

## Original Article



# Low Contrast and Low kV CTA Before Transcatheter Aortic Valve Replacement: A Systematic Review

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## ABSTRACT

**BACKGROUND:** Minimizing contrast dose and radiation exposure while maintaining image quality during computed tomography angiography (CTA) for transcatheter aortic valve replacement (TAVR) is desirable, but not well established. This systematic review compares image quality for low contrast and low kV CTA versus conventional CTA in patients with aortic stenosis undergoing TAVR planning.

**METHODS:** We performed a systematic literature review to identify clinical studies comparing imaging strategies for patients with aortic stenosis undergoing TAVR planning. The primary outcomes of image quality as assessed by the signal-to-noise ratio (SNR) and contrast-to-noise ratio (CNR) were reported as random effects mean difference with 95% confidence interval (CI).

**RESULTS:** We included 6 studies reporting on 353 patients. There was no difference in cardiac SNR (mean difference, -1.42; 95% CI, -5.71 to 2.88;  $p = 0.52$ ), cardiac CNR (mean difference, -3.83; 95% CI, -9.98 to 2.32;  $p = 0.22$ ), aortic SNR (mean difference, -0.23; 95% CI, -7.83 to 7.37;  $p = 0.95$ ), aortic CNR (mean difference, -3.95; 95% CI, -12.03 to 4.13;  $p = 0.34$ ), and ileofemoral SNR (mean difference, -6.09; 95% CI, -13.80 to 1.62;  $p = 0.12$ ) between the low dose and conventional protocols. There was a difference in ileofemoral CNR between the low dose and conventional protocols with a mean difference of -9.26 (95% CI, -15.06 to -3.46;  $p = 0.002$ ). Overall, subjective image quality was similar between the 2 protocols.

**CONCLUSIONS:** This systematic review suggests that low contrast and low kV CTA for TAVR planning provides similar image quality to conventional CTA.

**Keywords:** Aortic valve; Contrast media; Heart valve prosthesis

## INTRODUCTION

Transcatheter aortic valve replacement (TAVR) is an established treatment method for patients with symptomatic severe aortic stenosis regardless of their surgical risk.<sup>1</sup> Computed tomography angiography (CTA) is often used for TAVR planning to comprehensively assess valve anatomy and vascular access to facilitate patient selection and prosthesis sizing.<sup>2,3</sup> Patients undergoing TAVR are often elderly with multiple comorbidities including renal dysfunction.<sup>4</sup> The use of iodinated contrast media in patients with renal dysfunction

undergoing TAVR can cause acute kidney injury that is associated with increased morbidity and mortality.<sup>5</sup> Therefore, reducing the amount of contrast media administered during CTA for TAVR planning is an active area of research.<sup>6</sup> The use of radiation protection strategies by using low tube voltage settings has shown promise to reduce contrast media volumes for CTA.<sup>7</sup> However, there is limited data on image quality using low contrast and low kV CTA for TAVR planning. The objective of this systematic review was to assess image quality for low contrast and low kV CTA versus conventional CTA in patients for TAVR planning.

## METHODS

This study did not require ethical approval because only public published data was used. The reporting of this systematic review follows the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement.<sup>8</sup> This review was submitted to PROSPERO for registration.

### Search strategy

We performed a systematic literature review to identify randomized and nonrandomized clinical studies that reported qualitative and quantitative image outcomes using low contrast and low radiation CT protocols in patients before TAVR. Searches were limited to peer-reviewed articles published in English up to March 1<sup>st</sup>, 2022. This research involved human subjects and described image quality on patients undergoing CT for TAVR assessment. We developed the search strategy according to available guidance from the Cochrane Collaboration.

The search strategy in PubMed explored Medical Subject Heading (MeSH) terms related to patients undergoing low contrast and low kV CTA for TAVR assessment. The exact search strategy is available as a supplementary file. The articles found to be relevant during the search were stored in EndNote. References from full-text articles were also evaluated and considered for inclusion.

### Study selection

Articles were selected for inclusion based on predefined criteria, which included low contrast and radiation dose CT protocols for TAVR assessment. The primary outcome was quantitative image quality as assessed by the signal-to-noise ratio (SNR) or contrast-to-noise ratio (CNR). We excluded feasibility studies, case series, studies that did not report on our primary outcome, and studies that did not have different contrast volumes between groups.

Two authors (SL, MB) independently completed literature searches and screened abstracts to choose potentially relevant articles. Selected articles underwent full evaluation to assess their potential inclusion in the systematic review.

### Risk for bias

The risk for bias was assessed using the Cochrane tool for assessing the risk for bias in randomized controlled trials (RCTs).<sup>9</sup> The risk of bias for non-randomized trials was evaluated using the ROBINS-I tool.<sup>10</sup> The risk for bias was assessed by 2 independent reviewers (SL, MB).

### Statistical analysis

Review Manager software 5.4 was used for data analysis. Pooled estimates were calculated using Mantel-Hansel methods. The DerSimonian and Laird<sup>11</sup> approach was used for random effects model estimation. We preferred the random effects model to account for potential statistical heterogeneity. Pooled estimates were presented as mean difference with 95% confidence intervals (CI). A  $\chi^2$  test of heterogeneity was calculated for each pooled analysis. The  $I^2$  measure of statistical heterogeneity was also estimated, with higher  $I^2$  values representing greater heterogeneity. All statistical tests were two-sided and p-values < 0.05 were considered significant.

## RESULTS

### Literature search

Our search yielded 126 abstracts. We excluded 107 studies at the abstract level and selected 19 full-text articles for detailed assessment; 6 studies were ultimately included in our systematic review. **Figure 1** describes the flow-chart of included studies.

### Baseline characteristics of the studies

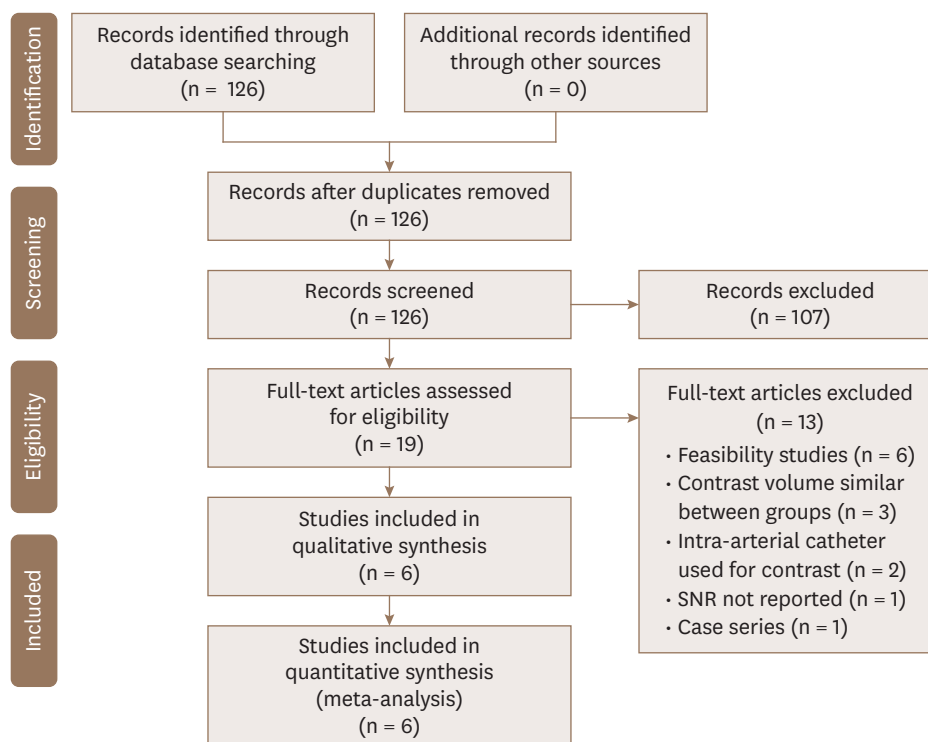
**Table 1** shows the baseline characteristics of the included studies. All studies were published between 2017 and 2020. The 6 studies included 353 patients and the median age of the participants was 82.5 years interquartile range (IQR) (80.4–83.8). The median percentage of men was 51.0 IQR (41.4–54.5). The median BMI for participants was 26.0 IQR (24.8–27.3). In the study by Felmy et al., 25% of the patients were in atrial fibrillation. While in the studies by Franzesi et al. and Ippolito et al., 100% of the patients were in sinus rhythm. In the study by Suchá et al., the low-dose group had significantly worse kidney function with median creatinine of 1.6 compare to median creatinine of 1.0 in the conventional group.

**Table 1.** Baseline characteristics of the included studies

Study, author, year	Location	Sample size	Group	Group size	Mean age, years	Men, %	BMI (kg/m <sup>2</sup> )	Atrial fibrillation (%)	Sinus rhythm (%)	HR (bpm)	LV EF (%)	Creatinine (mg/dL)	GFR (mL/min/1.73m <sup>2</sup> )
Felmly et al., <sup>13</sup> 2017	SC, USA	40	Group B (low-dose)	20	78.6 ± 7.6	75	28.1 ± 5.4	25	NR	62.6 ± 12.4	62.6 ± 17.1	NR	NR
			Group A (standard)	20	81.8 ± 6.2	70	24.8 ± 3.8	25	NR	68.7 ± 11.2	58.3 ± 15.9	NR	NR
Franzesi et al., <sup>14</sup> 2018	Italy	79	Study group	42	83.6 (79–89)	45	24.8 ± 2.9	NR	100	73 ± 6.8	NR	NR	NR
			Control group	37	84.3 (81–90)	54	25.4 ± 3.8	NR	100	75 ± 8.7	NR	NR	NR
Hachulla et al., <sup>15</sup> 2019	Switzerland	84	Protocol 2: 60 mL contrast	42	83 ± 6.8	52	27.2 ± 4.6	NR	NR	75.5 ± 11.0	NR	NR	NR
			Protocol 1: 120 mL contrast	42	86.3 ± 6.5	43	26.5 ± 5.0	NR	NR	76.3 ± 12.2	NR	NR	NR
Onoda et al., <sup>16</sup> 2019	Japan	40	70 kV group	30	84.2 ± 5.8	37	20.3 ± 3.6	NR	NR	NR	62.4 ± 12.2	0.91 ± 0.44	61.4 ± 25.5
			120/120 kV group	10	83.5 ± 6.0	30	23.4 ± 4.8	NR	NR	NR	66.1 ± 5.4	0.67 ± 0.19	75.0 ± 28.4
Ippolito et al., <sup>17</sup> 2020	Italy	60	Study group	32	73.32 (43–78)	56	NR	NR	100	76 ± 6.05	NR	NR	NR
			Control group	28	76.14 (56–88)	50	NR	NR	100	75 ± 8.7	NR	NR	NR
Suchá et al., <sup>18</sup> 2020	CA, USA	50	Low-CM	25	82 ± 7.9	52	27.3 ± 6.4	NR	NR	NR	NR	1.6 (1.5–1.9)	36 (32–44)
			Standard-CM	25	81 ± 7.2	36	28.0 ± 5.1	NR	NR	NR	NR	NR	1.0 (0.9–1.1)

Values are mean ± SD, median (interquartile range), or number (%).

BMI: body mass index, bpm: beats per minute, CA: California, GFR: glomerular filtration rate, HR: heart rate, LV EF: left ventricle ejection fraction, NR: not reported, SC: South Carolina, USA: United States of America.



**Figure 1.** Flow chart of the included studies. SNR: signal-to-noise ratio.

**Risk for bias**

The risk for bias revealed adequate randomization and allocation concealment in the 1 RCT included in this study. Blinding for qualitative image assessment was not addressed in the RCT due to the nature of the study. The non-RCTs included

in this study had appropriate selection and ascertainment approaches, while blinded assessments were deficient. Overall, the risk for bias in the measurement of the quantitative image outcomes was low.

**Low contrast dose strategies for CTA for TAVR planning**

The included studies reported image quality in patients undergoing low contrast CTA for TAVR planning as summarized in **Table 2**.<sup>12,17</sup> Felmly et al.<sup>12</sup> low-dose protocol consisted of a third-generation dual-source CT at 70 kV with 40 ml contrast compared to a conventional protocol with a second-generation dual-source CT at 100 kV and 60 mL contrast. Aortoiliac SNR was higher for the low-dose protocol (low dose:  $15.4 \pm 6.7$ , conventional:  $13.1 \pm 6.8$ ,  $p = 0.0003$ ), whereas cardiac SNR (conventional:  $15.6 \pm 9.0$ , low dose:  $12.2 \pm 4.5$ ,  $p = 0.0003$ ) and cardiac CNR (conventional:  $20.2 \pm 13.4$ , low dose:  $15.3 \pm 6.7$ ,  $p = 0.0181$ ) were higher for the conventional protocol. Subjective image quality was similar between the 2 groups for cardiac and aortoiliac attenuation and noise. However, there was significantly increased subjective image quality for aortoiliac image noise for the low dose protocol (low dose: 4.42 [IQR 4.0–5.0], conventional 4.12 [IQR 4.0–5.0],  $p = 0.037$ ). All TAVR candidates were safely and effectively evaluated using the low-dose protocol.<sup>12</sup> Franzesi et al.<sup>13</sup> low-dose protocol consisted of 100 kV, 50 mL contrast, whole-body retrospective ECG-gating, and iterative reconstruction algorithm compared to a conventional protocol using 120 kV, 100 mL contrast, ECG-gating for chest, and Filtered back Projection reconstruction. All patients were examined with a 256-multidetector CT. Higher mean attenuation values were achieved with the low-dose protocol compared to the conventional protocol. There was no significant difference in subjective image quality between the 2 groups.<sup>13</sup> Hachulla et al.<sup>14</sup> low-dose protocol consisted of 60 mL contrast compared to a conventional protocol using 120 mL contrast. All patients were examined with a 128-slice multidetector CT using automated 80-140 kV

tube voltage. Although higher aortic mean attenuation was achieved with the conventional protocol, there was similar overall image quality between the 2 groups.<sup>14</sup> Onoda et al.<sup>15</sup> low-dose protocol consisted of a mean 31 mL contrast with 70 kV tube voltage compared to a conventional protocol with a mean 78 mL contrast with 120 or 100 kV tube voltage using a 192-slice dual-source CT. The low-dose protocol maintained adequate objective image quality.<sup>15</sup> Ippolito et al.<sup>16</sup> low-dose protocol consisted of 80 kV, 60 mL contrast, and iterative reconstruction compared to a conventional protocol using 100 kV, 80 mL contrast, and iDose4 reconstruction. All patients were examined with a 256-row multidetector CT. Higher mean attenuation values were achieved with the low-dose protocol and there were no significant differences in subjective image quality between the 2 groups.<sup>16</sup> Suchá et al.<sup>17</sup> low-dose protocol consisted of a median 69 mL contrast and 90 kV tube voltage compared to a conventional protocol with a median 116 mL contrast and 100 kV tube voltage. All patients were examined with dual-source CT. The low-dose protocol achieved good image quality. However, the conventional protocol was shown to have higher measurement reproducibility. Therefore, the authors recommended reserving low-dose protocols for patients at high risk for postcontrast acute kidney injury.<sup>17</sup>

The average contrast volume (low dose  $51.6 \pm 14.2$  mL, conventional  $92.3 \pm 23.6$ ,  $p = 0.001$ ) and tube voltage (low dose  $82 \pm 13.0$  kV, conventional  $108 \pm 11.0$  kV,  $p = 0.018$ ) was decreased for the low dose protocols when compared to the conventional protocols as shown in **Table 3**. Pooled estimates of the included studies revealed no difference in cardiac SNR between the low dose and conventional protocols with

**Table 2.** Summary of study design, protocols, and outcomes for the included studies

Study, author, year	Sample size	Study design	Low-dose protocol	Conventional protocol	Image acquisition	Outcomes
Felmly et al., <sup>13</sup> 2017	40	Prospective cohort	3rd-gen. DSCT, 70 kV, 40 mL contrast, 2.5 mL/s flow rate, iodine: 10.8 g	2nd-gen. DSCT, 100 kV, 60 mL contrast, 4.0 mL/s flow rate, iodine: 16.1 g	See protocols	Vascular attenuation, noise, SNR, CNR, image quality
Franzesi et al., <sup>14</sup> 2018	79	Prospective cohort	100 kV, whole-body retrospective ECG-gating, 50 mL contrast, iterative reconstruction algorithm, iodine: 350 mg/mL	120 kV, ECG-gating for chest, 100 mL contrast, FBP reconstruction, iodine: 350 mg/mL	256-MDCT	Image quality, vascular enhancement, radiation dose, image noise
Hachulla et al., <sup>15</sup> 2019	84	Retrospective cohort	60 mL contrast, iodine: 350 mg/mL	120 mL contrast, iodine: 350 mg/mL	128-slice MDCT automated 80–140 kV	Mean attenuation, image quality, SNR, CNR, CAD prediction
Onoda et al., <sup>16</sup> 2019	40	Prospective cohort	70 kV, mean 31 mL contrast, iodine: 300 mg/mL	120/100 kV, mean 78 mL contrast, iodine: 300 mg/mL	192-slice DSCT	Vascular attenuation, image noise, CNR, renal function
Ippolito et al., <sup>17</sup> 2020	60	Randomized controlled trial	80-kV ECG-gated, 60 mL contrast, IMR reconstruction, iodine: 350 mg/mL	100 kV, 80 mL contrast, iDose4 reconstruction, iodine: 350 mg/mL	256-row MDCT	Subjective and objective image quality, radiation dose
Suchá et al., <sup>18</sup> 2020	50	Retrospective cohort	Median 69 mL contrast, 90 kV, iodine: 300 mg/mL	Median 116 mL contrast, 100 kV, iodine: 300 mg/mL	DSCT (Slice thickness/increment, mm: 0.75/0.7)	Image quality, pre-TAVR measurement interobserver variability, renal function change

CNR: contrast-to-noise ratio, DSCT: dual-source CT, ECG: electrocardiogram, MDCT: multidetector CT, SNR: signal-to-noise ratio.

Low Dose CTA Before TAVR

**Table 3.** Average contrast volume and tube voltage for the included protocols

Measurement	Low-dose protocols	Conventional protocols	p-value
Average contrast volume (mL)	51.6 ± 14.1	92.3 ± 23.6	0.001
Average tube voltage (kV)	82 ± 13.0	108 ± 10.9	0.018

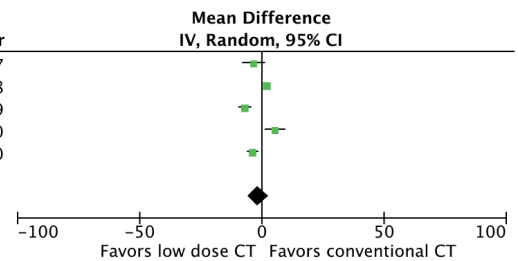
a mean difference of -1.42 (95% CI, -5.71 to 2.88, p = 0.52). The forest plot for cardiac SNR is shown in **Figure 2**. There was no difference in cardiac CNR between the low dose and conventional protocols with a mean difference of -3.83 (95% CI, -9.98 to 2.32, p = 0.22). The forest plot for cardiac CNR is shown in **Figure 3**. The statistical heterogeneity was high for the reported cardiac SNR and CNR outcomes with I<sup>2</sup> values of 95% and 93%, respectively. There was no difference in aortic SNR between the low dose and conventional protocols with a mean difference of -0.23 (95% CI, -7.83 to 7.37, p = 0.95). The forest plot for aortic SNR is shown in **Supplementary Figure 1**. There was no difference in aortic CNR between the low dose and

conventional protocols with a mean difference of -3.95 (95% CI, -12.03 to 4.13, p = 0.34). The forest plot for aortic CNR is shown in **Supplementary Figure 2**. The statistical heterogeneity was high for the reported aortic SNR and CNR outcomes with I<sup>2</sup> values of 95% and 93%, respectively. There was no difference in ileofemoral SNR between the low dose and conventional protocols with a mean difference of -6.09 (95% CI, -13.80 to 1.62, p = 0.12). The forest plot for ileofemoral SNR is shown in **Supplementary Figure 3**. There was a significant difference in ileofemoral CNR between the low dose and conventional protocols with a mean difference of -9.26 (95% CI, -15.06 to -3.46, p = 0.002). The forest plot for ileofemoral CNR is shown in **Figure 4**. The statistical heterogeneity was high for the reported ileofemoral SNR and CNR outcomes with I<sup>2</sup> values of 85% and 80%, respectively.

Qualitative image assessments were completed by the included studies as summarized in **Table 4**. Overall, qualitative image

Study or Subgroup	Low Dose CT			Conventional CT			Weight	Mean Difference IV, Random, 95% CI	Year
	Mean	SD	Total	Mean	SD	Total			
Felmly 2017	12.2	4.5	20	15.6	9	20	17.9%	-3.40 [-7.81, 1.01]	2017
Franzesi 2018	15.7	2.11	42	13.7	1.42	37	21.9%	2.00 [1.21, 2.79]	2018
Hachulla 2019	17	5	42	24	6	42	20.7%	-7.00 [-9.36, -4.64]	2019
Ippolito 2020	22.63	7.78	32	17.25	7.84	28	18.6%	5.38 [1.42, 9.34]	2020
Suchá 2020	12.8	3.2	25	16.6	4.1	25	21.0%	-3.80 [-5.84, -1.76]	2020
<b>Total (95% CI)</b>	<b>161</b>			<b>152</b>			<b>100.0%</b>	<b>-1.42 [-5.71, 2.88]</b>	

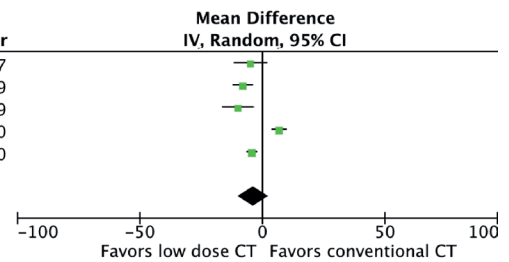
Heterogeneity: Tau<sup>2</sup> = 21.77; Chi<sup>2</sup> = 78.62, df = 4 (P < 0.00001); I<sup>2</sup> = 95%  
Test for overall effect: Z = 0.65 (P = 0.52)



**Figure 2.** Forest plot of cardiac signal-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning. CI: confidence interval, CT: computed tomography, SD: standard deviation.

Study or Subgroup	Low Dose CT			Conventional CT			Weight	Mean Difference IV, Random, 95% CI	Year
	Mean	SD	Total	Mean	SD	Total			
Felmly 2017	15.3	6.7	20	20.2	13.4	20	17.9%	-4.90 [-11.47, 1.67]	2017
Hachulla 2019	30	7	42	38	11	42	20.5%	-8.00 [-11.94, -4.06]	2019
Onoda 2019	21.6	6.7	30	31.6	9.2	10	18.3%	-10.00 [-16.19, -3.81]	2019
Ippolito 2020	19.25	6.92	32	12.43	4.14	28	21.4%	6.82 [3.97, 9.67]	2020
Suchá 2020	10.6	3.3	25	14.9	3.7	25	21.9%	-4.30 [-6.24, -2.36]	2020
<b>Total (95% CI)</b>	<b>149</b>			<b>125</b>			<b>100.0%</b>	<b>-3.83 [-9.98, 2.32]</b>	

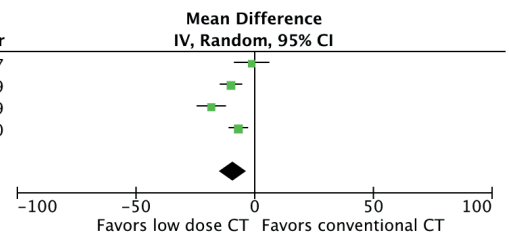
Heterogeneity: Tau<sup>2</sup> = 43.84; Chi<sup>2</sup> = 58.09, df = 4 (P < 0.00001); I<sup>2</sup> = 93%  
Test for overall effect: Z = 1.22 (P = 0.22)



**Figure 3.** Forest plot of cardiac contrast-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning. CI: confidence interval, CT: computed tomography, SD: standard deviation.

Study or Subgroup	Low Dose CT			Conventional CT			Weight	Mean Difference IV, Random, 95% CI	Year
	Mean	SD	Total	Mean	SD	Total			
Felmly 2017	23.9	11.5	20	25.2	11.7	20	21.4%	-1.30 [-8.49, 5.89]	2017
Hachulla 2019	29	7	42	39	13	42	26.8%	-10.00 [-14.47, -5.53]	2019
Onoda 2019	17.4	7.5	30	35.7	8.5	10	23.9%	-18.30 [-24.21, -12.39]	2019
Suchá 2020	15.9	6.1	25	22.8	7.7	25	27.9%	-6.90 [-10.75, -3.05]	2020
<b>Total (95% CI)</b>	<b>117</b>			<b>97</b>			<b>100.0%</b>	<b>-9.26 [-15.06, -3.46]</b>	

Heterogeneity: Tau<sup>2</sup> = 27.52; Chi<sup>2</sup> = 15.23, df = 3 (P = 0.002); I<sup>2</sup> = 80%  
Test for overall effect: Z = 3.13 (P = 0.002)



**Figure 4.** Forest plot of ileofemoral contrast-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning. CI: confidence interval, CT: computed tomography, SD: standard deviation.



**Table 4.** Summary of qualitative image assessment for the included studies

Study	Parameter	Low-dose protocol	Conventional protocol	p-value
Felmly et al., <sup>13)</sup> 2017	Cardiac attenuation	4.71 (IQR 5.0–5.0)	4.67 (IQR 4.0–5.0)	0.720
	Image noise	4.08 (IQR 4.0–4.0)	3.88 (IQR 4.0–4.0)	0.238
	Aortoiliac attenuation	4.53 (IQR 4.0–5.0)	4.33 (IQR 4.0–5.0)	0.213
	Aortoiliac noise	4.42 (IQR 4.0–5.0)	4.12 (IQR 4.0–5.0)	0.037
Franzesi et al., <sup>14)</sup> 2018	Image quality, reader 1	3.48 ± 0.71	3.45 ± 0.67	0.378
	Image quality, reader 2	3.45 ± 0.67	3.41 ± 0.62	0.621
Hachulla et al., <sup>15)</sup> 2019	Global quality, excellent	38/42	37/42	NR
	Global quality, moderate	4/42	5/42	NR
Ippolito et al., <sup>17)</sup> 2020	Image quality, reader 1	3.44 ± 0.73	3.48 ± 0.71	0.840
	Image quality, reader 2	3.50 ± 0.53	3.45 ± 0.67	0.562
Suchá et al., <sup>18)</sup> 2020	Annular image quality	3.0 (range 2–4)	3.5 (range 2–4)	NR
	Iliofemoral image quality	3.5 (range 2–4)	4.0 (range 2.5–4)	NR

IQR: interquartile range, NR: not reported.

assessment was similar for the low dose protocols when compared to conventional protocols. Felmly et al.<sup>12)</sup> did report increased subjective image quality for aortoiliac noise for their low dose protocol when compared to the conventional protocol (low dose: 4.42 [IQR, 4.0–5.0], conventional 4.12 [IQR, 4.0–5.0],  $p = 0.037$ ).

## DISCUSSION

This systematic review suggests adequate image quality for low contrast and low kV CTA compared to conventional CTA for TAVR planning. Our findings are derived from 6 studies reporting on image quality in 353 patients with aortic stenosis undergoing CTA for TAVR planning. There was no difference in image quality as assessed by cardiac, aortic, and iliofemoral SNR and cardiac and aortic CNR between imaging protocols for this patient population. There was decreased iliofemoral CNR for the low dose protocols when compared to the conventional protocols. This is likely from improved CNR with higher contrast and kV amounts used in the conventional protocols at this specific anatomic location. Interestingly, Ippolito et al.<sup>16)</sup> were able to achieve significantly higher SNR and CNR with their low dose protocol combined with a model-based iterative reconstruction algorithm, specifically IMR.<sup>16)</sup> Overall subjective image quality was similar with high quality scores for both protocols. Image quality was acceptable for TAVR planning using the low dose protocols in all of the included studies.

These findings are consistent with additional studies that achieved adequate image quality using intra-arterial catheters to deliver low contrast volumes compared to conventional CTA for TAVR planning.<sup>18)19)</sup> Recently developed CT technology using virtual monoenergetic images at lower kV levels has been shown to result in better image quality at lower doses of contrast material.<sup>20)21)</sup> This is feasible since the K-edge of iodine

at 33.3 kV is closer to lower x-ray energy spectra, therefore increased soft tissue contrast can be obtained at decreased keV levels.<sup>22)23)</sup> Cavallo et al.<sup>24)</sup> described a method of using 40 keV reconstruction with low contrast volume compared to conventional 120 kV images with the same contrast volume in patients undergoing TAVR planning. The 40 keV method with low dose contrast achieved better image quality as assessed by SNR and CNR. Mangold et al. developed another 40 keV virtual monoenergetic image reconstruction technique that obtained higher image quality when compared to a conventional protocol for TAVR planning.<sup>25)</sup> Several studies have also demonstrated the feasibility of low radiation and low contrast CTA for TAVR planning, although comparisons to conventional imaging were not made.<sup>26-28)</sup> These findings also support a recent expert consensus document that recommends optimizing scanning protocols to achieve lower overall contrast volumes.<sup>29)</sup>

This systematic review provides important insights on using a low contrast and low kV CTA strategy for patients undergoing TAVR planning that may inform decisions in clinical practice. Using low radiation through low-tube-voltage techniques maintains image quality at lower contrast volumes. Minimizing contrast volume can help make TAVR available to patients with renal dysfunction who would otherwise not be candidates since there is a dose-dependent association with contrast-induced nephropathy.<sup>30)</sup> As TAVR expands to younger patient populations, the reduced radiation dose used by this imaging strategy may also become a more relevant clinical benefit.

The limitations of this systematic review are influenced by the limitations of the included studies. The included studies had small sample sizes likely due to the complexity of image quality research for this patient population. One major limitation is the variability in the low dose and conventional protocols between studies. Various contrast amounts, tube voltages, and image acquisition techniques were used and severely limits the generalizability of

the aggregate data. Variations in clinical settings, population characteristics, and imaging protocols are potential sources of heterogeneity. We used a random effects model to account for these variations. CIs for estimates were wide in spite of pooling likely due to high heterogeneity and low sample sizes.

In conclusion, this systematic review suggests that low contrast and low kV CTA for TAVR planning provides similar image quality to conventional CTA.

## SUPPLEMENTARY MATERIALS

### Supplementary Figure 1

Forest plot of aortic signal-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning.

[Click here to view](#)

### Supplementary Figure 2

Forest plot of aortic contrast-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning.

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### Supplementary Figure 3

Forest plot of ileofemoral signal-to-noise ratio in patients undergoing computed tomography angiography for transcatheter aortic valve replacement planning.

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#### Conflict of Interest

The authors have no financial conflicts of interest.

#### Author Contributions

Conceptualization: Lacy S, Benjamin M, Syed M, Kinno M; Data curation: Lacy S; Formal analysis: Lacy S; Investigation: Lacy S; Methodology: Lacy S; Software: Lacy S; Supervision: Benjamin M, Syed M, Kinno M; Writing - original draft: Lacy S, Benjamin M; Writing - review & editing: Benjamin M, Osman M, Syed M, Kinno M.

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