

Upper Paleolithic tuberculosis: A probable case illustrated by paleoimaging methods (Azilian site of Les Iboussières, France)

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1	<u>Title</u> : UPPER PALEOLITHIC TUBERCULOSIS: A PROBABLE CASE ILLUSTRATED BY
2	PALEOIMAGING METHODS (AZILIAN SITE OF LES IBOUSSIERES, FRANCE)
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21 Abstract:

The aim of this paper is to present the results of µCT-scan and 3D imaging analyses of two 22 skeletal lesions observed on human remains of one of the last European hunter-gatherers from 23 the late Paleolithic (Azilian period): a sacroiliac osteoarthritis and a femoral lesion suggestive 24 25 of a soft tissue abscess imprint. These two skeletal elements (fused left sacrum and coxal bone, and right femur) displayed osteometric criteria indicating that they belonged to the same 26 individual. These two associated lesions are consistent with a low-grade osteoarticular 27 infection, and suggest a diagnosis of pelvic tuberculosis with a cold abscess of the thigh. If 28 29 molecular confirmation is obtained, this case would be the first evidence of tuberculosis among 30 Upper Palaeolithic populations. Keywords: 31

- 32 Tuberculosis; Hunter-gatherers; Upper Paleolithic; Azilian; Paleoimaging; 3D methods
- 33

35 **Text:**

36 1. Introduction

Tuberculosis is presented as one of the oldest human infections. Paleopathology can find its
traces in human remains from ancient times [1].

39 Although the Neolithic period was for a long time considered as the date of its emergence in the human species and animal domestication its main cause [2], the phylogenetic study of the 40 41 group of bacteria responsible for human and animal tuberculosis (*M. tuberculosis* complex) showed a very different evolutionary history, demonstrating an ancestry of the human-42 adapted strain compared to the animal-adapted strains [3]. This result allowed authors to 43 estimate the emergence of the human-adapted TB strain in the early period of Paleolithic [4]. 44 However, paleopathological research, whether morphological or molecular, has so far failed 45 to yield any convincing evidence of pre-Neolithic human tuberculosis [5]. Its earliest evidence 46 47 is dated to the emergence of the Neolithic in the Fertile Crescent, before domestication [6]. A convincing case of tuberculosis infection in the Paleolithic has been presented, but it concerns 48 49 a North American Pleistocene bison [7]. To resolve this paradox, paleopathologists must focus 50 their attention on Paleolithic human remains in search of evidence of tuberculosis infection, 51 using morphological and/or molecular methods. Thanks to paleoimaging methods that 52 brought an added-value to retrospective diagnosis in paleopathology [8], we present here, 53 while awaiting molecular confirmation, a probable case of pelvic tuberculosis on one 54 individual from the last European hunter-gatherers population (final Paleolithic, Azilian 55 period, South-Eastern France).

56

57 2. Material and methods

58 2.1. Archaeological background

59 The site "Aven des Iboussières" is located near Malataverne, a small town of in South-Eastern France near the city of Montélimar. The plain of Malataverne is overhung by an Urgonian 60 limestone massif, carved by an extensive karstic network, which is more than 600 meters long, 61 including several caves. This network is very unstable, making its exploration very dangerous 62 63 for speleologists. In 1990, speleologists explored the upper hall and discovered some human 64 remains. The instability of the network only allowed one rescue excavation, carried out in 1994 by Bernard Gély. Short time after this excavation, the collapse of the cavity vault prohibited 65 all further research. The east-west stratigraphic section distinguishes 5 layers (the fourth level 66

67 being itself subdivided into three sublevels A, B and C). The human remains and the great 68 majority of the archaeological and faunal material that have been collected, were coming from this layer 4C. The ¹⁴C dates of animal and human bones from this layer provided similar results 69 dating this site at 10210±80 BP [9]. This site is exceptional due to the quantity and the diversity 70 71 of the objects of ornament and is considered as the richest in Western Europe [10]. This 72 material comes from layers 4C and 5 and is represented by ochre stones and engraved bone 73 pendants, perforated and decorated hedgehog hemi-mandibles, incised, engraved and ochre bones, pierced teeth of deer, and marine shells. 74

75 2.2. Faunal diversity

Faunal remains are abundant and diversified [11]. Fourteen species were identified: Bison, Auroch, Ibex, Chamois, Moose, Deer, Roe deer, Wild boar, Leopard, Lynx and smaller mammals. Some of these remains presented marks of cutting, other were used as ornaments such as anterior teeth of deer, perforated to be sewed on a garment, or tied like pearls for a necklace, and small mammals' mandibles decorated with striations.

81 2.3. Human remains

The anthropological material was constituted by 291 bone fragments and 15 isolated teeth, representing a minimum number of 9 individuals, 4 adults recognized as males and 5 children. The general anthropological features are matched with those of Upper Paleolithic huntergatherers including general robustness, dolichocrania, broad face, low and rectangular orbits, platycnemia and high stature [12,13] (**figure 1**).

Paleogenomics analyses were performed on 3 individuals and showed that they belonged to the U lineage of the N mitochondrial DNA haplogroup such as some of the more ancient European hunter-gatherers (Dolni Vestonice) and the majority of the Holocene huntersgatherers [14].

91 2.4. Paleopathological observations

Among the pathologies previously noted [15], two skeletal lesions were reexamined that both have an infectious pattern. The observation of two lesions of infectious origin on two different bones in a small osteo-archaeological series (only 4 adult individuals), leads us to follow a principle of parsimony and propose that these are two manifestations of the same infectious process affecting the same individual. To support this hypothesis, we checked whether the dimensions of the femoral head corresponded to the size of the acetabulum. These two bones

98 being opposed (left acetabulum and right femoral head), it was not possible to test the 99 anatomical congruence of the coxofemoral re-articulation directly, so we used an indirect method by applying the formulae established by Plavcan et al [16]. These regression formulas 100 101 can be used to predict the diameter of an individual's femoral head from the size of his 102 acetabulum. Since there is no difference in size between the left and right femoral heads of the same person [17], we can bypass the question of laterality here. By obtaining, from 103 104 acetabular size (diameter: 53.5mm), a calculated value of 47.175 mm for the femoral head diameter, for a measured value of 47,4 mm, we can conclude that these two bones most 105 106 probably belong to the same individual. We have therefore re-examined these two pathological bones from the same pathophysiological perspective, which enabled us to enrich 107 108 the diagnostic discussion.

The first lesion is a left sacro-iliac fusion which had been the subject of a first study by medical tomography made in 1995, with, therefore, a resolution of low precision but which nevertheless allowed to perform a first 3D reconstruction [18] and the second lesion, which had not been analyzed by paleoimaging methods until now, which is present on the posterior side of the right femur: it shows an irregular aspect, with periosteal reaction and polylobulated imprints.

115 2.5. Paleoimaging methods

116 Microtomography acquisitions were performed using a GE v|tome|x S microtomograph. Coxal 117 bone was scanned at a voltage of 120 kV and a current of 200 μ A using a copper filter of 118 0.1mm. Images were acquired at a pixel size of 112 μ m.

Femoral diaphysis was scanned at a voltage of 100 kV and a current of 200 μA using a copper
filter of 0.1mm. Images were acquired at a pixel size of 105 μm.

121 The images were reconstructed with the TIVMI[®] v2.3 software based on the HMH (Half-122 Maximum Height) algorithm [19] extended to 3D [20]. The segmentation of the inner 123 structures was processed manually by the first author (HC).

124

125 3. Results

126 As for left sacro-iliac fusion, the 3D reconstruction from μ CT slices revealed 2 distinct cysts,

127 one anterior and one posterior, overlapping the sacroiliac joint, having a mirror aspect, and

located on the iliac and sacral sides of the joint. The latter is partially fused (figure 2).

The posterior cyst is in direct communication with remnant of sacroiliac joint, constituting an
external pathway to lumbo-gluteal soft tissues (figure 3).

As for femur, it presents at middle shaft periosteal reactions and lobulated imprints of the cortical surface (**figure 4**). The examination of the μ CT slices shows a typical aspect of cortical osteitis under the lobulated imprints with adjacent periosteal reaction. It can be observed that the medullar space and endosteum are strictly normal and that the periosteal reaction is not circumferential but limited. This aspect of osteoperiostitis is very different from the one of septic osteomyelitis which is characterized by invasion of the medullary zone and a circumferential periosteal reaction.

138

139 4. Discussion

Diagnostic discussion focuses first on sacroiliac lesions. These are characterized by fusion of 140 141 the sacroiliac joint and the presence of two intraosseous cystic cavities. Unilateral ankylosis of the sacroiliac joint can have three causes: degenerative, inflammatory or infectious. 142 Degenerative joint disease and DISH can lead to sacroiliac fusion, but are accompanied by 143 144 osteophytes with destruction of the joint in the first case, and thick ossification of the 145 sacroiliac ligaments without destruction of the joint space in the second; these two conditions 146 can be associated in elderly patients. Inflammatory spondyloarthropathies are very often 147 responsible for damage to the sacroiliac joint. The joint is affected, and fusion takes place 148 through fine, regular ossification of the sacroiliac ligaments. Age-related and inflammatory 149 causes can therefore be ruled out, and only infectious causes remain to be discussed. 150 Infectious sacroiliitis results either from pyogenic germs (Staphylococcus aureus, 151 Streptococcus, gram-negative bacilli), or from specific germs such as those of tuberculosis or brucellosis. Pyogenic germs have a cellular toxicity that makes them highly destructive to bone 152 153 tissue, with a strong inflammatory reaction that increase bone lysis; chronicity is rare, and is accompanied by bone sequestration and pronounced periosteal reactions; death was 154 frequent in the past in absence of treatment [21]. The aspect of the sacroiliac fusion is 155 therefore more suggestive here of a low-grade infection. Tuberculosis and brucellosis 156 157 infection have a granulomatous mode, the bone tissue reaction aimed at limiting the 158 progression of infection around the granulomas by peripheral sclerosis. Unlike brucellosis, 159 ankylosis is the natural history of sacroiliac tuberculosis [22]

160 As for cystic lesions, the discussion is more theoretical, as the main diseases responsible for 161 intraosseous cyst formation (aneurysmal bone cyst, unicameral bone cyst, cystic angiomatosis) or bone tumours or dysplasias having a cystic appearance (non-ossifying 162 fibroma, fibrous dysplasia, giant cell tumor, chondroblastoma, chondromyxoid fibroma, or 163 even osteoid osteoma) show different localizations, aspect and are never responsible for joint 164 lesions and ankylosis. The association of intraosseous cystic lesions of the ilium and sacrum in 165 166 close contact with partially fused sacroiliac arthritis has no other explanation than an infection. Brodie's abscess, corresponding to a subacute or chronic osteomyelitis with a "cyst 167 168 shape" may be discussed, as a very few cases were reported on posterior ilium in the literature [23], some of them with gluteal syndrome, but sacroiliac joints were normal in these reported 169 cases. Brodie's abscess of ilium is presented as very unusual, on the contrary of cystic bone 170 lesions due to tuberculosis which are much more frequent, whatever the skeletal location: 171 172 spine [24]; distal epiphysis [25]; iliac bone near sacroiliac joint [26]. Moreover, tuberculosis is known to produce mirror-image bone lesions on either side of an intervertebral disc, 173 174 metaphyseal plate or joint space [27], an aspect which is observed on our specimen, on the 175 two sides of the sacroiliac joint. This feature is not observed for Brodie's abscess. Therefore, 176 our conclusion is that the lesions we observed on this Azilian specimen (partial ankylosis of 177 sacroiliac joint with two osseous cysts overlapping the joint with a mirror aspect) is more in 178 favor of a tuberculous sacroiliitis than all the other discussed conditions.

179 Another element that adds to the discussion is the lesions observed on the posterior part of 180 the right femoral shaft. The first question is whether it could be septic osteomyelitis, 181 responsible for these irregular periosteal reactions. In this case, the infection starts in the 182 medullary spaces, crosses the cortical bone via the canaliculi and forms an abscess that detaches the periosteum, stimulating osteogenesis and leading to the formation of an 183 184 involucrum. The analysis of µCT slices excluded a possible diagnosis of septic osteomyelitis as they showed an aspect of moderate osteoperiostitis which is very different (see results and 185 186 figure 4).

The interpretation is that the periosteal and cortical bone changes observed on this femoral shaft are due to a slow-growing polylobulated mass located in the soft tissues of the posterior thigh and in close contact with the bone. The moderate periosteal and cortical changes observed on slices A to D (figure 4) are compatible with infectious abscess of low-grade, so called "cold abscess".

192 To summarize, anthropological and paleopathological criteria converge to suggest that these 193 two bone elements belong to the same subject, and to advance a diagnosis of low-grade 194 infection, with sacroiliac osteoarthritis and cold abscess of the thigh. If this semiology was observed on a today patient, the most probable diagnosis would be tuberculosis. The 195 196 sacroiliac joint is indeed a classic location for TB osteoarthritis, and cold abscess of the thigh has been described in medical literature as a consequence of pelvic skeletal tuberculosis with 197 198 gluteal involvement [28–31]. We can assume that the cold abscess of the thigh results here from a posterior migration of pus from the infected left sacroiliac joint under the gluteal 199 200 muscles, responsible for a flow of cold abscess from the sacral region to the posterior aspect of the right thigh through the various intermuscular spaces of the gluteal muscles and the 201 202 lodges of thigh muscles, anterior and posterior. This led to the development of a cold abscess 203 in soft tissues of the middle part of the thigh. The abscess was in close contact with the 204 posterior part of the femur, developed in the space between the *fasciae* separating the posterior muscle compartment from the anterior and medial compartments of the thigh 205 206 muscles (figure 5). The fact that the abscess was located on the right side of the thigh, whereas 207 the starting point was located on the left sacroiliac side of the thigh, was due to pain and 208 gravitational logic: the subject had to present a pain on the left side bothering him in a dorsal 209 or left lateral decubitus sleeping position. He had therefore to adopt an antalgic resting and 210 sleeping position on the right side, leading by gravity the passage of the tubercular pus from 211 the left side of the sacrum towards the spaces between the muscle lodges of the anterior and 212 posterior part of the right thigh.

To broaden the etiological discussion beyond the exclusive responsibility of M. tuberculosis, 213 214 the paleopathological lesions observed on these Paleolithic human remains are indicative of skeletal infection by a low-grade pathogen, more likely a bacterial one. Best profile would be 215 216 a telluric bacillus, having a wild animal reservoir and granulomatous inflammatory mode 217 responsible for chronic lesions involving bones and joints. Ideally, it should have a triple way of transmission (pulmonary, digestive and cutaneous). The bacteria having this profile belong 218 219 to the Actinobacteria phylum such as the genera Mycobacterium at the first place, followed 220 by Nocardia and Rhodococcus.

The archaeological context of the hunters-gatherers of the Iboussières site showed that they were in close contact with a large spectrum of wild fauna, which are currently a natural reservoir of these Actinobacteria, giving opportunity for microbial exchanges.

The aspect of the lesions is suggestive of a chronic stage of the infection, which is compatible with a human-adapted infection, as are some of the strains of *Mycobacterium tuberculosis* complex. This observation is in line with several phylogenetic models, according to which human populations in Eurasia were infected with tubercle bacilli as early as the beginning of the Upper Paleolithic [32]. The precise identification of the strain by paleomicrobiological methods is in progress. These future results will be of great interest to understand more precisely the evolution of human tuberculosis in Europe at the end of the Paleolithic.

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233 References:

- 234 [1] Pálfi G, Dutour O, Perrin P, Sola C, Zink A. Tuberculosis in evolution. Tuberculosis
 235 2015;95:S1–3. https://doi.org/10.1016/j.tube.2015.04.007.
- 236 [2] Pales L. Paléopathologie et pathologie comparative. Paris: Masson et Cie; 1930.
- 237 [3] Brosch R, Gordon SV, Marmiesse M, Brodin P, Buchrieser C, Eiglmeier K, et al. A new
- evolutionary scenario for the Mycobacterium tuberculosis complex. PNAS 2002;99:3684–
 9. https://doi.org/10.1073/pnas.052548299.
- [4] Gutierrez MC, Brisse S, Brosch R, Fabre M, Omaïs B, Marmiesse M, et al. Ancient Origin
 and Gene Mosaicism of the Progenitor of Mycobacterium tuberculosis. PLoS Pathog
 2005;1:e5. https://doi.org/10.1371/journal.ppat.0010005.
- [5] Buzic I, Giuffra V. The paleopathological evidence on the origins of human tuberculosis: a
 review. Journal of Preventive Medicine and Hygiene 2020;Vol. 61 No. 1s1:E3 Pages.
- 245 https://doi.org/10.15167/2421-4248/JPMH2020.61.1S1.1379.
- [6] Baker O, Lee OY-C, Wu HHT, Besra GS, Minnikin DE, Llewellyn G, et al. Human
 tuberculosis predates domestication in ancient Syria. Tuberculosis 2015;95:S4–12.
 https://doi.org/10.1016/j.tube.2015.02.001.
- [7] Rothschild BM, Martin LD, Lev G, Bercovier H, Bar Gal GK, Greenblatt C, et al.
 Mycobacterium tuberculosis Complex DNA from an Extinct Bison Dated 17,000 Years
 before the Present. CLIN INFECT DIS 2001;33:305–11. https://doi.org/10.1086/321886.
- [8] Coqueugniot H, Dutailly B, Dutour O. The third dimension in palaeopathology: How can
- three dimensional imaging by computed tomography bring an added value to
- retrospective diagnosis? Int J Osteoarchaeol 2020;30:538–50.
- 255 https://doi.org/10.1002/oa.2881.
- [9] Gély B, Morand P. Les sépultures épipaléolithiques de l'aven des Iboussières à
 Malataverne (Drôme, France): Premiers résultats. Ardèche Archéologie 1998:13–8.
- 258 [10] D'Errico F, Vanhaeren M. Le mobilier funéraire de l'Aven des Iboussières et l'identification
- des marqueurs culturels à l'Epipaléolithique. Les derniers chasseurs-cueilleurs d'Europe
 occidentale 13000-5500 av. J.-C., vol. 699, Annales Littéraires de l'Université de Besançon;
 2000, p. 325–42.
- [11] Chaix L. Unusual and eclectic: the epipalaeolithic fauna from the Aven des Iboussières
 (Drôme, France). Munibe (Antropologia-Arkeologia) 2005:411–20.
- 264 [12] Dutour O. Etude anthropologique préliminaire. In: Gély B, Morand P, editors. Les
- 265 sépultures épipaléolithiques de l'aven des Iboussières à Malataverne (Drôme, France):
 266 Premiers résultats., 1998, p. 13–8.
- [13] Aymard I, Ardagna Y, Lalys L, Signoli M, Gély B, Dutour O. Etude anthropologique du site
 azilien des Iboussières, Malataverne, Drôme. Un siècle de construction du discours

- scientifique en Préhistoire, vol. 3, Avignon: Société Préhistorique Française; 2007, p. 537–
 44.
- 271 [14] Posth C, Renaud G, Mittnik A, Drucker DG, Rougier H, Cupillard C, et al. Pleistocene
- 272 Mitochondrial Genomes Suggest a Single Major Dispersal of Non-Africans and a Late
- Glacial Population Turnover in Europe. Current Biology 2016;26:827–33.
- 274 https://doi.org/10.1016/j.cub.2016.01.037.
- [15] Dutour O, Palfi G, Panuel M, Gély B. Paleopathological study of Upper Paleolithic remains
 from Southeastern France. Journal of Paleopathology 1995:98.
- 277 [16] Plavcan JM, Hammond AS, Ward CV. Brief Communication: Calculating hominin and
- 278 nonhuman anthropoid femoral head diameter from acetabular size: Reconstructing Fossil
 279 Femoral Head Size. Am J Phys Anthropol 2014;155:469–75.
- 280 https://doi.org/10.1002/ajpa.22591.
- [17] Pfeufer D, Gililland J, Ajinkya R, Duensing I, Anderson MB, Peters CL, et al. Radiographic
 Native Femoral Head Size Correlates with Acetabular Cup Size in Primary Total Hip
- 283 Arthroplasty. J Hip Surg 2019;03:197–202. https://doi.org/10.1055/s-0039-3400481.
- [18] Coqueugniot H, Dutailly B, Desbarats P, Boulestin B, Pap I, Szikossy I, et al. Three dimensional imaging of past skeletal TB: From lesion to process. Tuberculosis
- 286 2015;95:S73–9. https://doi.org/10.1016/j.tube.2015.02.004.
- 287 [19] Spoor CF, Zonneveld FW, Macho GA. Linear measurements of cortical bone and dental
- enamel by computed tomography: Applications and problems. Am J Phys Anthropol
 1993;91:469–84. https://doi.org/10.1002/ajpa.1330910405.
- 290 [20] Dutailly B, Coqueugniot H, Desbarats P, Gueorguieva S, Synave R. 3D surface
- reconstruction using HMH algorithm. Proceedings of IEEE International Conference on
 Image Processing, Le Caire, Egypte, 7-10 Novembre 2009: 2009, p. 2505–8.
- [21] Dutour O. Paleopathology of Infectious Diseases. In: Grauer AL, editor. The Routledge
 Handbook of Paleopathology, Routledge; 2022, p. 324–37.
- 295 [22] Ramlakan RJS, Govender S. Sacroiliac joint tuberculosis. International Orthopaedics (SICO
- 296 2007;31:121–4. https://doi.org/10.1007/s00264-006-0132-9.
- 297 [23] Behera G, Poduval M, Patro D, Sahoo S. Brodie's Abbscess of Posterior Ilium with Gluteal
- 298 Syndrome, an Unusual Cause of Paediatric Low Back Pain: A Case Report. MOJ
- 299 2017;11:68–71. https://doi.org/10.5704/MOJ.1707.009.
- 300 [24] Zhen P, Li X, Lu H. Single Vertebra Tuberculosis Presenting with Solitary Localized
- 301 Osteolytic Lesion in Young Adult Lumbar Spines: Single Vertebra Tuberculosis.
- 302 Orthopaedic Surgery 2013;5:105–11. https://doi.org/10.1111/os.12037.
- 303 [25] Elmi A, Tabrizi A, Tolouei FM. Skeletal Tuberculosis Presenting as a Small Cystic Lesion in
- the Medial Femoral Condyle. . NUMBER 2013;1.

- 305 [26] Ismail M. Isolated iliac bone tuberculosis: a case report n.d.
- 306 [27] Akgul T. Cystic transphyseal bone tuberculosis: a report of two cases. Acta Orthop
- 307 Traumatol Turc 2012;46:316–9. https://doi.org/10.3944/AOTT.2012.2698.
- 308 [28] Bevan PG. Tuberculosis of Pubis Presenting as Cold Abscess of Thigh. BMJ 1955;2:832–3.
 309 https://doi.org/10.1136/bmj.2.4943.832.
- 310 [29] Garg PK, Chaurasia A, Singh AP. An Unusual Cause of Thigh Swelling: Tuberculosis of the
- Pubis. The American Journal of Tropical Medicine and Hygiene 2013;88:1–2.
- 312 https://doi.org/10.4269/ajtmh.2013.12-0358.
- 313 [30] Chahbouni M, Eloukili I, Lamrani MO, Kharmaz M, Ismail F, Mahfoud M, et al. Abcès
- tuberculeux de la face postérieure de la cuisse: à propos d'un cas. Pan Afr Med J 2014;19.
 https://doi.org/10.11604/pamj.2014.19.50.4683.
- 316 [31] Papagelopoulos PJ, Papadopoulos ECh, Mavrogenis AF, Themistocleous GS, Korres DS,
- Soucacos PN. Tuberculous sacroiliitis. A case report and review of the literature. Eur Spine
 J 2005;14:683–8. https://doi.org/10.1007/s00586-004-0831-0.
- 319 [32] Dutour O. The paleopathology and paleoepidemiology of Upper paleolithic tuberculosis:
- Review of evidence and hypotheses. Tuberculosis 2023;this special issue:xxx.
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- 329

- 330 Figure captions:
- Figure 1. Skull of male adult from the Iboussières site, facial view (A) and right lateral view (B).
- 332 Credits: O. Dutour
- 333 Figure 2. (A): 3D reconstruction of the sacrum and left ilium in superior view; (B): Transparency
- of the 3D reconstruction of the two bones with superimposition of axial μ CT section. It reveals
- two cysts: one anterior (red) and one posterior (green) overlapping the sacroiliac joint (dark
- blue) which is partially fused; (C): Same axial μ CT section with the two cysts only. Credits: H.
- 337 Coqueugniot
- 338 Figure 3. Axial μ CT section showing the communication of the posterior cyst with the gluteal
- 339 space: (1) sacrum, (2) left ilium, (3) anterior remnant of the sacroiliac joint, (4) posterior cyst;
- 340 (5) communication of the posterior cyst through posterior remnant of the sacroiliac joint
- towards the gluteal space (6). Credits: H. Coqueugniot
- Figure 4. Portion of the right femoral shaft showing the level of µCT slices from A to D. They reveal a typical aspect of cortical osteitis (1) under the lobulated imprints (2) with adjacent periosteal reaction (3). It can be observed that the medullar space and endosteum are strictly normal and that the periosteal reaction is not circumferential but limited to the part underlying the remodeled surface at the posterior face of femoral diaphysis. Credits: H. Coqueugniot
- Figure 5. Reconstructed section of the right thigh showing the development of the cold abscess (1) in the space between the anterior (A) and posterior-medial (B) muscular lodges in close contact with femur (2). Credits: O. Dutour
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- 353