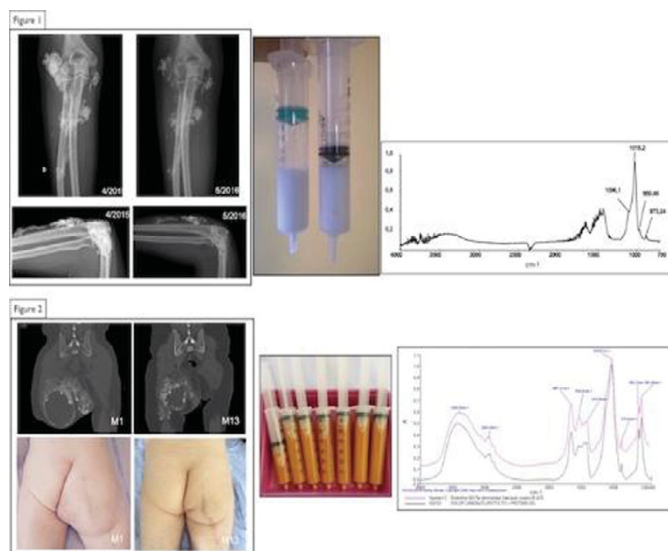


**Methods:** We report two cases of successful intra-lesional injections of sodium thiosulfate (STS).

**Results:** Case 1: A 44-years old woman, with history of dermatomyositis developed in 2009 several TC involving the extensor surfaces of the right elbow and forearm, the two ischiatic regions, fingers and lumbar back. As TC kept growing despite successive treatments, including intravenous STS, we proposed weekly intra-lesional injections of 10% STS in the elbow's lesion. The injected volume varied from 10 to 30 mL at each session. Fourier transform infrared spectroscopy (FTIR) of the aspirations confirmed that TC was composed of carbonated-apatite crystals. After 6-month treatment, we observed clinical and radiological regression of elbow TC whereas bottom TC and finger calcifications were unchanged (Fig 1). No side effect was observed except a subcutaneous infection occurring after an injection during the fifth month. This infection resolved with anti-staphylococcal antibiotics. Calcium, bicarbonate and chlorus serum levels, anion gap and eGFR remained unchanged during STS treatment.

Case 2: A 42-year old male presented with a prolonged history of hyperphosphatemic familial TC, confirmed by genetic analysis revealing homozygous mutations in the gene encoding the fibroblast growth factor 23. The extraosseous calcifications comprised a plain lesion on the right side of the left tibia and a massive heterogeneous lesion in the right buttock, making it the sitting position impossible. Treatment with maximal dose of phosphate binders, diet prescription and an attempt to surgically remove of the buttock's lesion were unsuccessful. Topical application of STS led to a near complete disappearance of the tibial lesion but was inefficient to treat the TC at the buttock. Considering the potential efficacy of STS in this patient, local injections of 25% STS (12 mL every week) were performed. The calcified material aspiration's analysis confirmed that TC was composed of carbonated-apatite crystals. Calcium and bicarbonate serum levels, anion gap and eGFR remained unchanged. After a 12-month treatment of STS injections, the lesion had significantly regressed (Fig 2), allowing the patient to sit again with no pain.



**Conclusions:** Intra-lesional injection of STS seems to be a promising treatment for TC. More studies are needed to confirm these results, and to understand the mechanisms implicated in the calcinosis resorption.

**Disclosure of Interest:** None declared

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#### AB0883 THE SENSITIVITY OF GOUT SPECIFIC ULTRASOUND SIGNS AT METATARSOPHALANGEAL JOINTS WOULD BE BETTER BY THE DORSAL SURFACE EXAMINATION

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**Objectives:** To compare the prevalence of ultrasonographic gout specific signs at the dorsal and plantar surfaces of the metatarsophalangeal joints (MTP).

**Methods:** This is a cross-sectional study which includes 15 patients with chronic gout, defined according to the American College of Rheumatology criteria (ACR 1977). Ultrasound (US) examination was performed using a high-frequency linear probe (Toshiba Xario<sup>®</sup>, frequency (8–14 MHz)) in B mode. 150 articular sites were studied at their dorsal and plantar surfaces. The ultrasound has objectified the presence of two signs: hyperechoic band over the superficial margin of the articular cartilage described as a double contour (DC) and the tophaceous deposits at the joint cavity. We compared the prevalence of the two signs between the dorsal and palmar surfaces at each site studied.

**Results:** The mean age at onset was 54.7±12,6 years, and the median diagnosis duration was 0 (0.3) years.

The results of the US examination are summarized in Table 1.

Table 1. Prevalence comparison of DC and tophaceous deposits between dorsal and plantar surfaces at MTP joints

Joints (N=150)	Double contour (%)			Tophaceous deposits (%)		
	Dorsal surface	Plantar surface	P	Dorsal surface	Plantar surface	P
MTP 1 (N=30)	33,3	10	0,03	56,7	6,7	0,01
MTP 2 (N=30)	13,3	0	<0.001	13,3	0	<0.001
MTP 3 (N=30)	6,7	0	<0.001	3,3	0	<0.001
MTP 4 (N=30)	0	0	<0.001	6,7	0	<0.001
MTP 5 (N=30)	3,3	0	<0.001	10	0	<0.001

**Conclusions:** Our study suggests that globally, DC predilect significantly in dorsal than in plantar surfaces of MTP joints. These results should be verified on a larger population.

**Disclosure of Interest:** None declared

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#### AB0884 PREVALENCE OF ULTRASONOGRAPHIC GOUT SPECIFIC SIGNS OF HAND AND FINGERS JOINTS

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**Objectives:** To evaluate the prevalence of ultrasonographic gout specific signs of hand and fingers joints.

**Methods:** This is a cross-sectional study which includes 15 patients with chronic gout, defined according to the American College of Rheumatology criteria (ACR 1977). Ultrasound (US) examination was performed using a high-frequency linear probe (Toshiba Xario<sup>®</sup>, frequency (8–14 MHz)) in B mode. 540 articular sites were studied at their dorsal surface. The ultrasound has objectified the presence of two signs: hyperechoic band over the superficial margin of the articular cartilage described as a double contour (DC) and the tophaceous deposits at the joint cavity.

**Results:** The mean age at onset was 54.7±12,6 years, and the median diagnosis duration was 0 (0.3) years.

The results of the US examination are summarized in Table 1.

Table 1. Prevalence of ultrasonographic gout specific signs of the wrist, MCP, PIP and DIP joints in the studied population

Joints (N=540)	Double contour (%)	Tophaceous deposits (%)
<b>Wrist joints (N=120)</b>	12,6	36
Radiocarpal (N=30)	20	43,3
Ulnocarpal (N=30)	13,3	50
Scaphotrapezial (N=30)	3,3	16,7
Trapeziometacarpal (N=30)	13,3	43,3
<b>MCP (N=150)</b>	8	18
MCP 1 (N=30)	3,3	16,7
MCP 2 (N=30)	13,3	33,3
MCP 3 (N=30)	6,7	16,7
MCP 4 (N=30)	6,7	3,3
MCP 5 (N=30)	10	20
<b>PIP (N=150)</b>	4	16,6
IP (N=30)	3,3	30
PIP 2 (N=30)	6,7	16,7
PIP 3 (N=30)	6,7	20
PIP 4 (N=30)	3,3	10
PIP 5 (N=30)	0	6,7
<b>DIP (N=120)</b>	0,8	10
DIP 2 (N=30)	0	13,3
DIP 3 (N=30)	3,3	13,3
DIP 4 (N=30)	0	6,7
DIP 5 (N=30)	0	6,7

**Conclusions:** This study showed a predilection for the gout specific ultrasound signs (DC and tophaceous deposits) of the wrist and MCP joints. The contribution of musculoskeletal ultrasound seems to be very interesting to objectify the presence of gout specific signs in the hand and fingers joints.

**Disclosure of Interest:** None declared

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#### AB0885 PREVALENCE OF ULTRASONOGRAPHIC GOUT SPECIFIC SIGNS OF FOOT JOINTS

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**Objectives:** To evaluate the prevalence of ultrasonographic gout specific signs of foot joints.

**Methods:** This is a cross-sectional study which includes 15 patients with chronic gout, defined according to the American College of Rheumatology criteria (ACR 1977). Ultrasound (US) examination was performed using a high-frequency linear probe (Toshiba Xario<sup>®</sup>, frequency (8–14 MHz)) in B mode. 330 articular sites were

studied at their dorsal surface. The ultrasound has objectified the presence of two signs: hyperechoic band over the superficial margin of the articular cartilage described as a double contour (DC) and tophaceous deposits at the joint cavity. **Results:** The mean age at onset was 54.7±12.6 years, and the median diagnosis duration was 0 (0.3) years.

The results of the US examination are summarized in Table 1.

Table 1. Prevalence of ultrasonographic gout specific signs of hind-foot, mid-foot and Metatarsophalangeal joints in the studied population

Joints (N=330)	Double contour (%)	Tophaceous deposits (%)
Hind-foot	8,8	10
Talocrural (N=30)	13,3	13,3
Subtalar:		
Lateral (N=30)	3,3	10
Medial (N=30)	10	6,7
Mid-foot	14,4	32,2
Talonavicular (N=30)	13,3	13,3
Intertarsal (N=30)	6,7	43,3
Tarsometatarsal (N=30)	23,3	40
MTP (N=150)	11,3	18
MTP 1 (N=30)	33,3	56,7
MTP 2 (N=30)	13,3	13,3
MTP 3 (N=30)	6,7	3,3
MTP 4 (N=30)	0	6,7
MTP 5 (N=30)	3,3	10

**Conclusions:** This study showed a predilection for the gout specific ultrasound signs (DC and tophaceous deposits) in the tarsometatarsal and metatarsophalangeal joints, especially in the first MTP. The contribution of musculoskeletal ultrasound seems to be very interesting to objectify the presence of gout specific signs of the foot joints.

**Disclosure of Interest:** None declared

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**AB0886 PHARMACOKINETICS, PHARMACODYNAMICS, AND TOLERABILITY OF VERINURAD (RDEA3170), A SELECTIVE URIC ACID REABSORPTION INHIBITOR, IN HEALTHY ADULT MALE SUBJECTS**

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**Background:** Verinurad (RDEA3170) is a selective uric acid reabsorption inhibitor in clinical development for the treatment of gout and asymptomatic hyperuricemia.

**Objectives:** The aim of this study was to evaluate the pharmacokinetics, pharmacodynamics, and tolerability of verinurad following single and multiple doses in healthy adult males.

**Methods:** This was a Phase 1, randomized, double-blind, placebo-controlled, single- and multiple-ascending dose study. Panels of 8 male subjects (6 active, 2 placebo) received a single oral dose of verinurad or placebo in either a fasted (2 mg, 5 mg, 20 mg, 40 mg) or fed (5 mg, 20 mg) state and panels of 12 male subjects (9 active, 3 placebo) received ascending doses of once-daily verinurad (1 mg, 5 mg, and 10 mg) or placebo in a fasted state for up to 10 days. Verinurad was administered as an oral solution for 1 and 2 mg doses and in tablet form for doses >2 mg. Serial plasma/serum and urine samples were assayed for verinurad and uric acid at predetermined time points. Safety was assessed by adverse event (AE) reports, laboratory tests, vital signs, and electrocardiograms (ECGs).

**Results:** A total of 81 adult males aged 18–54 years enrolled and completed the study. Following single oral doses of verinurad, absorption was rapid and exposure (maximum plasma concentration [C<sub>max</sub>] and area under the plasma concentration-time curve [AUC]) increased in a dose-proportional manner up to the maximum dose tested; C<sub>max</sub> was achieved at 0.5–0.75 hours post-dose in the fasted state, and was slightly delayed to 1.25 hours post-dose in the fed state. Food appeared to decrease AUC by about 23% and C<sub>max</sub> by about 50%. Following multiple daily doses, there was modest accumulation of verinurad. Urinary excretion of verinurad accounted for approximately 2% of the administered dose, suggesting that renal excretion is a minor elimination pathway for unchanged verinurad. Reductions in serum uric acid (sUA) correlated with dose. Under fasted conditions, single-dose administration of verinurad 2, 5, 20, or 40 mg reduced sUA levels by 16%, 24%, 48%, and 62%, respectively. Following multiple once-daily dosing for 10 days, verinurad reduced sUA levels by 22%, 44%, and 61% for the 1, 5, and 10 mg doses, respectively. A persistent pharmacologic effect (>15% fractional excretion of uric acid relative to baseline) was evident for at least 24 hours after dosing for verinurad doses of 2 mg or above. Verinurad was well tolerated at all doses. No serious AEs, severe AEs, discontinuations due to AEs or clinically significant laboratory or ECG abnormalities were reported.

**Conclusions:** Single and multiple doses of verinurad were well tolerated, absorption was rapid and exposure was dose-proportional. Verinurad increased urinary uric acid elimination and resulted in sustained reductions in sUA. These data support further clinical evaluation of once-daily verinurad as a treatment for gout.

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**AB0887 PHARMACOKINETICS, PHARMACODYNAMICS, AND TOLERABILITY OF VERINURAD (RDEA3170), A SELECTIVE URIC ACID REABSORPTION INHIBITOR, IN HEALTHY JAPANESE MALE SUBJECTS**

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**Background:** Chronic gout is a significant clinical problem in Asia, including Japan, where many patients remain suboptimally treated with currently available therapies. Verinurad (RDEA3170) is a selective uric acid reabsorption inhibitor in clinical development for the treatment of gout and asymptomatic hyperuricemia. **Objectives:** The aim of this study was to evaluate the pharmacokinetics, pharmacodynamics, and tolerability of verinurad in healthy Japanese and non-Asian adult male subjects.

**Methods:** This was a Phase 1, randomized, single-blind, placebo-controlled study (NCT01872832). Panels of 8 Japanese male subjects were randomized in a 3:1 ratio to receive a modified-release formulation of oral verinurad (2.5 mg, 5 mg, 10 mg, 15 mg) or placebo administered as a single dose in a fasted state and as multiple once-daily doses in a fed state for 7 days. A panel of 8 non-Asian male subjects received single and multiple doses of oral verinurad (10 mg) or placebo. Serial plasma/serum and urine samples were assayed for verinurad and uric acid at predetermined time points. Safety was assessed by adverse event (AE) reports, laboratory tests, vital signs, and electrocardiograms (ECGs).

**Results:** Of 48 randomized subjects, 46 (Japanese: 39, non-Asian: 7) completed the study. Treatment groups were generally well balanced; however, mean body weight and body mass index were approximately 14% and 7% lower, respectively, in Japanese than non-Asian subjects. Following single- or multiple-oral doses of verinurad in Japanese subjects, exposure (maximum plasma concentration [C<sub>max</sub>] and area under the plasma concentration-time curve [AUC]) increased in a near dose-proportional manner under fasted or fed conditions. The time to C<sub>max</sub> (T<sub>max</sub>) was approximately 1.25–2.0 hours post-dose under fasted conditions. A moderate-fat meal delayed T<sub>max</sub> up to 5 hours post-dose and increased plasma verinurad exposures up to 109%. Following once-daily multiple doses, there was modest accumulation of verinurad. C<sub>max</sub> and AUC were 38% and 23% higher, respectively, in Japanese versus non-Asian subjects, largely due to the difference in body weight. Mean reductions in serum uric acid following once-daily multiple dosing of verinurad 10 mg were 62% and 58% at maximum reduction and 46% and 44% at 24 hours post-dose in Japanese and non-Asian subjects, respectively. Verinurad was well tolerated at all doses. One Japanese subject discontinued verinurad due to an AE of urticaria that resolved after 11 days. No serious AEs, Grade 3 or 4 AEs, or clinically significant laboratory or ECG abnormalities were noted.

**Conclusions:** Verinurad significantly lowered serum uric acid and was well tolerated in both healthy Japanese and non-Asian males, despite small differences in plasma pharmacokinetics. These data support further evaluation of once-daily verinurad as a treatment for hyperuricemia with or without gout in the Japanese population.

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**AB0888 TENOFOVIR INDUCED OSTEOMALACIA: A PROFILE BASED ON THREE PATIENTS WITH LOW PHOSPHORUS AND NORMAL LEVELS OF VITAMIN D AND PARATHYROID HORMONE**

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**Background:** Tenofovir can induce proximal renal tubular changes that result in varying expression of Fanconi syndrome<sup>1,2</sup>. Less commonly, osteomalacia related to hypophosphatemia<sup>2,3</sup> can occur and has been documented with bone biopsy<sup>3</sup>. The clinical details of 28 reported cases of tenofovir induced osteomalacia, some of whom had vitamin D deficiency and secondary hyperparathyroidism, was recently summarized<sup>3</sup>.

**Objectives:** To describe the clinical presentation and course of three HIV patients with tenofovir induced hypophosphatemic osteomalacia and compare to cases previously reported.

**Methods:** The clinical, laboratory, and radiologic features of three HIV patients referred for evaluation of pain and osteoporosis were reviewed.

**Results:** All three patients were male, had diffuse pain, suffered multiple clinical fractures, and were on combination long and short acting opioids at the time of presentation. Two were in wheelchairs and two had neuropathy.

All patients had hypogonadism and proteinuria and case 1 had glycosuria. All patients had normal 25-OH vitamin D, vitamin B12, PTH, serum protein electrophoresis, magnesium, CBC, calcium, CPK, sedimentation rate, TSH. Case 1 had fractures of the hip, sacrum, and humerus; case 2 hip and ribs; and case 3 ribs, pelvis and knee. The technetium bone scan showed a similar pattern of increased uptake in multiple ribs, calcaneus, metatarsal bones, knees, and