American College of Radiology
ACR Appropriateness Criteria®
Management of Acute Pulmonary Embolism

Variant: 1 Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.

Procedure	Appropriateness Category
Anticoagulation	Usually Appropriate
Systemic thrombolysis	Usually Appropriate
Catheter-directed therapy pulmonary artery	Usually Appropriate
Surgical embolectomy pulmonary artery	Usually Appropriate
Extracorporeal membrane oxygenation	May Be Appropriate

Variant: 2 Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.

Procedure	Appropriateness Category
Anticoagulation	Usually Appropriate
Catheter-directed therapy pulmonary artery	Usually Appropriate
Surgical embolectomy pulmonary artery	May Be Appropriate (Disagreement)
Systemic thrombolysis	Usually Not Appropriate
Extracorporeal membrane oxygenation	Usually Not Appropriate

Variant: 3 Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

Procedure	Appropriateness Category
Anticoagulation	Usually Appropriate
Catheter-directed therapy pulmonary artery	Usually Not Appropriate
Extracorporeal membrane oxygenation	Usually Not Appropriate
Surgical embolectomy pulmonary artery	Usually Not Appropriate
Systemic thrombolysis	Usually Not Appropriate

Variant: 4 Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.

Procedure	Appropriateness Category
Anticoagulation	Usually Appropriate
Catheter-directed therapy pulmonary artery	Usually Not Appropriate
Systemic thrombolysis	Usually Not Appropriate
Extracorporeal membrane oxygenation	Usually Not Appropriate
Surgical embolectomy pulmonary artery	Usually Not Appropriate

Variant: 5 Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial

hemorrhage. Initial therapy.

Procedure	Appropriateness Category
Catheter-directed therapy pulmonary artery	Usually Appropriate
Extracorporeal membrane oxygenation	May Be Appropriate
Surgical embolectomy pulmonary artery	May Be Appropriate (Disagreement)
Anticoagulation	May Be Appropriate
Systemic thrombolysis	Usually Not Appropriate

Variant: 6 Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.

Procedure	Appropriateness Category
Anticoagulation	Usually Appropriate
Catheter-directed therapy right heart	Usually Appropriate
Surgical embolectomy right heart	Usually Appropriate
Systemic thrombolysis	May Be Appropriate
Extracorporeal membrane oxygenation	May Be Appropriate

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Summary of Literature Review

Introduction/Background

Acute pulmonary embolism (PE) is a subclass of venous thromboembolism (VTE), in which blood clots typically formed in the deep veins of the extremities embolize to the pulmonary arterial circulation. In 2019, there were an estimated 1,036,000 documented cases of VTE in the United States, of which 393,000 involved PE [1]. Risk factors for VTE include inheritable thrombophilias, such as Factor V Leiden, or acquired risk factors, such as recent surgery, trauma, malignancy, pregnancy, hormone therapy, aging, or obesity [2]. During the COVID-19 pandemic, increased VTE and PE prevalence was reported in patients hospitalized with COVID-19 infection [3].

For a discussion of the role of imaging in the diagnosis of PE, please refer to the separate ACR Appropriateness Criteria® topic on "Suspected Pulmonary Embolism" [4]. Once a diagnosis of PE is established, risk stratification is commonly performed to help guide subsequent management. These algorithms typically distinguish 3 categories of PE severity: 1) massive or high-risk, 2) submassive or intermediate risk, and 3) low risk [5-7]. Multidisciplinary PE response teams have emerged to help streamline patient management and coordinate care [8,9].

Initial Therapy Definition

Initial therapy is defined as a first-line treatment option for the medical condition defined by the variant. More than one option can be considered usually appropriate as the initial therapy when:

- There are equivalent alternatives (ie, only one option will be planned to effectively manage the patient's care).

OR

- There are complementary therapies (ie, more than one treatment option is planned to be performed simultaneously or in sequence during the same setting, wherein the therapies provide synergistic or complementary benefits to effectively manage the patient's care).

Discussion of Procedures by Variant

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.

This variant depicts a patient with massive or high-risk PE. The criteria for massive/high-risk PE as established by the American Heart Association (AHA) is sustained hypotension (systolic blood pressure <90 mm Hg for at least 15 minutes) or need for vasopressor support [6]. The immediate goal of intervention is to reduce pulmonary arterial clot burden and to relieve right heart strain.

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.
A. Anticoagulation

Prompt initiation of systemic anticoagulation is considered standard of care for massive/high-risk PE [10]. Although the goal of anticoagulation is to neutralize the clotting cascade and prevent the propagation of thrombus, anticoagulation does not have a direct impact on reducing the existing clot burden, which is the objective of the subsequently described therapies.

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.
B. Catheter-directed therapy pulmonary artery

Catheter-directed therapy (CDT) can involve a combination of mechanical and/or pharmacologic endovascular techniques targeted at reducing the existing PE burden. Many of the published clinical trials evaluating the use of CDT have focused on intermediate/submassive risk PE and are discussed to greater detail in the subsequent variant. Although fewer in number, trials including high-risk PE do support the use of catheter-directed techniques in appropriately selected patients within this cohort. The Pulmonary Embolism Response to Fragmentation, Embolectomy, and Catheter Thrombolysis (PERFECT) multicenter registry [11] and the Single-Arm, Multicenter Trial of Ultrasound-Facilitated, Catheter-Directed, Low-Dose Fibrinolysis for Acute Massive and Submassive Pulmonary Embolism (SEATTLE II) trial [12] included patients with high-risk or intermediate-risk pulmonary emboli. These trials published data supporting the use of catheter-directed delivery of thrombolytic agents directly into the pulmonary arterial system, with the SEATTLE II trial demonstrating a decrease in thrombus burden, improvement in right heart strain, and reduced pulmonary hypertension [12]. The FlowTriever All-Comer Registry for Patient Safety and Hemodynamics (FLASH) registry, a multinational, multicenter registry of patients with high- and intermediate-risk PE treated with the FlowTriever mechanical thrombectomy system, has reported a 7.6 mm Hg mean drop in mean pulmonary artery pressure (-23.0%; $P < .0001$) and decreased right ventricular (RV)/left ventricular (LV) ratio (1.23-0.98; $P < .0001$) at 48 hours posttreatment, with a favorable adverse event risk profile [13]. The Higher-Risk Pulmonary Embolism Thrombolysis

(HI-PEITHO) multinational multicenter randomized control trial, targeted to complete patient enrollment in December 2023, will evaluate the clinical benefits of ultrasound (US)-facilitated catheter-directed thrombolysis with anticoagulation versus anticoagulation alone in high- to intermediate-risk PE, including primary outcome PE-related mortality as well as quality of life indicators and functional status over a 12-month follow-up period [14].

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.
C. Extracorporeal membrane oxygenation

In high-risk PE with hemodynamic instability, mechanical circulatory support via venoarterial extracorporeal membrane oxygenation (ECMO) is an emerging therapy that is gaining traction in the management of these critically ill patients. This may be considered as a rescue therapy in patients with refractory cardiogenic shock despite previous treatment with thrombectomy or thrombolysis [15].

Treatment with ECMO may also serve a role as a primary or standalone therapy in highly unstable patients presenting with profound shock and/or requirement for cardiopulmonary resuscitation. In an institutional retrospective review of 17 patients over a 2-year duration, 77% of survivors of high-risk PE received ECMO and anticoagulation alone [16]. Furthermore, a separate retrospective review of 83 patients did not demonstrate a statistically significant difference in survival to discharge for patients receiving systemic thrombolysis before ECMO versus those receiving ECMO alone (88.9% versus 84.6%; $P = .94$) [17].

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.
D. Surgical embolectomy pulmonary artery

Although surgical embolectomy has traditionally been considered a salvage therapy in the setting of failed attempts at the management of high-risk/massive PE, modern surgical techniques to promptly remove the thrombus have been demonstrated to be associated with a lower rate of bleeding complications as compared with thrombolysis [18]. Furthermore, a meta-analysis review of 17 studies comparing mechanical reperfusion strategies (of which 85.9% was surgical embolectomy) to other strategies, including systemic thrombolysis, CDT, or standalone ECMO, found a significant reduction in mortality in the mechanical reperfusion group compared with other strategies (22.6% versus 41.8%; odds ratio [OR], 0.439; 95% confidence interval [CI], 0.237-0.816; $P = .009$) [19]. Although reported in-hospital mortality rates for surgical embolectomy remain higher than for systemic thrombolysis or CDTs (20% versus 16% versus 7%, $P < .01$) [20], this may reflect a selection bias with physicians opting toward surgical thrombectomy or systemic thrombolysis over CDT for higher severity PE.

Variant 1: Adult. Extensive acute bilateral central pulmonary emboli. Sustained hypotension for more than 15 minutes. Initial therapy.
E. Systemic thrombolysis

A meta-analysis review of 15 clinical trials concluded that although systemic thrombolytic therapy was associated with a significant reduction of overall mortality compared with anticoagulation alone (OR, 0.59; 95% CI, 0.36-0.96), it was also associated with an increased risk of fatal, intracranial, or major hemorrhage [21]. Another meta-analysis of 33 studies also demonstrated a reduction in overall mortality in patients treated with thrombolysis compared with anticoagulation alone (2.29% versus 4.03%; OR, 0.57; 95% CI, 0.35-0.92) but with increased risk of major bleeding

events (9.46% versus 3.75%; OR, 2.70; 95% CI, 1.83-3.97) [22].

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.

This variant describes a patient with submassive or intermediate-risk PE. The criteria for submassive/intermediate-risk PE as outlined by the AHA is RV strain without hypotension [6]. RV strain can be demonstrated by echocardiography or CT angiography of the pulmonary arteries (measured RV/LV ratio >0.9). Alternatively, RV strain can be demonstrated by an elevation in cardiac biomarkers (troponin, brain natriuretic peptide). The goal of intervention is to improve patient outcome by reducing clot burden and relieving right heart strain.

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.
A. Anticoagulation

Prompt initiation of systemic anticoagulation is considered the standard of care for submassive/intermediate-risk PE [10].

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.
B. Catheter-directed therapy pulmonary artery

Given that intermediate-risk/submassive PE is associated with right heart strain, CDTs facilitate a targeted approach to attempt to reduce the clot burden, thereby relieving the elevated right heart pressure effectively and expediently.

Data supporting the use of pharmacological CDT in high-risk/massive PE detailed previously also extends to the use of these techniques for intermediate-risk/submassive PE. The Ultrasound Accelerated Thrombolysis of Pulmonary Embolism (ULTIMA) trial concluded that CDT was superior to anticoagulation alone in reversing RV dilation at 24 hours in patients with intermediate-risk/submassive PE [23]. In 2015, the PERFECT multicenter registry [11] and the SEATTLE II trial [12] were published supporting the use of catheter-directed delivery of thrombolytic agents directly into the pulmonary arterial system, with the latter demonstrating a decrease in thrombus burden, improvement in right heart strain and reduced pulmonary hypertension [12]. The Optimum Duration of Acoustic Pulse Thrombolysis Procedure in Acute Intermediate-Risk Pulmonary Embolism (OPTALYSE PE) trial demonstrated that CDT can be performed safely with lower-dose thrombolytics and for shorter durations than previously used [24].

Because the market for catheter infusion systems expanded, the Standard Versus Ultrasound-Assisted Thrombolysis for Submassive Pulmonary Embolism (SUNSET sPE) multicenter randomized clinical trial was commenced to evaluate conventional multiside-hole lysis catheters versus US-assisted delivery systems and found them to be comparable in their efficacy in the reduction of thrombus burden at 48 hours posttreatment [25]. The HI-PEITHO multinational multicenter randomized control trial, targeted to complete patient enrollment in December 2023, will evaluate the clinical benefits of US-facilitated catheter-directed thrombolysis with anticoagulation versus anticoagulation alone in high- to intermediate-risk PE, including primary outcome PE-related mortality as well as quality of life indicators and functional status over a 12-month follow-up

period [14].

Catheter-directed mechanical thrombectomy has also demonstrated efficacity in the treatment of PE. The FlowTriever Pulmonary Embolectomy Clinical Study (FLARE) trial evaluated the effectiveness of large bore catheter aspiration thrombectomy in intermediate-risk/submassive PE and demonstrated a statistically significant 25% reduction in RV/LV ratio at 48 hours postprocedure [26]. The EXTRACT-PE (A Prospective Multicenter Trial to Evaluate the Safety and Efficacy of the Indigo Aspiration System in Acute Pulmonary Embolism) trial, a prospective multicenter study of a commercially available small bore catheter–based aspiration system, showed a significant reduction in RV/LV ratio and a low major adverse event rate while avoiding the use of intraprocedural thrombolytic agents in 98.3% of treated patients [27]. In addition, the FLASH registry, a multinational, multicenter registry of patients with high- and intermediate-risk PE treated with the FlowTriever mechanical thrombectomy system, has reported a 7.6 mm Hg mean drop in mean pulmonary artery pressure (-23.0%; $P < .0001$) and decreased RV/LV ratio (1.23-0.98, $P < .0001$) at 48 hours posttreatment, with a favorable adverse event risk profile [13].

In a matched retrospective cohort analysis of 470 patients with submassive PE, patients receiving anticoagulation plus CDT (thrombolysis or aspiration thrombectomy) demonstrated lower mortality at 1 year (7.6% versus 9.8%; $P = .004$) and no difference in bleeding complications at 30 days (2.9% versus 1.6%, $P = .28$) or development of chronic thromboembolic pulmonary hypertension (10.6% versus 9.5%, $P = .44$) [28].

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.

C. Extracorporeal membrane oxygenation

There is no relevant literature to support ECMO as the primary treatment for hemodynamically stable patients with intermediate-risk/submassive PE.

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.

D. Surgical embolectomy pulmonary artery

Although surgical embolectomy has traditionally been considered as a treatment option for hemodynamically unstable patients with massive PE, the literature also lends support to the surgical management of intermediate-risk/submassive central PE in patients with increased bleeding risk from thrombolysis [29]. A retrospective review of 133 patients with submassive PE undergoing either surgical embolectomy or catheter-directed treatment demonstrated reduced incidence of bleeding complications in the surgical patients compared with those receiving endovascular therapy (1.4% versus 9.7%, $P < .05$) [30].

Variant 2: Adult. Acute bilateral pulmonary emboli. RV/LV ratio greater than 0.9 on CTA. Evidence of right heart strain on echocardiogram. Elevated troponin level. No hypotension. Initial therapy.

E. Systemic thrombolysis

The use of systemic thrombolytics in the patient with intermediate-risk/submassive PE is controversial. The multicenter PEITHO randomized control trial evaluated the use of systemic thrombolysis with standard anticoagulation versus standard anticoagulation alone in patients with

intermediate-risk PE [31]. Whereas the systemic thrombolysis group did demonstrate a reduced incidence of death or hemodynamic compromise compared with the anticoagulation group alone (2.6% versus 5.6%; OR, 0.44; 95% CI, 0.23-0.87; *P* = .02), thrombolytic therapy was associated with a high rate of major adverse events including hemorrhagic stroke (2.0% versus 0.2%; *P* = .003) and major extracranial hemorrhage (6.3% versus 1.2%; *P* < .001). Subsequent meta-analysis reviews, although demonstrating an overall reduction in mortality for systemic thrombolysis over anticoagulation alone when both high- and intermediate-risk PE cases were included, found that this benefit disappeared when high-risk PE cases were excluded [21,22]. This would imply that the risk of major bleeding adverse events with systemic thrombolysis would outweigh the mortality benefit for patients with hemodynamically stable intermediate-risk/submassive PE.

Long-term follow-up of patients in the PEITHO trial after a period of at least 24 months (median 37.8 months) from thrombolysis versus placebo for intermediate-risk PE failed to demonstrate a significant difference in mortality rate (20.3% versus 18.0%; *P* = .43), functional limitation (36.0% versus 30.1%; *P* = .23), or chronic thromboembolic pulmonary hypertension detected at echocardiography (2.1% versus 3.2%; *P* = .79) [32].

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

This variant illustrates a patient with low-risk PE, defined by the AHA as not meeting high- or intermediate-risk categories [6].

The goal of intervention is to provide appropriate treatment of acute low-risk PE. Appropriate intervention improves patient outcome by recognizing low-risk clot burden and providing clinical management to prevent escalation to a higher-risk category.

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

A. Anticoagulation

Anticoagulation is typically considered the standard of care for low-risk PE [10]. In patients with low-risk isolated subsegmental PE, a meta-analysis of pooled data fails to demonstrate clear harm or benefit for anticoagulation, suggesting that the decision of whether to commence anticoagulation may be determined by a case-by-case clinical judgment [33].

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

B. Catheter-directed therapy pulmonary artery

There is no relevant literature to support CDT as the primary treatment for patients with low-risk PE.

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

C. Extracorporeal membrane oxygenation

There is no relevant literature to support ECMO as the primary treatment for patients with low-risk

PE.

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

D. Surgical embolectomy pulmonary artery

The literature does not support surgical embolectomy as the primary treatment for patients with low-risk PE. A single-center review including 779 patients with central or peripheral PE presented an algorithm for appropriate surgical management for patients with PE [34]. In this algorithm, patients with peripheral PE were not surgical candidates because of unfavorable anatomic accessibility and were recommended for medical therapy. Patients with central PE and without hemodynamic compromise or evidence of RV dysfunction (low-risk per AHA classification) were also recommended for medical therapy.

Variant 3: Adult. Acute bilateral pulmonary emboli. RV/LV ratio less than 0.9 on CTA. No right heart strain on echocardiogram. Normal troponin level. No hypotension. Initial therapy.

E. Systemic thrombolysis

There is no relevant literature to support systemic thrombolysis as the primary treatment for patients with low-risk PE.

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.

This variant describes a patient with low-risk central PE. The literature evaluating the implications of central versus peripheral PE distribution is controversial. Although a central distribution of PE may be more likely to be associated with RV strain [35], central PE does not appear to be an independent predictor of mortality [36]. In fact, in a single-center retrospective review of 174 patients with central PE that was either saddle or nonsaddle in distribution, the nonsaddle PE had greater clinical RV dysfunction [37]. Thus, the management of saddle PE should be guided by appropriate risk stratification rather than appearance on imaging [38].

The goal of intervention is to provide appropriate treatment of appropriately risk stratified PE, by understanding that clinical parameters have greater weight in determining risk classification than imaging findings alone. Appropriate intervention improves patient outcome by providing appropriate risk stratified care and preventing escalation to a higher-risk category.

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.

A. Anticoagulation

Anticoagulation is considered the standard of care for central PE [10].

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.

B. Catheter-directed therapy pulmonary artery

There is no relevant literature to support the use of CDT as the primary treatment for patients with low-risk central PE without hypotension or clinical or imaging evidence of right heart strain.

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.

C. Extracorporeal membrane oxygenation

There is no relevant literature to support the use of ECMO as the primary treatment for patients with low-risk central PE without hypotension or clinical or imaging evidence of right heart strain.

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.
D. Surgical embolectomy pulmonary artery

The literature does not support the use of surgical embolectomy as the primary treatment for patients with low-risk central PE without hypotension or clinical or imaging evidence of right heart strain. A single-center review including 103 patients with central PE, defined as involving the main, primary, or both branch pulmonary arteries, presented an algorithm for appropriate surgical management for patients with central PE [34]. Although RV dysfunction was demonstrated in the majority of patients with central PE who underwent echocardiography (28 of 47; 60%), there was a subset of patients who had central PE and normal RV function and thus would be classified according to the AHA as "low-risk." In this algorithm, patients with central PE and without hemodynamic compromise or evidence of RV dysfunction were recommended for treatment with medical therapy.

Variant 4: Adult. Acute saddle pulmonary embolism. Normal RV/LV ratio on CTA. Normal troponin level. No hypotension. Initial therapy.
E. Systemic thrombolysis

There is no relevant literature to support the use of systemic thrombolysis as the primary treatment for patients with low-risk central PE without hypotension or clinical or imaging evidence of right heart strain.

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.

Major bleeding events are a significant contributor to morbidity and mortality of patients receiving thrombolytics for the treatment of acute PE. A risk stratification score, the PE-CH score, was designed to stratify the risk of intracranial hemorrhage in patients receiving thrombolytics, and includes 4 independent prognostic factors: 1) preexisting peripheral vascular disease, 2) age >65 years, 3) prior stroke with residual deficit, and 4) prior myocardial infarction [39].

The goal of intervention is to provide appropriate treatment of appropriately risk stratified PE, in the setting of contraindication to anticoagulation or thrombolysis. Appropriate intervention improves patient outcome by providing alternative therapies to reduce clot burden and improve right heart strain in the context of increased risk of adverse bleeding events.

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.
A. Anticoagulation

The use and timing of anticoagulation as well as the selection of an anticoagulant agent following a recent intracranial bleeding event is controversial and often depends on case-by-case assessment. Although case studies have documented the use of heparin and oral anticoagulants for VTE and PE in patients with intracranial hemorrhage [40,41], there currently exist no large randomized clinical trials evaluating the use of anticoagulation for PE in the context of recent intracranial hemorrhage.

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.
B. Catheter-directed therapy pulmonary artery

Optimal pharmacochemical therapy is suggested to improve outcomes while minimizing risk of bleeding [42]. Although the SEATTLE II and ULTIMA trials evaluating US-assisted catheter-directed thrombolysis excluded patients deemed high risk for thrombolytic therapy, subsequent studies have evaluated the safety profile of localized thrombolysis in this patient population. Lee et al [43] performed an institutional retrospective review of 91 patients undergoing treatment for PE, of whom 17 were deemed high risk for thrombolysis because of such variables as surgery, hemorrhage, or cardiac arrest. There was a total of 7 major bleeding events, including 3 within the high-risk cohort. Conversely, mechanical thrombectomy techniques without the use of adjunctive systemic or locally delivered thrombolytics have been demonstrated as effective in reducing RV/LV ratio with a minimized bleeding risk profile [27].

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.
C. Extracorporeal membrane oxygenation

Whether used as a salvage therapy in the setting of failed thrombolysis or thrombectomy, as a bridge to surgical embolectomy, or as a standalone therapy, ECMO is emerging as a therapeutic option in patients with a high risk of bleeding [44]. ECMO does require full anticoagulation to maintain patency of the system, thereby warranting careful patient selection.

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.
D. Surgical embolectomy pulmonary artery

Although surgical embolectomy may be considered as a treatment option for massive or high-risk central PE in a subset of patients with increased bleeding risk, in the particular setting of acute intracranial hemorrhage, the necessary use of intraoperative high-dose systemic anticoagulation has been associated with risks of further bleeding and neurologic deterioration [45].

Variant 5: Adult. Acute bilateral central pulmonary emboli. Evidence of right heart failure on echocardiogram. Sustained a syncopal event with head trauma and acute intracranial hemorrhage. Initial therapy.
E. Systemic thrombolysis

Intracranial hemorrhage is an absolute contraindication to the use of systemic thrombolysis.

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.

Thrombus from the deep veins of the lower extremities, once mobilized, may temporarily lodge in the right-sided cardiac chambers as it transits toward the pulmonary arterial circulation. This is termed "clot-in-transit" and represents an unstable variant of VTE, which is prone to distal embolization. Right-sided cardiac thrombus has been associated with a higher mortality, approaching 20%, as compared with controls [46].

The goal of intervention is to provide appropriate treatment of acute thrombus-in-transit within

the right heart. Appropriate surgical or nonsurgical intervention improves patient outcome by reducing clot burden and minimizing the risk of significant distal clot embolization with resultant right heart strain.

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.
A. Anticoagulation

Standard anticoagulation is a mainstay in the treatment of PE, including PE with right heart thrombus, or "clot-in-transit." However, the role of anticoagulation as a standalone treatment for right heart thrombus is controversial. Right heart thrombus is associated with a higher-risk for clinical deterioration due to its propensity for clot migration leading to hemodynamic compromise, and thus more advanced therapies may be warranted.

A retrospective review of 325 patients with PE and right heart thrombi failed to demonstrate a lower risk of all-cause death or PE-rated mortality for reperfusion treatment versus anticoagulation alone (6.2% versus 14%; *P* = .15; 4.7% versus 7.8%; *P* = .47) [47]. Multiple additional retrospective reviews, in contrast, demonstrate an improved mortality for treatment with either thrombolysis or surgical embolectomy compared with anticoagulation alone [48-51].

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.
B. Catheter-directed therapy right heart

There are limited data in the literature to evaluate the use of CDTs in the management of right heart clot-in-transit. Case reports have described the use of CDTs in the management of right heart thrombus [52].

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.
C. Extracorporeal membrane oxygenation

There are limited data in the literature to evaluate the use of ECMO in the management of right heart clot-in-transit. Given the propensity of right heart thrombus for distal migration leading to cardiogenic shock, ECMO may be considered as a treatment strategy in hemodynamically unstable patients.

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.
D. Surgical embolectomy right heart

Given the high risk of clinical deterioration, surgical thrombectomy has been employed as a treatment option for patients with high- and intermediate-risk PE and right heart thrombus [53]. A particular advantage to surgical embolectomy for the management of clot-in-transit is that the surgeon can address both the intracardiac clot and the frequently associated central PE concurrently.

Variant 6: Adult. Acute thromboembolism in transit. Thrombus in the right atrium. Sustained hypotension for more than 15 minutes. Initial therapy.
E. Systemic thrombolysis

The use of systemic thrombolysis has been described as a treatment option for right heart thrombus-in-transit, with favorable mortality compared with no treatment or anticoagulation alone [48-51].

Summary of Highlights

This is a summary of the key recommendations from the variant tables. Refer to the complete narrative document for more information.

- **Variant 1**: For massive or high-risk PE, there are a variety of first-line treatment options available. Anticoagulation, systemic thrombolysis, CDTs, and surgical embolectomy are all considered usually appropriate techniques to reduce clot burden and relieve right heart strain, with management strategy typically determined based on patient factors and local expertise.
- **Variant 2**: For submassive or intermediate-risk PE, anticoagulation and CDTs are considered usually appropriate management techniques.
- **Variants 3 and 4**: For patients presenting with low-risk PE, without hypotension or clinical or imaging evidence of right heart strain, anticoagulation alone is considered the usually appropriate management technique. This applies to both central and peripheral pulmonary emboli because management should be guided by appropriate risk stratification rather than appearance on imaging.
- **Variant 5**: For patients with acute high- or intermediate-risk PE necessitating treatment but with a contraindication to thrombolysis, catheter-directed techniques using mechanical thrombectomy without pharmacologic thrombolytics is considered usually appropriate, provided the thrombus is in a distribution accessible to catheter techniques.
- **Variant 6**: For acute right atrial thrombus-in-transit, first-line management with anticoagulation, CDTs, or surgical embolectomy are considered usually appropriate techniques to reduce clot burden and minimize risk of distal embolization causing right heart strain.

Supporting Documents

The evidence table, literature search, and appendix for this topic are available at https://acsearch.acr.org/list. The appendix includes the strength of evidence assessment and the final rating round tabulations for each recommendation.

For additional information on the Appropriateness Criteria methodology and other supporting documents, please go to the ACR website at https://www.acr.org/Clinical-Resources/Clinical-Tools-and-Reference/Appropriateness-Criteria.

Gender Equality and Inclusivity Clause

The ACR acknowledges the limitations in applying inclusive language when citing research studies that predates the use of the current understanding of language inclusive of diversity in sex, intersex, gender, and gender-diverse people. The data variables regarding sex and gender used in the cited literature will not be changed. However, this guideline will use the terminology and definitions as proposed by the National Institutes of Health.

Appropriateness Category Names and Definitions

Appropriateness Category Name	Appropriateness Rating	Appropriateness Category Definition
Usually Appropriate	7, 8, or 9	The imaging procedure or treatment is indicated in

		the specified clinical scenarios at a favorable risk-benefit ratio for patients.
May Be Appropriate	4, 5, or 6	The imaging procedure or treatment may be indicated in the specified clinical scenarios as an alternative to imaging procedures or treatments with a more favorable risk-benefit ratio, or the risk-benefit ratio for patients is equivocal.
May Be Appropriate (Disagreement)	5	The individual ratings are too dispersed from the panel median. The different label provides transparency regarding the panel's recommendation. "May be appropriate" is the rating category and a rating of 5 is assigned.
Usually Not Appropriate	1, 2, or 3	The imaging procedure or treatment is unlikely to be indicated in the specified clinical scenarios, or the risk-benefit ratio for patients is likely to be unfavorable.

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54. Measuring Sex, Gender Identity, and Sexual Orientation.

Disclaimer

The ACR Committee on Appropriateness Criteria and its expert panels have developed criteria for determining appropriate imaging examinations for diagnosis and treatment of specified medical condition(s). These criteria are intended to guide radiologists, radiation oncologists and referring physicians in making decisions regarding radiologic imaging and treatment. Generally, the complexity and severity of a patient's clinical condition should dictate the selection of appropriate imaging procedures or treatments. Only those examinations generally used for evaluation of the patient's condition are ranked. Other imaging studies necessary to evaluate other co-existent diseases or other medical consequences of this condition are not considered in this document. The availability of equipment or personnel may influence the selection of appropriate imaging procedures or treatments. Imaging techniques classified as investigational by the FDA have not been considered in developing these criteria; however, study of new equipment and applications should be encouraged. The ultimate decision regarding the appropriateness of any specific radiologic examination or treatment must be made by the referring physician and radiologist in light of all the circumstances presented in an individual examination.

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