K, Hobohm L, Ebner M, Kresoja KP, Munzel T, Konstantinides SV, Lankeit M. Trends in thrombolytic treatment and outcomes of acute pulmonary embolism in Germany. *Eur Heart J*. 2020;41:522-9.
3. Meyer G, Vicaut E, Danays T, Agnelli G, Becattini C, Beyer-Westendorf J, Bluhmki E, Bouvaist H, Brenner B, Couturaud F, Dellas C, Empen K, Franca A, Galie N, Geibel A, Goldhaber SZ, Jimenez D, Kozak M, Kupatt C, Kucher N, Lang IM, Lankeit M, Meneveau N, Pacouret G, Palazzini M, Petris A, Pruszczyk P, Rugolotto M, Salvi A, Schellong S, Sebbane M, Sobkowicz B, Stefanovic BS, Thiele H, Torbicki A, Verschuren F, Konstantinides SV; PEITHO Investigators. Fibrinolysis for patients with intermediate-risk pulmonary embolism. *N Engl J Med*. 2014;370:1402-11.
4. Pruszczyk P, Klok FA, Kucher N, Roik M, Meneveau N, Sharp ASP, Nielsen-Kudsk JE, Obradovic S, Barco S, Giannini F, Stefanini G, Tarantini G, Konstantinides S, Dudek D. Percutaneous treatment options for acute pulmonary embolism: a clinical consensus statement by the ESC Working Group on Pulmonary Circulation and Right Ventricular Function and the European Association of Percutaneous Cardiovascular Interventions. *EuroIntervention*. 2022;18:e623-38.
5. Farmakis I, Barco S, Giannakoulas G, Keller K, Valerio L, Tichelbäcker T, Partovi S, Ahrens I, Konstantinides S, Hobohm L. A Nationwide Registry of reperfusion Therapies for Pulmonary Embolism in Older Patients with Frailty. *EuroIntervention*. 2023;19:772-81.
6. Sista AK, Horowitz JM, Tapson VF, Rosenberg M, Elder MD, Schiro BJ, Dohad S, Amoroso NE, Dexter DJ, Loh CT, Leung DA, Bieneman BK, Perkowski PE, Chuang ML, Benenati JF; EXTRACT-PE Investigators. Indigo Aspiration System for Treatment of Pulmonary Embolism: Results of the EXTRACT-PE Trial. *JACC Cardiovasc Interv*. 2021;14:319-29.
7. Toma C, Bunte MC, Cho KH, Jaber WA, Chambers J, Stegman B, Gondi S, Leung DA, Savin M, Khandhar S, Kado H, Koenig G, Weinberg M, Beasley RE, Roberts J, Angel W, Sarosi MG, Qaqi O, Veerina K, Brown MA, Pollak JS. Percutaneous mechanical thrombectomy in a real-world pulmonary embolism population: Interim results of the FLASH registry. *Catheter Cardiovasc Interv*. 2022;99:1345-55.
8. Kopec G, Araszkiwicz A, Kurzyrna M, Slawek-Szmyt S, Stepniewski J, Roik M, Darocha S, Golebiowski M, Jaguszewski M, Jankiewicz S, Kaluzna-Oleksy M, Kuliczowski W, Lewicka E, Mularek-Kubzdela T, Pietrasik A, Protasiewicz M, Przybylski R, Pleskot P, Tycinska A, Zielinski D, Podolec P, Trzeciak P, Grygier M, Mroczek E, Pruszczyk P. Role of catheter-directed therapies in the treatment of acute pulmonary embolism. Expert opinion of the Polish PERT Initiative, Working Group on Pulmonary Circulation, Association of Cardiovascular Interventions, and Association of Intensive Cardiac Care of the Polish Cardiac Society. *Kardiol Pol*. 2023;81:423-40.
9. Klok FA, Piazza G, Sharp ASP, Ni Ainle F, Jaff MR, Chauhan N, Patel B, Barco S, Goldhaber SZ, Kucher N, Lang IM, Schmidtman I, Sterling KM, Becker D, Martin N, Rosenfield K, Konstantinides SV. Ultrasound-facilitated, catheter-directed thrombolysis vs anticoagulation alone for acute intermediate-high-risk pulmonary embolism: Rationale and design of the HI-PEITHO study. *Am Heart J*. 2022;251:43-53.
10. Gonsalves CF, Gibson CM, Stortecky S, Alvarez RA, Beam DM, Horowitz JM, Silver MJ, Toma C, Rundback JH, Rosenberg SP, Markovitz CD, Tu T, Jaber WA. Randomized Controlled Trial of Mechanical Thrombectomy Versus Catheter-directed Thrombolysis for Acute Hemodynamically Stable Pulmonary Embolism: Rationale and Design of the PEERLESS Study. *Am Heart J*. 2023;266:128-37.