

Optimization of dual energy computed tomography post-processing to reduce lower limb artifacts in gout

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Background: In gout, several types of dual-energy computed tomography (DECT) artifacts have been described (nail bed, skin, beam hardening, submillimeter and vascular artifacts), which can lead to overdiagnosis. The objective of this study was to determine the optimal DECT settings for post processing in order to reduce the frequency of some common artifacts in patients with suspected gout.

Methods: Seventy-seven patients hospitalized for suspected gout (feet/ankles and/or knees) who received a DECT imaging were included (final diagnosis of 43 gout and 34 other rheumatic disorders). Different post-processing settings were evaluated using Syngovia software: nine settings (R1 to R9) were evaluated with a combination of different ratio (1.28, 1.36 and 1.55) and attenuation coefficient (120, 150, 170 HU).

Results: Among the nine settings tested, the R2 setting (170 HU, ratio =1.28) significantly reduced the presence of knee and foot/ankle artifacts compared to the standard R1 setting (85% and 94% decrease in beam hardening and clumpy artifacts in the ankle and foot, respectively (P<0.001); a decrease of 71%, 60% and 88% respectively of meniscal beam hardening, beam hardening and submillimeter artifacts in the knee (P<0.001). Compared to standard settings, the use of R2 settings decreased sensitivity [0.79 (95% CI: 0.65, 0.88) versus 0.90 (95% CI: 0.78, 0.96)] and increased specificity [0.86 (95% CI: 0.71, 0.93) versus 0.63 (95% CI: 0.47, 0.77)] (P<0.001). Settings using an attenuation coefficient to 120 HU and/or a ratio to 1.55 were all associated with a significant increasing of artifacts, especially clumpy and beam hardening artifacts.

Conclusions: Applying a ratio of 1.28 and a minimum attenuation of 170 HU in DECT post-processing eliminates the majority of artifacts located in the lower limbs, particularly clumpy artifacts and beam hardening.

Keywords: Gout; artifacts; dual energy computed tomography; post-processing

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Introduction

Gout, one of the most common inflammatory rheumatic conditions, is characterized by intra-articular or periarticular monosodium urate (MSU) deposits (1). The prevalence of gout, which is estimated at 0.9% in France (2), is increasing due to the aging of the population and dietary habits that have shifted towards foods which are richer in purines and fructose.

The gold standard for confirming a diagnosis of gout is based on the detection of MSU crystals under polarized light microscopy in synovial fluid aspirated from the patient's joint. However, joint puncture has certain limitations (sensitivity, inaccessible joint site) (3), may not be available in inter-critical gout, and remains poorly used in primary care practice (4). Imaging offers an alternative method to detect the presence of MSU crystals in such clinical situations (5). Among the existing imaging techniques, dual-energy computed tomography (DECT) has demonstrated over the last decade its ability to detect MSU crystals deposited in the soft tissue surrounding joints. The principle of DECT is based on the difference in X-ray absorption between two given materials (e.g., calcium and MSU). The difference in attenuation between the two materials at two different energy levels (80 and 140 kV) makes it possible to determine the nature of the material being studied.

The American College of Rheumatology/European League Against Rheumatism gout task force acknowledged the good performance of DECT for the diagnosis of gout (6-9) by including it in the 2015 classification criteria (10). A meta-analysis based on 8 studies (510 patients/268 controls) found a sensitivity and specificity of 0.847 and 0.936, respectively (11). DECT and ultrasound had comparable diagnostic performance (12). However, reproducibility is generally excellent in DECT, and better than for ultrasound (6,13).

Nevertheless, not all material color coded as urate on DECT corresponds to locations of urate deposition. These artifacts, related to either the patient or the CT acquisition (14-17), can lead to an over-diagnosis of gout (false positives), but this issue has been insufficiently evaluated in patients with suspected gout.

Artifacts were defined in the 2015 ACR/EULAR criteria (10) as follows: nail bed deposits, skin deposits, beam hardening artifacts, submillimeter points [defined as "clumpy" artifacts in the ankle and foot (18)] and vascular and movement artifacts. Identification of the most common

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artifacts and artifact-reduction techniques are therefore essential for the radiologist to avoid overdiagnosis and the unnecessary implementation of urate-lowering therapy (14,18).

Optimization of post-processing settings is an essential prerequisite for any imaging. For DECT, the increase or decrease of the attenuation coefficient (HU) has shown a decrease in some artifacts (17,18). Increasing the range allows to smooth the image, making it possible to reduce certain artifacts, but at the cost of a reduction in spatial resolution. Modifying the post-processing parameters thus makes it possible to increase the volume of the MSU deposits (increase the range, decrease the minimum HU but at the cost of an increase in the frequency of artifacts) or, on the contrary, to reduce the artifacts but at the cost of a reduction in the volume of the "real" MSU deposits. It is therefore critical to find post-processing parameters that significantly reduce the occurrence of artifacts while maintaining the sensitivity of DECT for the detection of MSU deposits in patients suspected of gouty arthritis.

The aims of this study were to provide a descriptive analysis of the artifacts in gout patients and controls and to propose an optimization of the parameters during the postprocessing of the data in order to reduce the occurrence of artifacts in lower limb dual energy CT scan.

Methods

Study design

Retrospective, monocentric study conducted in the radiology and rheumatology departments of the Dijon University Hospital (France).

All consecutive patients hospitalized for suspicion of gouty arthritis who had a DECT exam of the lower limbs (feet/ankles and/or knees) from February 2015 to April 2020 were enrolled. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by Institutional Review Board and individual consent for this study was waived.

Collected data

For each patient, demographic characteristics were collected. Cardiovascular comorbidities, serum urate level, gout duration, ultrasound findings for gout (presence of the double contour sign), radiographic data [Kellgren and Lawrence stage for knee osteoarthritis (19)] were also collected when available. A total of 102 patients underwent DECT scans for evaluation and quantification of their MSU crystal volume, but reconstructions from native sections were not possible for 25 patients who were therefore excluded from the analysis. Finally, 77 patients were included in the final analysis. Three patients had both ankle and knee DECT (80 DECT in total).

Patient classification

Patients were classified in the gout or non-gout group (other rheumatic disorders) according to the ACR 77 classification criteria (20) (results of joint puncture when performed or characteristics of the flare). The ACR/EULAR 2015 classification criteria were not used because they incorporate DECT data.

DECT and post-processing

The DECT used was a dual energy "dual source" scanner (80 kV-140/Sn kV) (SIEMENS, SOMATOM Flash definition). The acquisition time was <2 minutes per joint. Mean (SD) delivered radiation dose (DLP) for the knee was 203±27.8 mGy·cm and for the ankle/foot 179±45.6 mGy·cm. The approximate effective radiation dose reported in other studies was 0.5 mSv (21). The scanner was equipped with 2×64 detectors and $2 \times (64\times0.6 \text{ mm})$ collimations. The number of reconstructed slices was 2×128 slices/turn. Concerning the "double energy lower extremity" protocol used, the rotation time was 0.5 seconds with a collimation of 0.32×0.6 mm. The post-processing of the data was performed using Syngo Via software (version Somaris/7 Syngo CT 2012 B, VH 10 ASL 79118 Siemens Healthineers). The standard parameters used during the acquisition were: ratio 1.36; HU minimum 150; HU maximum 500 HU; distance to air and distance to bone 5; range 4.

Image analysis

All DECT images were read by experienced musculoskeletal radiologists and blinded to clinical-biological patient data and diagnosis.

In order to reduce the occurrence of artifacts and to define the most relevant parameters to be modified, different ratio settings were tested: 1.28, 1.36, and 1.55 (22,23). Concerning the attenuation coefficient (HU), we systematically applied (in association with a ratio of 1.28, 1.36 and 1.55) different settings, calibrating the minimum attenuation to 120 HU, 150 HU (default) and 170 HU.

Comparison between the nine different settings ("R1" 1.36–150 HU; "R2" 1.28–170 HU; "R3" 1.28–120 HU; "R4" 1.28–150 HU; "R5" 1.36–120 HU; "R6" 1.36–170 HU; "R7" 1.55–120 HU; "R8" 1.55–150 HU; "R9" 1.55–170 HU) was performed. The maximum attenuation (500 HU), the distance to air, the distance to bone and the range were not modified.

Statistical analyses

The inter-observer reproducibility of each examination was achieved by determining Cohen's kappa coefficient. Excellent reproducibility is indicated by kappa >0.81, good from 0.80 to 0.71, moderate from 0.70 to 0.41, poor from 0.40 to 0.21, and very poor <0.20. No imputations were performed for missing data. Qualitative variables were compared using a Chi2 or Fisher's test when appropriate. Quantitative variables were compared using a Student's T-test or Mann Whitney test when appropriate. The frequencies of artifacts according to the different parameters applied were compared using a McNemar's test. A Friedman test was performed to compare volumes for matched populations. Correlations between the presence of the artifacts and clinical features (age, sex, high blood pressure, diabetes, BMI, and severe knee osteoarthritis and calcium pyrophosphate deposits for knee localization) were obtained with a Spearman's test. Diagnostic performance based on the calculation of sensitivity, specificity, positive predictive value, negative predictive value and ROC curve was then calculated for the standard settings and for all modified settings; all tests were considered significant if their P values were below 0.05. Analyses were performed using IBM SPSS v25 statistics software.

Results

Study population (Table 1)

Of the 77 patients, 43 met ACR77 criteria for the diagnosis of gout. Among them, 26 had a joint puncture (16 knees and 10 ankle/feet). Joint fluid analysis found MSU crystals in 18 of these patients. For the remaining 34 patients, the diagnoses retained were: chondrocalcinosis (n=9; based on presence of calcium pyrophosphate (CPP) crystals on joint fluid analysis), osteoarthritis flare-up (n=18; based on X-ray results and/or cytology of joint fluid aspiration), and other chronic inflammatory rheumatic diseases (n=7; rheumatoid

Table 1 Clinical-biological and DECT characteristics of the gout and non-gout groups

	Gout (n=43)	Controls (n=34)	Р
Age, years (mean ± SD)	66.1 (15.0)	72.8 (12.7)	
Male (%)	88.4	52.9	0.001
Uricemia, mg/dL (mean ± SD)	7.79 (2.21)	5.65 (2.08)	<0.001
Artifacts, n (%)	35 (81.4)	27 (79.4)	0.82
DECT MSU* deposits, n (%)			
Absence	13 (30.2)	30 (88.2)	<0.001
Intra-articular	25 (58.1)	4 (11.8)	<0.001
Extra-articular only	5 (11.6)	1 (2.9)	0.12
Extra-articular deposits, n (%)	25 (58.1)	3 (8.8)	<0.001
ULT, n (%)	13 (30.2)	4 (11.8)	0.05
Symptom duration, months (mean ± SD)	30.7 (51.9)	13.0 (18.6)	0.06
eGFR, mL/min (mean ± SD)	75.2 (26.1)	80.7 (22.8)	0.32
HBP, n (%)	25 (58.1)	20 (58.8)	0.95
History of stroke, n (%)	6 (14.0)	4 (11.8)	0.77
Myocardial infarction in past medical history, n (%)	6 (14.0)	5 (14.7)	0.92
Diabetes, n (%)	10 (23.3)	11 (32.4)	0.37
BMI, n (%)	28.4 (3.8)	28.02 (5.1)	0.69
Ultrasound DC sign, n (%)	9 (20.9)	1 (2.9)	0.02
Radiological OA grading (knee) KL III and IV, n (%)	6/18 (33.3)	9/20 (45)	0.15
Volume MSU deposits average in cm ³ (SD) (artifacts included)	0.56 (0.72)	0.15 (0.23)	0.003

Osteoarticular ultrasound was performed on 59 of 77 patients (76.7% of cases) looking for the double contour sign. Quantitative values are expressed as a mean (± standard deviation SD). Significance: P<0.05. *, Artifacts excluded. MSU, monosodium urate; ULT, urate-lowering therapy; HBP, high blood pressure; DC, double contour sign; KL, Kellgren and Lawrence radiological grading, severe for stages 3 and 4; BMI, body mass index (kg/m²); eGFR, estimated glomerular filtration rate (CKD-EPI).

arthritis, ankylosing spondylarthritis, psoriatic arthritis). Patient characteristics are presented in *Table 1*.

Diagnostic performance of R1 to R9 settings (Table 2 and Figures S1-S3)

Diagnostic performance of each setting was calculated separately for each location (ankle/foot and knees) and for combination of both ("full").

Best diagnostic performance was obtained with the R2 settings [AUC foot/ankle =0.81 (P<0.001); AUC knee =0.89 (P<0.001); AUC "foot/ankle + knee =0.84 (P<0.001)]. However, the use of R2 settings decreased sensitivity [0.79 (95% CI: 0.65, 0.88) versus 0.90 (95% CI: 0.78, 0.96)] and increased specificity [0.86 (95% CI: 0.71, 0.93) versus 0.63

(95% CI: 0.47, 0.77)] compared to R1 settings (P<0.001). Settings using 120 HU attenuation coefficients were all associated to an increase of artifacts especially clumpy and beam hardening artifacts.

Foot and ankle artifacts (42 DECT) (Table 3 and Table S1)

For inter-observer reproducibility of artifacts identification (using default "R1" settings), kappa coefficients were calculated and showed good to excellent reproducibility: Nail bed: kappa =1.00 (95% CI: 1.00, 1.00), skin: kappa =0.90 (95% CI: 0.73, 1.00), clumpy: kappa =0.81 (95% CI: 0.63, 0.99), beam hardening: kappa =0.84 (95% CI: 0.65, 0.95).

The application of "R2" settings in post-processing

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Table 2 Diagnostic performance from R1 (standard setting) to R9 (modified setting) in ankle/foot, knee DECT and the combination of both locations ("full")

	Sensitivity (95% CI)	Specificity (95% CI)	PPV (95% CI)	NPV (95% CI)	AUC
R1 ankle/foot	0.84 (0.66, 0.94)	0.75 (0.50, 0.89)	0.84 (0.69, 0.92)	0.75 (0.53, 0.88)	0.78 (0.65, 0.95)
R1 knee	1.00 (0.82, 1.00)	0.55 (0.34, 0.74)	0.66 (0.55, 0.76)	1.00 (1.00, 1.00)	0.77 (0.62, 0.93)
R2 ankle/foot	0.69 (0.50, 0.83)	0.93 (0.72, 0.98)	0.94 (0.72, 0.99)	0.65 (0.50, 0.77)	0.81 (0.68, 0.94)
R2 knee	0.94 (0.74, 0.99)	0.85 (0.63, 0.91)	0.85 (0.64, 0.91)	0.94 (0.74, 0.99)	0.89 (0.78, 1.00)
R3 ankle/foot	1.00 (0.87, 1.00)	NC	0.61 (0.47, 0.75)	NC	0.50 (0.32, 0.69)
R3 knee	1.00 (0.82, 1.00)	NC	0.47 (0.32, 0.62)	NC	0.50 (0.31, 0.69)
R4 ankle/foot	0.81 (0.62, 0.91)	0.44 (0.23, 0.66)	0.70 (0.52, 0.83)	0.58 (0.31, 0.80)	0.62 (0.44, 0.80)
R4 knee	1.00 (0.82, 1.00)	0.40 (0.22, 0.61)	0.60 (0.42, 0.75)	1.00 (0.67, 1.00)	0.7 (0.53, 0.87)
R5 ankle/foot	1.00 (0.87, 1.00)	NC	0.61 (0.47, 0.75)	NC	0.50 (0.32, 0.69)
R5 knee	1.00 (0.82, 1.00)	NC	0.47 (0.32, 0.62)	NC	0.50 (0.31, 0.69)
R6 ankle/foot	0.77 (0.57, 0.88)	0.75 (0.51, 0.89)	0.83 (0.64, 0.93)	0.67 (0.43, 0.83)	0.76 (0.60, 0.92)
R6 knee	1.00 (0.82, 1.00)	0.35 (0.18, 0.56)	0.58 (0.41, 0.73)	1.00 (0.64, 1.00)	0.67 (0.50, 0.85)
R7 ankle/foot	1.00 (0.62, 1.00)	NC	0.61 (0.47, 0.75)	NC	0.50 (0.32, 0.69)
R7 knee	1.00 (0.82, 1.00)	NC	0.47 (0.32, 0.62)	NC	0.50 (0.31, 0.69)
R8 ankle/foot	0.93 (0.75, 0.97)	0.14 (0.01, 0.28)	0.61 (0.45, 0.75)	0.33 (0.06, 0.79)	0.49 (0.31, 0.67)
R8 knee	1.00 (0.82, 1.00)	NC	0.47 (0.32, 0.62)	NC	0.50 (0.31, 0.69)
R9 ankle/foot	0.96 (0.81, 0.99)	0.19 (0.06, 0.43)	0.65 (0.49, 0.78)	0.75 (0.30, 0.95)	0.57 (0.39, 0.76)
R9 knee	1.00 (0.82, 1.00)	NC	0.47 (0.32, 0.62)	NC	0.50 (0.31, 0.69)
R1 "full"	0.90 (0.78, 0.96)	0.63 (0.47, 0.77)	0.75 (0.66, 0.82)	0.85 (0.68, 0.93)	0.77 (0.66, 0.88)
R2 "full"	0.79 (0.65, 0.88)	0.86 (0.71, 0.93)	0.87 (0.75, 0.94)	0.77 (0.65, 0.86)	0.84 (0.75, 0.93)

R1 setting =ratio 1.36; HU =150; bone distance =5; air distance =5; range =4; R2 setting =ratio 1.28; HU =170; bone distance =5; air distance =5; range =4; R3 setting=ratio 1.28; HU =120; bone distance =5; air distance =5; range =4; R4 setting=ratio 1.28; HU =150; bone distance =5; air distance =5; air distance =5; range =4; R5 setting=ratio 1.36; HU =120; bone distance =5; air distance =5; range =4; R6 setting=ratio 1.36; HU =170; bone distance =5; air distance =5; range =4; R6 setting=ratio 1.36; HU =170; bone distance =5; air distance =5; range =4; R7 setting=ratio 1.55; HU =120; bone distance =5; air distance =5; range =4; R8 setting=ratio 1.55; HU =150; bone distance =5; air distance =5; air distance =5; air distance =5; air distance =5; range =4; R8 setting=ratio 1.55; HU =150; bone distance =5; air dista

Table 3 Comparison of the frequency of different ankle and/or foot artifacts according to the standard setting (R1) and the modified setting R2 and R3 (n=42)

	R1	R2	R3	Р
Average volume of MSU cm ³ (SD)	0.33 (0.50)	0.14 (0.25)	0.94 (0.95)	<0.001
Nail bed n (%)	30 (71.4)	24 (57.1)	31 (73.8)	<0.001
Skin n (%)	6 (14.3)	1 (2.4)	11 (26.2)	<0.001
Beam hardening n (%)	14 (33.3)	2 (4.8)	17 (40.5)	<0.001
Clumpy n (%)	17 (40.5)	1 (2.4)	39 (92.9)	<0.001

R1 is the standard setting (ratio at 1.36 and minimum attenuation set to 150 HU). R2 is the modified setting (ratio at 1.28 and minimum attenuation set to 170 HU). R3 is the standard setting (ratio at 1.28 and minimum attenuation set to 120 HU). SD, standard deviation. Significance: P<0.05.



Figure 1 Ankle sagittal view. Beam hardening artifact (arrows) (calcaneum) (A) with a talus gout deposition. Application of "R2" settings led to artifact removal (arrows). Note the persistence of a pathological deposit of MSU on the posterior aspect of the talus at setting R2 (asterisk). (A) R1 settings (150 HU, ratio 1.36); (B) R2 settings (170 HU, ratio 1.28).

significantly reduced almost all the described ankle and foot artifacts. Nail bed artifacts were reduced by 20%, skin artifacts by 83.3%, beam hardening by 85.6%, and submillimeter artifacts by 94.1% (*Figure 1*). However, the application of the "R2" settings (170 HU, ratio 1.28) was accompanied by a significant decrease in total MSU volume (reduced by 58.7%) (*Table 3, Figure 2*).

The application of the "R3" setting with ratio of 1.28 and minimum attenuation of 120 HU found a significant increase in the volume of MSU deposits [mean R1 *vs.* mean R3; 0.33 ± 0.50 *vs.* 0.94 ± 0.95 cm³ (P<0.001)], a significant increase in the occurrence of clumpy artifacts (increased by 129%) and a non-significant increase in the frequency of skin artifacts, beam hardening and nail beds (*Table 3*). The results of "R4" to "R9" settings are summarized in Table S1. No correlations were found between foot/ankle artifacts and clinical features (Table S2).

Knee artifacts (38 DECT) (Table 4 and Table S3)

Kappa coefficients for inter-reader reproducibility showed moderate to good reproducibility: meniscal beam hardening: kappa =0.77 (95% CI: 0.54, 0.94), beam hardening: kappa =0.78 (95% CI: 0.56, 0.99), submillimeter artifacts: kappa=0.67 (95% CI: 0.47, 0.83), vascular artifacts: kappa=0.63 (95% CI: 0.25, 0.92). The application of R2 settings in post-processing significantly reduced the set of artifacts defined by the ACR/ EULAR 2015 criteria (*Figures 3,4*). Meniscal beam hardening artifacts were reduced by 71%, beam hardening by 60%,

submillimeter artifacts by 88.9%, and vascular artifacts by 100%. However, the application of the "R2" setting resulted in a significant reduction in the volume of MSU deposits, as in the ankle/foot area (volume reduced by 60%) (*Table 4*).

In contrast, application of R3 settings significantly increased meniscal beam hardening artifacts (38%) and essentially clumpy artifact (300%). The results of "R4" to "R9" settings are summarized in Table S3.

The presence of articular chondrocalcinosis (n=9) was significantly associated with the occurrence of meniscal beam hardening artifacts (Spearman's Rho σ =0.501; 95% CI: 0.307, 0.666; P=0.001). Similarly, BMI was associated with the occurrence of beam hardening artifacts (excluding menisci) in the knee (σ =0.430; 95% CI: 0.09, 0.668; P=0.007) (Table S4).

Discussion

The aim of this case-control study in a real-life setting was to optimize DECT post-processing parameters in order to reduce the frequency of some artifacts commonly occurring in DECT performed for gout assessment. According to our results, lowering the ratio to 1.28 and increasing the minimum attenuation to 170 HU could eliminate artifacts that interfere with the proper interpretation of the DECT, and thus reduce the risk of false positives by radiologists.

This adjustment of the ratio and attenuation coefficient settings showed a gain in specificity (from 63.9% to 86.1%) at the expense of sensitivity, which dropped from 90.9% Quantitative Imaging in Medicine and Surgery, Vol 12, No 1 January 2022



Figure 2 DECT axial section of ankle and foot. The application of "R2" settings showed a decrease in MSU deposit volume (head of arrow) (B,D,F) in comparison to standard settings "R1" (A,C,E).

Table 4 Com	parison of the	frequency	7 of different ki	nee artifacts	according to t	he standard :	setting (R1)	and the modified	setting	(R2)(n=38)
	1				0				0	

	R1	R2	R3	Р
Average MSU volume cm ³ (SD)	0.42 (0.64)	0.17 (0.35)	0.82 (0.97)	<0.001
Meniscal beam hardening, n (%)	21 (55.3)	6 (15.8)	29 (76.3)	<0.001
Beam hardening, n (%)	25 (65.8)	10 (26.3)	25 (65.8)	<0.001
Clumpy, n (%)	9 (23.7)	1 (2.6)	36 (94.7)	<0.001
Vascular, n (%)	7 (18.4)	0	8 (21.1)	0.001

R1 is the standard setting (ratio at 1.36 and minimum attenuation set to 150 HU). R2 is the modified setting (ratio at 1.28 and minimum attenuation set to 170 HU). R3 is the modified setting (ratio at 1.28 and minimum attenuation set to 120 HU). SD, standard deviation. Significance: P<0.05.

to 79.5%. The initial specificity (63.5%) is lower than what has been reported in previous DECT studies, but the studied populations are not comparable. Indeed, most of the authors who found specificity >80% were studying patients with a known diagnosis of long-lasting and severe gout. In patients with undifferentiated arthritis, Gamala *et al.* (24) found that DECT had a lower specificity (<60%), close to

our results.

Despite the R2 settings (ratio 1.28 and 170 HU), the most common artifacts (14) such as nail bed deposits and subcutaneous artifacts persist, but they are generally easily identifiable by MSK radiologists. As extra-articular manifestation of gout are common on DECT imaging, it seems particularly important to minimize the clumpy



Figure 3 Sagittal knee sections in DECT in a "non-gout" patient. (A) Enhancement of the beam hardening (arrows) and submillimiter artifacts (head of arrows) using the R3 settings; (B) presence of beam hardening (arrows) and submillimeter artifacts (head of arrows) with standard settings (R1); (C) absence of artifacts with setting R2 (HU=170; ratio =1.28).



Figure 4 Coronal view of meniscal beam hardening artifact (arrow) (A) with standard settings "R1" (ratio: 1.36, HU=150). (B) Artifact removal with application of "R2" settings (ratio 1.28, HU=170). Joint puncture demonstrated presence of multiple calcium pyrophosphate crystals with no evidence of MSU crystals.

artifacts that appear within tendons and ligaments in the foot and ankle and also meniscal artifacts. In routine practice they are easily confused with actual urate deposits within these fibro-cartilage structures. Another effective method to eliminate or reduce clumpy artifacts is the use of tin filter, as reported by Park *et al.* (18) and Jeon *et al.* (25). Nevertheless, in our experience, clumpy artifacts may persist despite the tin filter.

In order to optimize the post-processing parameters, Park *et al.* (18) suggested increasing the minimum attenuation from 130 to 150 HU. The application of these parameters makes it possible to decrease the frequency of some artifacts. The authors also found a significant decrease (P<0.007) in the volume of MSU deposits, which is also the case in our cohort. This limitation needs to be considered because attempting to optimize the specificity of the DECT in gout can reduce its sensitivity, as demonstrated again. Our results show a significant decrease in MSU volume using the "R2" setting compared to the "R1" setting (*Tables 3,4*). Despite this decrease in MSU volume, it appears that most of the large MSU deposits may remain (*Figure 2*). Nevertheless, in the case of minimal deposits, the use of these parameters on DECT may lead to a risk of false negatives for the diagnosis of gout. However, the tophaceous volume alone is probably not sufficient to explain the disappearance of R2 deposits, as the density of the deposits is likely to also play a role (26).

According to our results, submillimeter artifacts are significantly reduced or even eliminated in most cases by the settings retained (ratio at 1.28; HU=170). It appears that decreasing the ratio reduced beam hardening artifacts, and decreasing the attenuation coefficient reduced clumpy artifacts. These observations may be explained by tissue attenuation principle that depends on the density (Zeff) of tissue and X-ray beam energy (KeV). Indeed, trabecular bone, meniscal CPP deposits and UMS deposits have close densities (at 80 and 140 kV). Decrease the ratio allows to a better differentiation between calcium deposits (CPP and trabecular bone) and MSU deposits. On the contrary, increasing ratio do not allow to distinguish between them and lead to false green coloration deposits (meniscal CPP deposits).

The settings used in this study proved to be effective for the ankle and foot but also for the knee, which had not been reported until now. It is also of interest to use the same parameters for these two-common locations of gouty arthritis, taking into account that it is usual practice to request a DECT of the knee and ankle for the evaluation of suspected gout. Other studies have shown the interest of modifying other parameters in post-processing in order to reduce artifacts. Mallinson *et al.* (14) noted that reducing the distance to air and the distance to bone reduced subcutaneous and under-nail bed artifacts. These different parameters were not tested in our study because these common artifacts are quite simple to identify and remove in routine imaging practice.

Another study looked at the post-processing parameters of DECT data in typical gout disease (involvement of first metatarsophalangeal joint) (27). The application of a minimum attenuation at 120 HU would allow a better visualization of the MSU deposits and a better correlation in terms of volume with ultrasound. Indeed, DECT tends to underestimate the volume of a tophus by visualizing only the "crystalline" part, unlike ultrasound which also takes into account the adjacent soft parts (28). In our experience, the application of these parameters does not seem to be relevant to the rest of the foot, the ankle, and the knee, because it leads to a significant increase in noise that hinders interpretation (clumpy artifacts especially).

It appears that some artifacts are not very sensitive to post-processing modifications, such as artifacts related to metallic materials and movement artifacts. In these cases, the best way to limit these artifacts would be either to modify the parameters during image acquisition (modification of the rotation speed of the slats, use of specific kernel, tin filter, etc.) (14,15), or to ensure the correct positioning of the patient (29).

We recognize that this study has some limitations,

including the recruitment in a single university center and the low number of patients. The use of ACR 77 classification criteria can be an important limitation. Indeed, there is a risk that some patients classified as "controls" might have gout, notably in cases of short duration of the condition.

Another fundamental aspect for the better detection of artifacts is the identification of patient-related risk factors. Our results identified certain factors associated with the occurrence of knee artifacts, including the significant correlation between beam hardening artifacts and BMI value, which has not been reported previously. One possible explanation for the correlation between BMI and beam hardening artifact could be that the increase of soft tissue (with an increase of density) lead to beam hardening. Another possibility is be that cortical bone is thicker in patients with a high BMI (maybe due to higher mechanical strains) than in subjects with a normal BMI.

Similarly, a correlation was found between the frequency of meniscal beam hardening artifacts and articular chondrocalcinosis in the knee. On the other hand, and contrary to the hypothesis of Bongartz *et al.* (7), no correlation was found between the existence of severe knee osteoarthritis (stage 3 and 4 of Kellgren and Lawrence) and the presence of meniscal beam hardening artifacts. Our result could nevertheless be partly explained by a lack of power of the present study, which remains to be confirmed.

In conclusion, we confirmed the high frequency of artifacts in DECT for the evaluation of suspected gout. Artifact-reduction techniques are therefore essential for mitigating their occurrence and misinterpretation. Indeed, the application of a ratio of 1.28 and a minimum attenuation of 170 HU makes it possible to eliminate some DECT artifacts in the foot, ankle and knee joints, with a significant gain in specificity. Nevertheless, the application of R2 parameters results in a drop in sensitivity, especially for foot/ankle, which can be a limitation for these localizations. Further advances in DECT acquisition and post-processing techniques are necessary to increase the accuracy of this second-line diagnostic approach in cases where the diagnosis is uncertain.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by Institutional Review Board and individual consent for this study was waived.

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AUC: area under the curve; 95%CI: 95% confidence interval Significance: P<0.05

Figure S1 Comparative analysis of diagnostic performance from R1 to R9 settings for foot/ankle [ROC and area under the curve (AUC)].



AUC: area under the curve; 95%CI: 95% confidence interval Significance: P<0.05

Figure S2 Comparative analysis of diagnostic performance from R1 to R9 settings for knee [ROC and area under the curve (AUC)].



Setting	AUC	р	95%CI
R1	0.77	< 0.001	(0.66 ; 0.88)
R2	0.84	< 0.001	(0.75 ; 0.93)
R3	0.50	1.00	(0.37 ; 0.62)
R4	0.65	0.02	(0.52 ; 0.77)
R5	0.50	1.00	(0.37 ; 0.62)
R6	0.69	0.003	(0.57 ; 0.81)
R7	0.50	1.00	(0.37 ; 0.62)
R8	0.49	0.88	(0.36 ; 0.61)
R9	0.53	0.64	(0.40 ; 0.66)

AUC: area under the curve; 95% CI: 95% confidence interval Significance: P<0.05

Figure S3 Comparative analysis of diagnostic performance from R1 to R9 settings for foot/ankle + knee [ROC and area under the curve (AUC)].

Table S1 Comparison of the frequency of different ankle and/or foot artifacts according to the R1 to R9 settings (n=42)

•	-	•				-				
	R1	R2	R3	R4	R5	R6	R7	R8	R9	Р
Nail bed n (%)	30 (71.4)	24 (57.1)	31 (73.8)	23 (54.7)	28 (66.6)	21 (50.0)	29 (69.0)	28 (66.6)	26 (61.9)	<0.001
Skin n (%)	6 (14.3)	1 (2.4)	11 (26.2)	3 (7.1)	7 (16.6)	2 (4.7)	7 (16.6)	4 (9.5)	2 (4.7)	<0.001
Beam hardening n (%)	14 (33.3)	2 (4.8)	17 (40.5)	6 (14.2)	13 (30.9)	5 (11.9)	42 (100)	38 (90.4)	34 (80.9)	<0.001
Clumpy n (%)	17 (40.5)	1 (2.4)	39 (92.9)	10 (23.8)	37 (88.1)	1 (2.3)	40 (95.2)	6 (14.2)	1 (2.3)	<0.001
Average volume of MSU Cm ³ (SD)	0.33 (0.50)	0.13 (0.25)	0.94 (0.95)	0.26 (0.40)	1.18 (1.08)	0.20 (0.36)	2.57 (1.93)	1.00 (1.17)	0.67 (0.92)	<0.001

SD, standard deviation. Significance: P<0.05.

Table S2 Correlation between most common artifacts and clinical features for the foot/ankle

		Age	Sex	HBP	Diabetes	BMI
Nails	Spearman σ	0.16	0.08	0.18	0.22	-0.03
	95% CI	[-0.16, 0.47]	[-0.24, 0.39]	[-0.13, 0.48]	[-0.07, 0.41]	[-0.31, 0.27]
	Р	0.31	0.60	0.25	0.17	0.84
Skin	Spearman σ	-0.12	-0.05	0.02	-0.05	0.10
	95% CI	[-0.40, 0.19]	[-0.42, 0.25]	[-0.28, 0.32]	[-0.28, 0.31]	[-0.27, 0.44]
	Р	0.44	0.73	0.89	0.72	0.52
Beam	Spearman σ	0.20	-0.17	-0.02	0.14	-0.01
hardening	95% CI	[-0.12, 0.41]	[-0.57, 0.15]	[-0.33, 0.29]	[-0.18, 0.45]	[-0.32, 0.29]
	Р	0.21	0.29	0.90	0.40	0.95
Clumpy	Spearman σ	-0.18	0.24	-0.013	-0.04	0.20
	95%CI	[-0.47, 0.13]	[-0.05, 0.45]	[-0.32, 0.29]	[-0.34, 0.28]	[-0.13, 0.51]
	Р	0.26	0.12	0.93	0.77	0.21

HBP, high blood pressure; BMI, body mass index; 95% CI: 95% confidence interval. Significance: P<0.05.

Table S3 Comparison of the frequency of different knee artifacts according to the R1 to R9 settings (n=38)

*	*	•								
	R1	R2	R3	R4	R5	R6	R7	R8	R9	Р
Meniscal beam hardening n (%)	21 (55.3)	6 (15.8)	29 (76.3)	16 (42.1)	30 (78.9)	15 (39.5)	33 (86.8)	20/ (52.6)	19 (50)	<0.001
Beam hardening n (%)	25 (65.8)	10 (26.3)	25 (65.8)	15 (39.5)	27 (71.1)	17 (44.7)	38 (100)	38 (100)	38 (100)	<0.001
Clumpy n (%)	9 (23.7)	1 (2.6)	36 (94.7)	4 (10.5)	38 (100)	3 (7.9)	38	6 (15.8)	0 (0)	<0.001
Vascular n (%)	7 (18.4)	0 (0)	8 (21.1)	2 (5.3)	7 (18.4)	3 (7.9)	14 (36.8)	9 (23.7)	9 (23.7)	<0.001
Average volume of MSU, cm ³ (SD)	0.42 (0.64)	0.17 (0.35)	0.82 (0.97)	0.28 (0.54)	1.12 (1.17)	0.24 (0.42)	3.21 (2.16)	1.64 (1.31)	1.17 (0.95)	<0.001

Table S4 Correlation between most common artifacts and clinical features for the knee

		Age	Sex	HBP	Diabetes	BMI	Severe osteoarthritis	CPP
Beam hardening	Spearman σ	0.12	0.07	0.27	0.05	0.43	0.25	-0.025
	95% CI	[-0.20, 0.41]	[-0.26, 0.41]	[-0.05, 0.58]	[-0.28, 0.36]	[0.09, 0.66]	[-0.06, 0.54]	[–0.56, 0.12]
	Р	0.40	0.66	0.10	0.75	0.007	0.14	0.12
Meniscal beam	Spearman σ	0.138	-0.057	0.126	-0.021	-0.04	0.185	0.50
hardening	95% CI	[-0.20, 0.45]	[-0.38, 0.27]	[-0.18, 0.42]	[-0.32, 0.29]	[-0.36, 0.27]	[-0.12, 0.51]	[0.31, 0.66]
	Р	0.40	0.73	0.45	0.90	0.77	0.26	0.001
Clumpy	Spearman σ	0.13	0.19	-0.05	0.25	0.14	0.05	-0.16
(or submillimiter artifact)	95% CI	[-0.14, 0.41]	[-0.09, 0.40]	[-0.41, 0.27]	[-0.11, 0.56]	[-0.16, 0.42]	[-0.28, 0.41]	[–0.37, 0.15]
	Р	0.53	0.41	0.75	0.13	0.39	0.73	0.32
Vascular	Spearman σ	0.16	0.13	0.004	-0.20	-0.06	0.17	-0.10
	95% CI	[-0.11, 0.42]	[-0.16, 0.35]	[-0.36, 0.28]	[-0.40, 0.11]	[-0.33, 0.23]	[-0.17, 0.47]	[-0.33, 0.21]
	Р	0.34	0.43	0.98	0.23	0.71	0.30	0.53

HBP, high blood pressure; BMI, body mass index; CPP, Calcium Pyrophosphate deposits; 95% CI: 95% confidence interval. Significance: P<0.05.



Illustration of meniscal beam hardening artifact in knee (image A and B) (arrows).



Illustration of beam hardening artifact in ankle (A, B) and knee (C) (arrows). Beam hardening follows the path of the hardened beam and was seen mainly within cortical bone. The beam hardening artifact was defined by a linear green deposit following cortical bone.