



NARRATIVE REVIEW

Reduction of Iodinated Contrast Medium Dose in Computed Tomography Pulmonary Angiography and its Impact on Image Quality: A Narrative Review

Kleanthis Konstantinidis, MSc* 



Department of Radiology and Diagnostic Imaging, General Hospital of Attica KAT, Athens, Greece

*Corresponding author: Konstantinidis Kleanthis, MSc, Department of Radiology and Diagnostic Imaging, General Hospital of Attica KAT, 2 Nikis Street, Kifissia, 14561, Athens, Greece, Tel: +306940828219

Abstract

Objectives: This narrative literature review provides an overview of the different strategies that have been successfully used to reduce the dose of intravenous contrast media (ICM) while maintaining image quality of pulmonary arteries in computed tomography pulmonary angiography (CTPA). These strategies include optimizing the ICM dose, utilizing modern CT scanners capabilities, customizing patient-specific protocols, and using advanced image reconstruction techniques.

Materials and methods: Thirteen relevant studies published up to February 2024 were identified across PubMed and Scopus databases using a comprehensive search strategy that employed the search terms “CTPA,” “contrast,” “reduction,” and “minimization.” An additional manual search on the Research Gate platform identified eight more studies, which were included in the qualitative synthesis. The inclusion criteria focused on studies that compared image quality between CTPA protocols with reduced ICM dose and standard ICM dose CTPA, or CTPA protocols with different ICM doses.

Conclusions: The review revealed several strategies, including the optimization of ICM dose, leveraging the capabilities of modern computed tomography scanners, patient-specific protocol customization and advanced image reconstruction techniques, which have been successfully implemented to reduce ICM dose while maintaining image quality of pulmonary arteries in CTPA. In addition, scanning with low kVp has allowed reduction in both the required ICM dose and the radiation dose to the patient. Conclusively, reducing the dose of administered ICM in CTPA is feasible, with several techniques and protocols demonstrating efficacy in clinical settings.

Keywords

Computed tomography pulmonary angiography, Pulmonary embolism, Iodinated contrast media

Introduction

In recent years, advancements in medical imaging technology have revolutionized the field of diagnostics. Computed Tomography Pulmonary Angiography (CTPA) is a widely accepted diagnostic standard [1] for identifying and evaluating blood clots within the pulmonary arteries (PA) and has a higher sensitivity and specificity than the D-dimer test alone [2]. However, the use of iodinated contrast media (ICM) in contrast-enhanced imaging procedures requires caution to ensure patient safety and diagnostic accuracy. As the global healthcare community increasingly embraces a commitment to patient safety, there is a growing need to explore and implement practices, that promote patient safety within diagnostic imaging [3,4]. By optimizing scan parameters, refining imaging techniques and leveraging the technological advancements of modern CT scanners, radiological departments can significantly reduce the dose of ICM administered without compromising image quality in CTPA studies.

While ICM administration is necessary to obtain contrast-enhanced images of the PA, its use is not without potential risks and complications for the patient. Conditions such as food, drug and contrast-induced

allergies, hyperthyroidism, diabetes, chronic kidney disease and multiple myeloma should be documented and considered before the ICM administration, to avoid allergic reactions [5], contrast-induced nephropathy (CIN) [6], anaphylaxis [7], thyrotoxicosis [8] and cardiac arrhythmia [9], which can be even fatal for the patient. Additionally, ICM can impose an increased burden or even cause post-contrast acute kidney injury in patients with compromised renal function [10]. The literature demonstrates an interest in optimizing the dose of intravenous ICM used in CTPA, aiming to achieve a balance between image quality and patient safety. Moreover, by analyzing the current research landscape, this review aims to highlight the efficacy of low ICM dose imaging techniques in CTPA with respect to image quality and summarize their outcomes.

Pulmonary embolism (PE) occurs when a blood clot in a deep vein in lower extremities or pelvis is detached and travels through the blood circulation to the lungs, blocking the blood supply from the heart to the lungs. It is a common and serious medical condition with potentially fatal consequences for the patient. PE can present with a variety of symptoms, including shortness of breath, chest pain, rapid heart rate and cough [11]. However, the clinical presentation can be nonspecific, making the diagnosis challenging [12]. CTPA has become the standard imaging modality for diagnosing PE due to its high sensitivity and specificity [2]. It offers several advantages, including its non-invasive nature, rapid imaging acquisition, and high diagnostic accuracy. CTPA allows for the prompt identification and localization of pulmonary emboli from radiologists, guiding clinicians in making timely and informed decision-making [13]. Clinical guidelines recommend CTPA as the first-line imaging test for suspected PE, particularly in cases where the clinical probability is moderate or high, making the use of CTPA integral to the diagnostic algorithm for PE [14]. During CTPA, an ICM bolus is injected into the patient's peripheral vein or through a central venous access using a power injector. The injected ICM enhances the opacification of the main and peripheral PA, while the CT scanner acquires transversal images of the chest, which can be submitted in multiplanar and three-dimensional reconstructions for the detailed imaging of the PA [15]. Appropriate injection rate and technique, scan delay and scan timing are crucial parameters, which allow the radiology team to synchronize the scan with the peak of the contrast-enhancement in PA and achieve optimal CTPA studies [16].

Beyond diagnosis, CTPA plays a crucial role in risk stratification and treatment planning. It facilitates determine the extent of PE, guiding decisions on anticoagulant therapy and, in severe cases, interventions such as thrombolysis or embolectomy [14]. CTPA has revolutionized the diagnosis of PE, providing radiologists and clinicians with a powerful and efficient tool for accurate visualization of the PA. Its widespread adoption

is reflective of its diagnostic efficacy and its pivotal role in improving patient outcomes through timely and appropriate management of PE, with the contribution of radiographers to CTPA protocols optimization and manipulation of CT parameters to be considered significant, in order to obtain high quality images [17].

Factors Affecting the Dose of Iodinated Contrast Medium in CTPA

Patient characteristics

The age and clinical condition of the patient, as well as coexisting medical conditions, such as cardiovascular diseases, diabetes, and chronic kidney disease, can affect vascular anatomy, cardiac output, breathing, mobility and cooperation, making CTPA a challenging procedure in some cases. Some patients may have mobility issues, cognitive impairments, or encounter difficulties that need to be addressed, to ensure a successful and comfortable imaging experience [18]. In addition, the radiology team must be aware of the patient's hemodynamics, to ensure sufficient imaging of the PA. Changes in heart rate and blood circulation may vary with age, impacting the administration of ICM and timing considerations of image acquisition and contrast enhancement in CTPA for different age groups [19].

The body mass and weight of the patient are key factors in CTPA, influencing ICM dose, image quality, radiation dose and diagnostic accuracy. Larger individuals may require higher ICM doses and radiation exposure to achieve optimal PA enhancement [20]. Additionally, the injection rate may be adjusted based on the patient's body mass to ensure adequate contrast delivery [21]. Patients with increased body mass may require an increase in tube current and voltage, to optimize image quality and maintain diagnostic accuracy. Obesity can pose an increased burden on the diagnostic accuracy of CTPA, affecting the visualization of the PA [22]. The radiology team may need to employ specific imaging techniques or reconstruction algorithms to overcome these challenges in patients with an increased body mass.

CTPA protocols

Adopting a low ICM dose protocol contributes to mitigate the risk of adverse reactions and CIN [23]. These protocols involve lower dose or concentration of contrast agents compared to routine CTPA protocols while maintaining diagnostic image quality. Tailoring ICM doses based on patient-specific factors, including weight and clinical history, is part of individualized imaging protocols. CTPA protocols with individualized injection parameters of ICM may be feasible, providing sufficient image quality with a substantial radiation dose reduction [24]. In addition, weight-based ICM dosing may ensure that each patient receives adequate ICM for optimal imaging [20].

Technological advancements in computed tomography

Several technological advancements have revolutionized Computed Tomography (CT), offering radiologists and radiographers new capabilities to perform faster CT scans and minimize radiation and ICM exposure to the patient. Multi-detector (MDCT) and dual-source CT (DSCT) scanners have made significant contributions to the reduction of ICM volume in CTPA. Allowing faster scanning, improved spatial and temporal resolution, dual-energy (DE) scanning options, advanced reconstruction algorithms and monochromatic imaging options, these CT scanners have assisted radiologists, radiographers and researchers in reducing both radiation exposure and ICM volume, without any compromise in image quality compared to routine CTPA protocols [25-28]. Studies have recorded that a low tube voltage (kVp) CTPA protocol on a MDCT scanner may allow simultaneous reduction of radiation exposure and ICM volume while maintaining image quality [29]. On the other hand, studies investigating the efficacy of CTPA protocols employing either low iodine concentration agents or low ICM volume on DSCT scanners have concluded that these scanners may produce high-quality CTPA images alongside a significant reduction in iodine load for the patient [30-32]. Several studies have also investigated high-pitch CTPA protocols. High-pitch scanning offers several benefits during scanning, including reduced scan times and decreased radiation dose compared to standard scanning methods. These capabilities are particularly valuable when fast image acquisition and motion artifact reduction are essential, such as in CTPA [33-36]. Novel technological advancements in CT hardware, such as photon-counting detectors are promising, allowing for significant reduction of ICM and radiation dose in the diagnosis of PE, while maintaining good to excellent image quality [34,37].

Advanced reconstruction algorithms play a pivotal role in maintaining image quality while reducing ICM volume. Iterative reconstruction techniques, in particular, enable image noise reduction and enhance image spatial resolution, compensating for lower exposure parameters and contrast densities [28]. Dual-energy (DE) monochromatic image reconstruction can improve the overall signal-to-noise ratio (SNR), allowing high diagnostic quality even with lower ICM volume or iodine concentration [30]. These advances can compensate for low iodine concentration and reduced volume of ICM, ensuring that image quality in CTPA remains uncompromised.

Methods

Search strategy, identification and selection of studies

This review was conducted according to the

“Preferred Reporting Items for Systematic Reviews and Meta-Analyses” (PRISMA) guidelines [38]. PubMed and Scopus databases were searched in February 2024 without any time restriction for comparative studies and clinical trials that measured image quality of reduced ICM dose CTPA protocols, using the terms: “CTPA” combined with the term “contrast” and either with the term “reduction” or “minimization”. Any duplicates from the two searches were merged. On an additional manual search on the Research Gate platform, eight more studies were identified and included in the qualitative synthesis. As part of the screening process; any records that were not relevant to the subject of this review were removed.

Eligibility criteria

Full-text articles were considered for eligibility if they met the following inclusion criteria (Table 1).

Full-text articles not meeting those criteria were excluded. The procedure of study selection is depicted in the flow diagram in Figure 1 and the data of the twenty-one studies that fulfilled the study criteria are summarized in Table 2.

Results

This review includes 21 studies, both prospective and retrospective, that were published between 2010 and 2023. These studies involved 3000 adult patients suspected of having PE who underwent CTPA. Most studies used both qualitative and quantitative image evaluation to compare the image quality of different CTPA protocols with different ICM doses. However, in three studies, only qualitative or quantitative image evaluation was used. Most studies evaluated both subjective and objective image quality. The subjective evaluation relies on the interpretation of images by radiologists or trained observers, focusing on the opacification of PA, image noise levels and artifacts, which directly influence diagnostic confidence in CTPA. The observers used rating scales to rate the vasculature opacification across the included studies. The rating scales varied from 3 to 5-point, where the first number denotes excellent image quality and the last number denotes non-diagnostic image quality. On the other hand, objective evaluation employs quantitative metrics to measure image characteristics such as CNR and SNR, providing a standardized means to compare imaging protocols. While objective measures offer reproducibility and can guide protocol optimization, they may not fully capture the diagnostic utility perceived by human observers. Thus, a comprehensive assessment of CT image quality necessitates a balanced integration of both subjective and objective evaluations, ensuring that imaging techniques not only meet technical standards but also effectively support clinical decision-making. All studies presented in Table 2 demonstrate feasible CTPA protocols with reduced ICM doses.

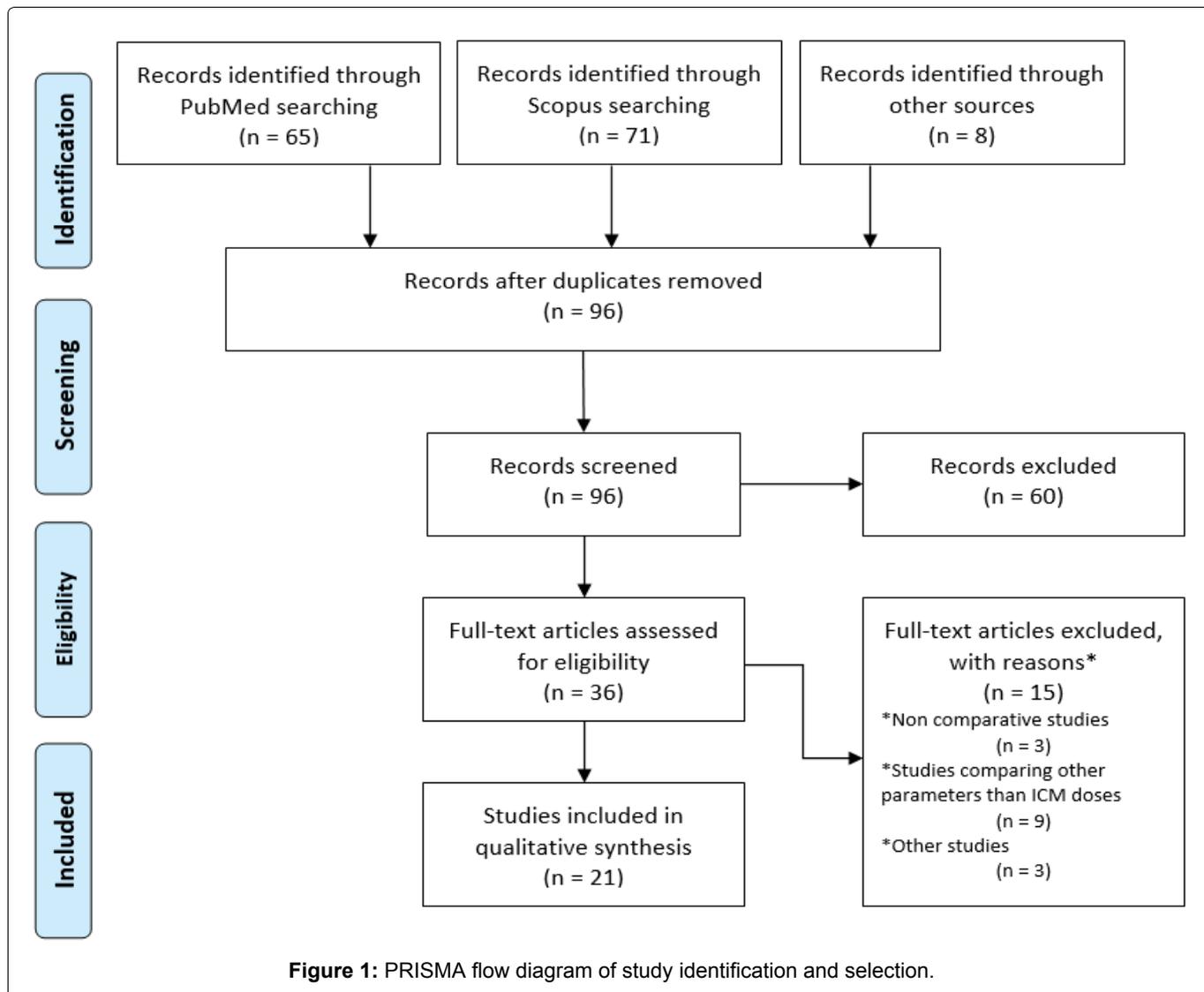


Table 1: Inclusion and exclusion criteria for study selection.

Inclusion criteria	Exclusion criteria	Rationale
All publications to date	n/a	To include all relevant studies
Full-text articles in English language	Articles not written in English language	Difficulty in comprehension
Prospective, retrospective studies and clinical trials	Case reports, reviews, meta-analyses	To include studies with original results only
Patients suspected of PE undergoing CTPA	Patients with other than PE pulmonary conditions	To assess image quality of PA
Studies comparing image quality of a CTPA protocol with reduced ICM dose to a standard CTPA protocol, employing either qualitative or quantitative image evaluation, or both	Non comparative studies, phantom studies, studies not assessing image quality for different ICM doses	This review aims to summarize the findings of comparative studies on CTPA image quality
Studies comparing image quality between CTPA protocols with various reduced ICM doses, employing either qualitative or quantitative image evaluation, or both	Non comparative studies, phantom studies, studies not assessing image quality for different ICM doses	This review aims to summarize the findings of comparative studies on CTPA image quality

In total, fifteen full text articles were excluded from the qualitative synthesis. Three studies assessed the efficacy of a CTPA protocol utilizing a low ICM dose, however, none of them conducted a comparative analysis with a standard protocol [28,35,39]. Nine studies

were investigating other parameters or aspects of CTPA protocols or contrast agents, such as injection rate and iodine concentration [31,40-47]. One study was a phantom study [48]. One study examined the diagnostic accuracy of a CTPA protocol with low radiation and ICM

Table 2: Summary of the characteristics of studies included in the review (published 2010-2023).

Author, year	Study design	Patients (n)	Intervention (test group)	Comparator (control group)	Qualitative image evaluation	Quantitative image evaluation
Schönfeld, et al. (2023) [33]	Retrospective study	n = 81	High-pitch CTPA, 20 ml ICM (n = 41)	Standard CTPA, 50 ml ICM (n = 40)	Good to excellent subjective image quality in over 90% of all exams with no significant difference between the groups.	Significantly lower mean contrast opacification, noise values and CNR in all segmented PA in test group compared to control group but within acceptable diagnostic limits.
Saeed, et al. (2023) [34]	Retrospective study	n = 105	High-pitch PCD-CTPA, 35 ml (n = 29), 45 ml (n = 62), 60 ml (n = 14) ICM	Comparison of image quality between the 3 groups	Ratings: 4.6/5 (group 1), 4.5/5 (group 2), 4.1/5 (group 3). Significant difference between groups 1 and 3 and between groups 2 and 3.	No significant difference in CNR, mean attenuation (HU), mean density (HU) in all evaluated PA locations in all groups.
Pannenbecker, et al. (2023) [37]	Retrospective study	n = 64	PCD-CTPA, 25 ml ICM (n = 32)	DE-CTPA, 50 ml ICM (n = 32)	Superior subjective image quality for 60-keV PCD scans (excellent or good ratings in 93.8% of PCD vs. 84.4% of 60 keV DE scans).	Significantly higher objective image quality parameters in the control group, both in the polychromatic reconstructions and at 60 keV.
Çeltikçi, et al. (2022) [49]	Retrospective study	n = 91	DE-CTPA, 40 ml ICM (n = 42)	Standard CTPA, 60ml ICM (n = 49)	No significant difference between the test and control group in a five-point scale scores for PA CE and image noise.	No significant difference between the test and control group in attenuation values (HU) in five PA locations, mean attenuation value (HU), mean background noise, SNR and CNR.
Wu, et al. (2020) [50]	Prospective observational study	n = 70	Biphasic time-enhancement curves approach, 80 kVp, 10 ml ICM (n = 35)	Test-bolus approach, 100 kVp, 20 ml ICM (n = 35)	Better PA image quality in the test group compared to the control group, with artifact reduction in the superior vena cava and subclavian vein.	Lower CT values, SNR and CNR of the evaluated PA and PV in the test group compared to the control group.
Silva, et al. (2020) [51]	Retrospective study	n = 176	CTPA with 20 ml ICM (n = 102)	CTPA with 40 ml ICM (n = 74)	Significant lower semi-qualitative scores for central and peripheral PACE for the test group. Comparable semi-qualitative image noise between the two groups.	Lower mean CE for the test group, though higher than the diagnostic threshold of 250 HU in both groups. Lower SNR and CNR for the test group compared to the control group.
Kamr, et al. (2020) [52]	Prospective study	n = 600	Test-bolus CTPA technique, 50 ml ICM (n = 300)	Bolus-tracking CTPA technique, 80-100 ml ICM (n = 300)	35% average diagnostic quality score increase from 1.75 in control group to 2.8 in test group.	Main PA average density increase from 260.5 HU in control group to 320 HU in test group. Ascending aorta average density decrease from 250 HU in control group to 130 HU in test group B.

Author, year	Study design	Patients (n)	Intervention (test group)	Comparator (control group)	Qualitative image evaluation	Quantitative image evaluation
Meyer, et al. (2018) [53]	Prospective study	n = 150	Optimized DE-CTPA, 45 ml mixture of ICM and saline [5.4 gr iodine load] (n = 50)	Standard CTPA or standard DE-CTPA, 80 ml ICM [32 gr iodine load] (n = 100)	No significant difference in the median image quality or the median image noise for:- Both the 40 keV and 50 keV VMS data set between both DE-CTPA protocol VMS datasets for the main and peripheral PA. -both DE-CTPA 40 keV and 50 keV protocols compared to the standard CTPA protocol.	Highest CNR of main PA at 50 keV and peripheral PA at 40 keV for both standard and optimized DE-CTPA. Significantly higher CNR values for the standard DE-CTPA.
Suntharalingam, et al. (2017) [54]	Retrospective study	n = 100	80 kVp CTPA, 25 ml ICM on a dual-source CT (n = 50)	100 kVp CTPA, 60 ml ICM on a dual-source CT (n = 50)	No significant difference in subjective image quality scores of PA between the two groups.	Objective image analysis revealed that signal intensities (SI), SNR and CNR of the PA were equal or significantly higher in the control group.
Chen, et al. (2017) [55]	Retrospective observational study	n = 127	60 ml ICM (n = 70)	75 ml ICM (n = 57)	n/a	No significant difference of mean opacification values (HU) in the main, right and left PA between the test and control group for the optimally opacified scans.
Hendriks, et al. (2016) [56]	Prospective study	n = 100	High-pitch CTPA, individual body-weight adjusted ICM dose [42~76 ml] (n = 50)	High-pitch CTPA, 90 ml ICM [75 ml bolus and 15 ml mixed phase] (n = 50)	Diagnostic image quality for all scans in both groups. All scans were graded as "good" or "excellent" at each anatomic level, except one.	No significant difference in attenuation values between the control and test group. No non-diagnostic scans with a minimum mean PA attenuation of 184 HU and 270 HU for the control and the test group respectively. Acceptable CNR for both groups, but better for the test group.
Boos, et al. (2016) [57]	Retrospective study	n = 70	Low-pitch dual-source CTPA at 70 kVp, SimDS, 40 ml ICM (n = 35)	High-pitch dual-source CTPA at 100~120 kVp, ATPS, 70 ml ICM (n = 35)	Diagnostic image quality for all examinations. No significant difference in subjective image quality between the control and test group.	Statistically significant difference between the two groups for the attenuation (HU) in the PT and LLSA. No statistically significant difference between the two groups for the SNR and CNR in the PT and LLSA.
Li, et al. (2015) [58]	Prospective study	n = 80	High-pitch dual-source CTPA at 70 kVp, SAFIRE reconstruction, 40 ml ICM (n = 40)	Low-pitch dual-source CTPA at 100 kVp, FBP reconstruction, 60 ml ICM (n = 40)	No significant difference in subjective image quality between the two groups. No difference in diagnostic accuracy between the two groups.	Higher CT values, SNR and CNR of PA in test group compared to control group.

Author, year	Study design	Patients (n)	Intervention (test group)	Comparator (control group)	Qualitative image evaluation	Quantitative image evaluation
Wang, et al. (2015) [59]	Prospective study	n = 60	Bolus triggering locator in the right atrium, spontaneous respiration, 40 ml ICM (n = 30)	Bolus triggering locator in the pulmonary trunk, suspended respiration, 70 ml ICM (n = 30)	Higher subjective image quality in the test group than the control group.	Significantly higher average CT values of main PA, RULA and RLPA and significantly lower CT values of AA, RUPV and RLPV in the test group than the control group. Significantly higher density between artery and vein pairs in the test group than the control group.
Szucs-Farkas, et al. (2014) [25]	Prospective randomized study	n = 501	80 kVp CTPA, 75 ml ICM (n = 246)	100 kVp CTPA, 100 ml ICM (n = 255)	No difference in subjective image quality and diagnostic confidence in both groups and all BW subgroups.	Decreased attenuation in the PT with increased BW in both groups. No differences in the PT attenuation between the two groups within each BW subgroup. Higher image noise in the test group in all BW subgroups. Higher CNR in the control group compared to the test group in all BW subgroups except for the 90-99 kg subgroup.
Lu, et al. (2014) [36]	Prospective study	n = 100	High-pitch CTPA at 80 kVp, SAFIRE reconstruction, 20 ml ICM (n = 50)	Low-pitch CTPA at 100 kVp, FBP reconstruction, 60 ml ICM (n = 50)	No significant difference in subjective image quality scores between two groups.	Higher mean CT numbers of PA in the test group compared to control group. Higher CNR and SNR of test group than those of control group.
Goble and Abdulkarim (2014) [60]	Retrospective study	n = 139	75 ml of 350 mg iodine/ml [26.25 gr iodine] ICM (n = 70)	100 ml of 300 mg iodine/ml [29.5 gr iodine] ICM (n = 69)	n/a	No significant difference in mean opacification of the main PA between the two groups. Significantly higher opacification in the right and left PA for the test group. No significant difference in the number of suboptimal opacified studies (opacification of less than 250 HU in main PA) between the groups.
Yuan, et al. (2012) [30]	Prospective study	n = 94	DE-CTPA, mixture of ICM and saline in 1:1 fashion resulting in 50% ICM dose and iodine load reduction and image reconstruction at 50 keV (n = 46)	Standard CTPA, undiluted ICM dose and 100 or 120 kVp tube voltage (n = 48)	Higher five-point score for standard CTPA image quality. Higher signal intensity in all PA, inferior noise only in segmental arteries, higher SNR and CNR for DE-CTPA.	n/a

Author, year	Study design	Patients (n)	Intervention (test group)	Comparator (control group)	Qualitative image evaluation	Quantitative image evaluation
Sodickson and Weiss (2012) [29]	Retrospective study	n = 152	100 kVp, 50 ml ICM (n = 53)	120 kVp, 75 ml ICM (n = 99)	n/a	Significant increase in main PA attenuation values by 96 HU and image noise for the test group compared to the control group. Comparable SNR for both groups.
Godoy, et al. (2011) [61]	Retrospective study	n = 50	Reduced contrast [RC] DE-CTPA [80-140 kVp], 50 ml ICM (n = 10)	DE-CTPA [80-140 kVp] CTPA (n = 20) and routine thoracic [RT] CT [80-140 kVp] (n = 20), 100~150 ml ICM	Significantly better central and peripheral vascular enhancement, image noise and global image quality scores in the 80 kVp images than 140 kVp images across all patient groups.	No significant difference in the SNRs in both 80 kVp and 140 kVp between either the CTPA and the RT group or the CTPA and the RC group.
Ramadan, et al. (2010) [62]	Prospective study	n = 90	60 ml (Protocol 1, n = 30), 55 ml (Protocol 2, n = 30), 50 ml (Protocol 3, n = 30) ICM	Comparison of image quality between the 3 protocols	All examinations evaluated as diagnostic in the subjective global image quality evaluation. Image quality was evaluated as diagnostic in 21 (23%) patients and excellent in 69 (77%) subjects. Subjective image quality was better in protocols 2 and 3 than in protocol 1.	Mean attenuation values for PA over 250 HU for all protocols. No difference between the attenuation levels between the three protocols. 90-100% success for protocols 2 and 3 where PA exceeds optimal attenuation (≥ 250 HU).

AA: Ascending Aorta; BW: Body Weight; CE: Contrast Enhancement, Contrast-Enhanced; CNR: Contrast-to-Noise Ratio; DE: Dual-Energy; HU: Hounsfield Units; LLSA: Lower Lobe Segmental Artery; MPA: Main Pulmonary Artery; PA: Pulmonary Arteries; PCD: Photon-Counting Detector; PT: Pulmonary Trunk; PV: Pulmonary Veins; SNR: Signal-to-Noise Ratio; RULA: Right Upper Lobe Artery; RLLPA: Right Lower Lobe Posterior Basal Segmental Artery; RUPV: Right Upper Pulmonary Vein; RLPV: Right Lower Pulmonary Vein; VMS : Virtual Monoenergetic Spectral

dose compared to a standard protocol for different body weights without assessing image quality [26]. One study compared a weight-adjusted contrast administration protocol to a standard CTPA protocol, using various ICM doses instead of using only a low dose of ICM [20].

Discussion

Achieving acceptable diagnostic image quality in CTPA with lower ICM dose involves a wide range of protocols and techniques, as well as utilizing the capabilities of the modern CT scanners and injectors. Several studies have demonstrated the effectiveness of simultaneous high-pitch technique with reduced ICM dose in maintaining image quality. High-pitch CTPA with low dose of ICM and reduced radiation dose may render comparable subjective image quality to standard CTPA with sufficient PA contrast opacification and CNR above diagnostic thresholds in most cases, despite the reduced objective image quality of this technique compared to standard CTPA [33]. At significantly low kVp the use of novel image reconstruction methods such as Sinogram Affirmed Iterative Reconstruction (SAFIRE) may provide comparable image quality and substantial radiation

dose reduction compared to a standard CTPA protocol with filtered-back projection reconstruction [58,63].

As evidenced by researchers, novel Photon-Counting Detector (PCD) technology in CT scanners has facilitated the reduction of ICM dose in CTPA even below 60 ml, providing acceptable image quality [34]. PCD-CTPA offers substantial advantages in terms of rapid image acquisition, radiation and ICM dose reduction and advanced imaging methods. The virtual mono energetic imaging on PCD-CT scanner offers the potential of ICM dose minimization and allows good to excellent image quality compared to a standard DE-CTPA protocol [37].

DE-CTPA presents another approach to reduce ICM dose and iodine load. Researchers, who compared DE-CTPA with standard CTPA, demonstrated that reduced iodine load [53] or ICM dose [49] does not significantly affect the opacification of PA, offering a potential path toward minimizing ICM use. Although aDE-CTPA protocol may result in high SNR and CNR, a slight reduction in image quality and increased image noise may be noted in some cases [30].

Many researchers have denoted the significance

of individualized CTPA protocols, tailored to patient's characteristics such as body weight [20]. A significant reduction in ICM dose was achieved through weight-adjusted contrast administration, as evidenced by Hendriks, et al. who emphasized the importance of tailoring ICM dose to patient weight [56]. They concluded that an individualized CTPA protocol can provide diagnostic image quality with a substantial reduction of ICM volume, especially for lower weight patients, compared to a CTPA protocol with a fixed ICM dose for these patients.

Exploring the feasibility of simultaneous low kVp and ICM dose, researchers have demonstrated the efficacy of such CTPA protocols with significantly reduced contrast dose and radiation exposure, maintaining sufficient image quality to exclude or diagnose PE. Suntharalingam, et al. have concluded that their submillisievert standard-pitch CTPA protocol with 25 ml ICM dose may obtain sufficient image quality while reducing radiation dose by approximately 71% compared to a standard protocol [54]. According to Szucs-Farkas, et al. reduced radiation and ICM dose can provide high vessel attenuation, while maintaining diagnostic image quality and diagnostic confidence [25]. Sodickson and Weiss have also demonstrated the efficacy of their CTPA protocol with low kVp and an appropriately designed ICM injection, to obtain diagnostic images, while reducing radiation exposure and ICM dose by 33% [29].

The concentration of ICM plays a key role in optimizing image quality in reduced ICM dose CTPA protocols. Higher concentrations of ICM allow for reduced volumes to be used while still achieving the necessary PA opacification for diagnostic imaging. This is particularly important in protocols aiming to minimize the dose of ICM to reduce the risk of nephrotoxicity and allergic reactions in vulnerable patients. Goble and Abdulkarim have demonstrated that a reduced volume of high-concentration ICM combined with multiphasic injection technique; allow contrast dose reduction without compromising CTPA accuracy [60]. Furthermore, the combination of low kVp and high-concentration ICM enables further reduction in contrast dose while maintaining or enhancing image quality, underscoring the importance of ICM concentration in achieving optimal diagnostic outcomes with minimal patient risk.

Reducing contrast agents is crucial for ensuring patient safety. Iodinated contrast agents can result in both mild and severe allergic reactions. Mild reactions include symptoms like rash, nausea, and itching, while severe reactions can lead to pulmonary edema, cardiac arrhythmia or arrest. To minimize the likelihood of triggering allergic responses, it is important to reduce the use of injected contrast agents. This is particularly important for patients with a history of allergies to contrast agents, food or drugs [5].

One of the most significant concerns associated with ICM is the development of contrast-induced nephropathy (CIN), particularly in individuals with chronic renal disease [64]. CIN is a serious complication of angiographic procedures following the administration of ICM and is characterized by a sudden impairment of kidneys function. Minimizing the dose or iodine load of the injected contrast agent, especially in patients with renal impairment, can reduce the risk of CIN [65].

Excessive iodine exposure can also affect thyroid function. Minimizing the use of ICM is particularly important for individuals with thyroid disorders like hyperthyroidism, or those at risk of developing thyroid dysfunction. Close monitoring and consideration of alternative imaging approaches may be warranted in such cases, to avoid thyroid dysfunction or thyrotoxicosis [8].

For patients who undergo multiple imaging procedures over time, the cumulative dose of ICM becomes a concern. Minimizing ICM exposure helps mitigate the potential of accumulating high doses of iodine, causing organ dysfunction, triggering allergic reactions and increasing the absorbed organ radiation dose [66], which could lead to contrast-induced adverse effects and risk of cancer.

Extravasation, the unintended leakage of an intravenously injected contrast agent into surrounding tissues, can cause mild skin reaction like inflammation or erythema, and more severe complications such as skin ulceration, tissue necrosis, and compartment syndrome [67]. Managing each patient according to her or his needs, establishing optimal venous access, minimizing the volume, injection rate and concentration of contrast agents, as well as employing proper injection techniques where needed, can reduce the potential of extravasation during CTPA [68,69].

Pregnant women are particularly vulnerable to the potential risks of ionizing radiation exposure. If a pregnant patient must undergo a CTPA, appropriate scan protocol selection and optimization of scan length may be needed, to minimize exposure of the patient and the developing fetus, whilst maintaining diagnostic quality [70]. Despite the use of ICM is considered safe during pregnancy [71], radiology professionals should carefully weigh the benefits of contrast-enhanced imaging against the potential risk of fetal hypothyroidism and submit pregnant patients in CTPA only when the clinical situation requires doing so, keeping the volume of ICM administered as low as possible [72].

Some patients may experience stress, anxiety or discomfort before or during CTPA. Sudden pain at the region of intravenous access, warmth or cold sensation during the administration of ICM may extend the patient's discomfort [73]. Vulnerable populations, such as pediatric patients, the elderly, and those with

multiple comorbidities, may be at a higher risk of contrast-induced adverse effects. Therefore, warming and tailoring ICM volume to the specific needs of these populations may reduce the risk of extravasation or allergic reactions, enhancing overall safety and contributing to a more positive patient experience, especially in cases where the diagnostic benefits of contrast-enhanced phase may be marginal [74]. Providing the patient with detailed information about the CTPA procedure and its potential risks, coupled with maintaining open communication throughout the ICM injection, is essential. This ensures that the patient feels safe and at ease, while simultaneously allowing the healthcare professional to monitor the patient effectively during the procedure [75]. In summary, healthcare professionals must carefully conduct a thorough screening of patients suspected of PE to assess their medical history, including any history of allergies, renal function, and pregnancy status, to determine the necessity of contrast-enhanced imaging, weighting the potential risks against the diagnostic benefits.

Conclusions

The investigation of reducing ICM dose in CTPA reflects a multifaceted approach aimed at optimizing patient care, diagnostic accuracy and patient safety. Numerous studies have delved into the clinical efficacy of CTPA with reduced doses of contrast agents, considering factors such as contrast volume and concentration, injection protocol and advancements in medical technology. The existing body of literature suggests that strategies to reduce ICM administration, while maintaining diagnostic image quality are feasible and potentially beneficial. Studies investigating lower ICM volumes, optimized timing protocols, and the implementation of advanced technologies like dual-energy and photon-counting CT have shown promise in achieving diagnostic accuracy comparable to standard approaches. Moreover, considerations for patient-specific factors, such as body weight, renal function and allergies, play a pivotal role in shaping individualized imaging protocols. The endeavor to reduce ICM is not solely about reducing doses but involves a careful balance between diagnostic accuracy and mitigating potential contrast-induced adverse effects. As medical technology continues to evolve, radiologists, radiographers and healthcare providers must remain attuned to the latest evidence-based guidelines and best practices. In navigating this complex landscape, it is essential for healthcare professionals to exercise clinical judgment, considering the specific clinical context, patient characteristics, and imaging goals. The ultimate goal is to deliver precise and accurate diagnoses while safeguarding patient well-being. As we move forward, collaboration between researchers, radiologists, radiographers and clinicians will contribute to the ongoing refinement of protocols, ensuring that

CTPA with reduced ICM dose continues to be a valuable and safe diagnostic tool in the realm of the PA imaging.

Funding

None.

Conflict of Interest

None.

Acknowledgments

None.

References

- Estrada-Y-Martin RM, Oldham SA (2011) CTPA as the gold standard for the diagnosis of pulmonary embolism. *Int J Comput Assist Radiol Surg* 6: 557-563.
- Nilsson T, Söderberg M, Lundqvist G, Cederlund K, Larsen F, et al. (2002) A comparison of spiral computed tomography and latex agglutination D-dimer assay in acute pulmonary embolism using pulmonary arteriography as gold standard. *Scand Cardiovasc J* 36: 373-377.
- Davenport MS, Perazella MA, Yee J, Dillman JR, Fine D, et al. (2020) Use of intravenous iodinated contrast media in patients with kidney disease: Consensus statements from the American College of radiology and the national kidney foundation. *Kidney Med* 2: 85-93.
- (2019) Patient safety in medical imaging: A joint paper of the European society of radiology (ESR) and the European federation of radiographer societies (EFRS). *Insights Imaging* 10: 45.
- Baerlocher MO, Asch M, Myers A (2010) Allergic-type reactions to radiographic contrast media. *CMAJ* 182: 1328.
- Turedi S, Erdem E, Karaca Y, Tatli O, Sahin A, et al. (2016) The high risk of contrast-induced nephropathy in patients with suspected pulmonary embolism despite three different prophylaxis: A randomized controlled trial. *Acad Emerg Med* 23: 1136-1145.
- Kim MH, Lee SY, Lee SE, Yang MS, Jung JW, et al. (2014) Anaphylaxis to iodinated contrast media: Clinical characteristics related with development of anaphylactic shock. *PLoS One* 9: e100154.
- Dunne P, Kaimal N, MacDonald J, Syed AA (2013) Iodinated contrast-induced thyrotoxicosis. *CMAJ* 185: 144-147.
- Diepenbroek SM, de Jonghe A, van Rees C, Seebus E (2021) Heart failure as a serious complication of iodinated contrast-induced hyperthyroidism: Case-report. *BMC Endocr Disord* 21: 207.
- Elias A, Aronson D (2021) Risk of acute kidney injury after intravenous contrast media administration in patients with suspected pulmonary embolism: A propensity-matched study. *Thromb Haemost* 121: 800-807.
- Tarbox AK, Swaroop M (2013) Pulmonary embolism. *Int J Crit Illn Inj Sci* 3: 69-72.
- MacDonald SLS, Mayo JR (2003) Computed tomography of acute pulmonary embolism. *Seminars in Ultrasound, CT and MRI* 24: 217-231.
- Doğan H, de Roos A, Geleijns J, Huisman MV, Kroft LJM (2015) The role of computed tomography in the diagnosis of acute and chronic pulmonary embolism. *Diagn Interv Radiol* 21: 307-316.

14. Erythropoulou-Kaltsidou A, Alkagiet S, Tziomalos K (2020) New guidelines for the diagnosis and management of pulmonary embolism: Key changes. *World J Cardiol* 12: 161-166.
15. Liu D, Cai X, Che X, Ma Y, Fu Y, et al. (2020) Visibility and image quality of peripheral pulmonary arteries in pulmonary embolism patients using free-breathing combined with a high-threshold bolus-triggering technique in CT pulmonary angiography. *J Int Med Res* 48: 0300060520939326.
16. Zhu J, Wang Z, Kim Y, Bae SK, Tao C, et al. (2017) Analysis of contrast time-enhancement curves to optimise CT pulmonary angiography. *Clin Radiol* 72: 340.e9-340.e16.
17. Mohammad BA, Alakhras MM, Reed W (2023) Assessing the knowledge of CT radiographers regarding how CT parameters affect patient dose and image quality. *Eur J Radiol* 166: 111023.
18. Nguyen ET, Hague C, Manos D, Memauri B, Souza C, et al. (2022) Canadian society of thoracic radiology/ Canadian association of radiologists best practice guidance for investigation of acute pulmonary embolism, part 2: Technical issues and interpretation pitfalls. *Can Assoc Radiol J* 73: 214-227.
19. Ridge CA, Mhuircheartaigh JN, Dodd JD, Skehan SJ (2011) Pulmonary CT angiography protocol adapted to the hemodynamic effects of pregnancy. *AJR Am J Roentgenol* 197: 1058-1063.
20. Ratnakanthan PJ, Kavvoudias H, Paul E, Clements WJ (2020) Weight-adjusted contrast administration in the computed tomography evaluation of pulmonary embolism. *J Med Imaging Radiat Sci* 51: 451-461.
21. Jamali L, Alikhani B, Getzin T, Ringe KI, Wacker FK, et al. (2020) Arterial attenuation in individualized computed tomography pulmonary angiography injection protocol adjusted based on the patient's body mass index. *J Res Med Sci* 25: 94.
22. Hawley PC, Hawley MP (2011) Difficulties in diagnosing pulmonary embolism in the obese patient: A literature review. *Vasc Med* 16: 444-451.
23. Zhang Y, Begum HA, Grewal H, Etxeandia-Ikobaltzeta I, Morgano GP, et al. (2022) Cost-effectiveness of diagnostic strategies for venous thromboembolism: A systematic review. *Blood Adv* 6: 544-567.
24. Tang Z, Fan K, Qiu L, Chen L, Qian Q, et al. (2022) Clinical value and feasibility of CT pulmonary angiography with personalized injection of contrast agent in pulmonary embolism. *Am J Transl Res* 14: 6774-6781.
25. Szucs-Farkas Z, Megyeri B, Christe A, Vock P, Heverhagen JT, et al. (2014) Prospective randomised comparison of diagnostic confidence and image quality with normal-dose and low-dose CT pulmonary angiography at various body weights. *Eur Radiol* 24: 1868-1877.
26. Szucs-Farkas Z, Christe A, Megyeri B, Rohacek M, Vock P, et al. (2014) Diagnostic accuracy of computed tomography pulmonary angiography with reduced radiation and contrast material dose: A prospective randomized clinical trial. *Invest Radiol* 49: 201-208.
27. Lee JW, Lee G, Lee NK, Moon JI, Ju YH, et al. (2016) Effectiveness of adaptive statistical iterative reconstruction for 64-slice dual-energy computed tomography pulmonary angiography in patients with a reduced iodine load: Comparison with standard computed tomography pulmonary angiography. *Journal of Computer Assisted Tomography* 40: 777-783.
28. Laqmani A, Kurfürst M, Butscheidt S, Sehner S, Schmidt-Holtz J, et al. (2016) CT pulmonary angiography at reduced radiation exposure and contrast material volume using iterative model reconstruction and iodose4 technique in comparison to FBP. *PLoS One* 11: e0162429.
29. Sodickson A, Weiss M (2012) Effects of patient size on radiation dose reduction and image quality in low-kVp CT pulmonary angiography performed with reduced IV contrast dose. *Emerg Radiol* 19: 437-445.
30. Yuan R, Shuman WP, Earls JP, Hague CJ, Mumtaz HA, et al. (2012) Reduced iodine load at CT pulmonary angiography with dual-energy monochromatic imaging: Comparison with standard CT pulmonary angiography -- A prospective randomized trial. *Radiology* 262: 290-297.
31. Wichmann JL, Hu X, Kerl JM, Schulz B, Frellesen C, et al. (2015) 70 kVp computed tomography pulmonary angiography: Potential for reduction of iodine load and radiation dose. *J Thorac Imaging* 30: 69-76.
32. Meier A, Higashigaito K, Martini K, Wurnig M, Seifert B, et al. (2016) Dual energy CT pulmonary angiography with 6g iodine-a propensity score-matched study. *PLoS One* 11: e0167214.
33. Schönfeld T, Seitz P, Kriehoff C, Ponorac S, Wötzel A, et al. (2023) High-pitch CT pulmonary angiography (CTPA) with ultra-low contrast medium volume for the detection of pulmonary embolism: A comparison with standard CTPA. *European Radiology* 34: 1921-1931.
34. Saeed S, Niehoff JH, Boriesosdick J, Michael A, Woeltjen MM, et al. (2024) Minimizing contrast media dose in ct pulmonary angiography with clinical photon counting using high pitch technique. *Acad Radiol* 31: 686-692.
35. Alobeidi H, Alshamari M, Widell J, Eriksson T, Lidén M (2020) Minimizing contrast media dose in CT pulmonary angiography with high-pitch technique. *Br J Radiol* 93: 20190995.
36. Lu GM, Luo S, Meinel FG, McQuiston AD, Zhou CS, et al. (2014) High-pitch computed tomography pulmonary angiography with iterative reconstruction at 80 kVp and 20 mL contrast agent volume. *Eur Radiol* 24: 3260-3268.
37. Pannenbecker P, Huflage H, Grunz JP, Gruschwitz P, Patzer TS, et al. (2023) Photon-counting CT for diagnosis of acute pulmonary embolism: Potential for contrast medium and radiation dose reduction. *European Radiology* 33: 7830-7839.
38. Moher D, Liberati A, Tetzlaff J, Altman DG (2009) Preferred reporting items for systematic reviews and meta-analyses: The PRISMA statement. *PLoS Med* 6: e1000097.
39. Han D, Shi W, Chen X, Zhou J, Yu Y, et al. (2020) Optimal monochromatic energy levels in dual-energy spectral CT pulmonary angiography with low contrast medium dosage. *I J Radiol* 17: e83479.
40. Mitchell DP, Rowan M, Loughman E, Ridge CA, MacMahon PJ (2017) Contrast monitoring techniques in CT pulmonary angiography: An important and underappreciated contributor to breast dose. *Eur J Radiol* 86: 184-189.
41. Brendlin AS, Winkelmann MT, Peisen F, Artzner CP, Nikolaou K, et al. (2021) Diagnostic performance of a contrast-enhanced ultra-low-dose high-pitch CT protocol with reduced scan range for detection of pulmonary embolisms. *Diagnostics* 11: 1251.
42. Nance JW, Henzler T, Meyer M, Apfaltrer P, Braunagel M, et al. (2012) Optimization of contrast material delivery for dual-energy computed tomography pulmonary angiography

- in patients with suspected pulmonary embolism. *Invest Radiol* 47: 78-84.
43. Aviram G, Rogowski O, Gotler Y, Bendler A, Steinvil A, et al. (2008) Real-time risk stratification of patients with acute pulmonary embolism by grading the reflux of contrast into the inferior vena cava on computerized tomographic pulmonary angiography. *J Thromb Haemost* 6: 1488-1493.
44. Harun HH, Karim MKA, Muhammad NA, Razak HRA, Sabarudin A, et al. (2020) Effect of iterative reconstruction algorithm associated with low contrast detectability performance from CT pulmonary angiography examinations. *J Phys: Conf Ser* 1505: 1-6.
45. Xue M, Zhang H, Kligerman S, Klahr P, D'Souza W, et al. (2013) Individually optimized uniform contrast enhancement in CT angiography for the diagnosis of pulmonary thromboembolic disease--a simulation study. *Med Phys* 40: 121906.
46. Rodrigues JCL, Joshi D, Lyen SM, Negus IS, Manghat NE, et al. (2014) Tube potential can be lowered to 80 kVp in test bolus phase of CT coronary angiography (CTCA) and CT pulmonary angiography (CTPA) to save dose without compromising diagnostic quality. *Eur Radiol* 24: 2458-2466.
47. Hu X, Ma L, Zhang J, Li Z, Shen Y, et al. (2017) Use of pulmonary CT angiography with low tube voltage and low-iodine-concentration contrast agent to diagnose pulmonary embolism. *Sci Rep* 7: 12741.
48. Harun HH, Karim MKA, Muhammad NA, Kechik MMA, Chew MT, et al. (2021) Task-based assessment on various optimization protocols of computed tomography pulmonary angiography examination. *Radiation Physics and Chemistry* 188: 109692.
49. Çeltikçi P, Hekimoğlu K, Kahraman G, Haberal KM, Kılıç D (2022) Dual-energy computed tomography pulmonary angiography with ultra-low dose contrast administration: Comparison of image quality with standard computed tomography pulmonary angiography. *Turk Gogus Kalp Damar Cerrahi Derg* 30: 549-556.
50. Wu H, Chen X, Zhou H, Qin B, Cao J, et al. (2020) An optimized test bolus for computed tomography pulmonary angiography and its application at 80 kV with 10 ml contrast agent. *Sci Rep* 10: 10208.
51. Silva M, Milanese G, Cobelli R, Manna C, Rasciti E, et al. (2020) CT angiography for pulmonary embolism in the emergency department: Investigation of a protocol by 20 ml of high-concentration contrast medium. *Radiol Med* 125: 137-144.
52. Kamr WH, El-Tantawy AM, Harraz MM, Tawfik AI (2020) Pulmonary embolism: Low dose contrast MSCT pulmonary angiography with modified test bolus technique. *Eur J Radiol Open* 7: 100254.
53. Meyer M, Haubenreisser H, Schabel C, Leidecker C, Schmidt B, et al. (2018) CT pulmonary angiography in patients with acute or chronic renal insufficiency: Evaluation of a low dose contrast material protocol. *Sci Rep* 8: 1995.
54. Suntharalingam S, Mikat C, Stenzel E, Erfanian Y, Wetter A, et al. (2017) Submillisievert standard-pitch CT pulmonary angiography with ultra-low dose contrast media administration: A comparison to standard CT imaging. *PLoS One* 12: e0186694.
55. Chen M, Mattar G, Abdulkarim JA (2017) Computed tomography pulmonary angiography using a 20% reduction in contrast medium dose delivered in a multiphasic injection. *World J Radiol* 9: 143-147.
56. Hendriks BMF, Kok M, Mihil C, Bekkers SCAM, Wildberger JE, et al. (2016) Individually tailored contrast enhancement in CT pulmonary angiography. *Br J Radiol* 89: 20150850.
57. Boos J, Kröpil P, Lanzman RS, Aissa J, Schleich C, et al. (2016) CT pulmonary angiography: simultaneous low-pitch dual-source acquisition mode with 70kVp and 40 ml of contrast medium and comparison with high-pitch spiral dual-source acquisition with automated tube potential selection. *Br J Radiol* 89: 20151059.
58. Li X, Ni QQ, Schoepf UJ, Wichmann JL, Felmlly LM, et al. (2015) 70-kVp High-pitch computed tomography pulmonary angiography with 40 mL contrast agent. *Acad Radiol* 22: 1562-1570.
59. Wang M, Li W, Lun-Hou D, Li J, Zhai R (2015) Optimizing computed tomography pulmonary angiography using right atrium bolus monitoring combined with spontaneous respiration. *Eur Radiol* 25: 2541-2546.
60. Goble EW, Abdulkarim JA (2014) CT pulmonary angiography using a reduced volume of high-concentration iodinated contrast medium and multiphasic injection to achieve dose reduction. *Clin Radiol* 69: 36-40.
61. Godoy MCB, Heller SL, Naidich DP, Assadourian B, Leidecker C, et al. (2011) Dual-energy MDCT: Comparison of pulmonary artery enhancement on dedicated CT pulmonary angiography, routine and low contrast volume studies. *Eur J Radiol* 79: e11-e17.
62. Ramadan SU, Kosar P, Sonmez I, Karahan S, Kosar U (2010) Optimisation of contrast medium volume and injection-related factors in CT pulmonary angiography: 64-slice CT study. *Eur Radiol* 20: 2100-2107.
63. Sieren JP, Hoffman EA, Fuld MK, Chan KS, Guo J, et al. (2014) Sinogram affirmed iterative reconstruction (SAFIRE) versus weighted filtered back projection (WFBP) effects on quantitative measure in the COPD Gene 2 test object. *Med Phys* 41: 091910.
64. Isaka Y, Hayashi H, Aonuma K, Horio M, Terada Y, et al. (2020) Guideline on the use of iodinated contrast media in patients with kidney disease 2018. *Clin Exp Nephrol* 24: 1-44.
65. Mohammed NMA, Mahfouz A, Achkar K, Rafie IM, Hajar R (2013) Contrast-induced nephropathy. *Heart Views* 14: 106-116.
66. Mazloumi M, Van Gompel G, Kersemans V, de Mey J, Buls N (2021) The presence of contrast agent increases organ radiation dose in contrast-enhanced CT. *Eur Radiol* 31: 7540-7549.
67. Roditi G, Khan N, van der Molen AJ, Bellin MF, Bertolotto M, et al. (2022) Intravenous contrast medium extravasation: Systematic review and updated ESUR contrast media safety committee guidelines. *Eur Radiol* 32: 3056-3066.
68. Shigematsu S, Oda S, Sakabe D, Matsuoka A, Hayashi H, et al. (2022) Practical preventive strategies for extravasation of contrast media during CT: What the radiology team should do. *Acad Radiol* 29: 1555-1559.
69. Tonolini M, Campari A, Bianco R (2012) Extravasation of radiographic contrast media: Prevention, diagnosis, and treatment. *Curr Probl Diagn Radiol* 41: 52-55.
70. Hendriks BMF, Schnerr RS, Milanese G, Jeukens CRLPN, Niesen S, et al. (2019) Computed tomography pulmonary angiography during pregnancy: Radiation dose of commonly used protocols and the effect of scan length optimization. *Korean J Radiol* 20: 313-322.

71. Perelli F, Turrini I, Giorgi MG, Renda I, Vidiri A, et al. (2022) Contrast agents during pregnancy: Pros and cons when really needed. *Int J Environ Res Public Health* 19: 16699.
72. Van Welie N, Portela M, Dreyer K, Schoonmade LJ, van Wely M, et al. (2021) Iodine contrast prior to or during pregnancy and neonatal thyroid function: A systematic review. *Eur J Endocrinol* 184: 189-198.
73. McCullough PA, Capasso P (2011) Patient discomfort associated with the use of intra-arterial iodinated contrast media: A meta-analysis of comparative randomized controlled trials. *BMC Med Imaging* 11: 12.
74. Zhang B, Liu J, Dong Y, Guo B, Lian Z, et al. (2018) Extrinsic warming of low-osmolality iodinated contrast media to 37°C reduced the rate of allergic-like reaction. *Allergy Asthma Proc* 39: e55-e63.
75. Lange S, Mędrzycka-Dąbrowska W, Małecka-Dubiela A (2023) Patient experience during contrast-enhanced computed tomography examination: Anxiety, feelings, and safety. *Safety* 9: 69.