

## Article

# Recognizing and Distinguishing the phenomenon referred to as meningioma

Bruce M. Rothschild \*

Carnegie Museum, 4400 Forbes Ave, Pittsburgh, PA 15213, USA

\* Correspondence: spondylair@gmail.com; Fax: 0000-0003-1327-6615

**Abstract: Background:** The reliability of a recent review of meningiomas in the archeologic record was difficult to assess, given the inverted sex ratio of the report and other contents apparently at variance with anatomical/medical findings in scientifically-identified cases. It therefore seemed appropriate to reexamine the nature of meningiomas and derive improved criteria for their recognition in the archeologic record and distinguish them from hemangiomas and bone marrow hyperplasia (recognized in the form of porotic hyperostosis). **Methods:** Medically-documented cases of meningiomas, were examined to establish a macroscopic standard that distinguishing among them. Alleged cases in the archeologic record were examined for conformity with those criteria. **Results:** An en face pattern of uniform mesh with contained whorls appears pathognomonic for meningiomas. This contrasts with the non-uniform marrow expansion displacement of trabeculae in porotic hyperostosis and non-uniform vascular displacement of trabeculae in hemangiomas. Reassessment of past attributions revealed few cases of meningiomas that could be confidently diagnosed. Those identified has sex ratios parsimonious with medical literature reports. **Conclusions:** Criteria suggested for identifying meningiomas permits distinguishing from hemangiomas, bone marrow hyperplasia (porotic hyperostosis) and from the macroscopically-observable surface spicules characteristic of osteosarcomas. Examination for fulfillment of criteria for meningiomas and hemangiomas seems to provide a picture (including sex ratios) mirroring that of the clinical literature, concluding that Cook and Danforth's disparate ratios were related to less fastidious case selection. Additionally, confidence in recognizing porotic hyperostosis may be compromised because of apparent similar macroscopic alterations to those seen with hemangiomas.

**Keywords:** meningioma; hemangioma; cranial pathology; porotic hyperostosis; differential diagnosis; osteosarcoma

## 1. Introduction

I have long sought clarification of post hoc examples and the literature related to meningiomas. A variety of cranial pathologies of disparate sizes and morphologies (e.g., 18,87) have been so-attributed. Given past experience with the trans-phylogenetic uniformity of findings for a given disease [63,64], it seems reasonable to conduct an intensive examination of what is being referred to as meningiomas. The purpose of this review of documented clinical cases and review the anthropologic literature to for diagnosis accuracy by:

1. Examining medically-documented cases of meningioma to develop evidence-based criteria for their recognition and for distinguishing them from the disorders in their differential
2. Reviewing the archeologic record to identify validly diagnosed occurrences, distinguishing them from hemangiomas and cranial bone marrow hyperplasia.

### 1.1. Derivation

Meninges, the membranes enclosing the brain, arise from multi-potential mesenchymal cells. Thus, they can therefore differentiate into a variety of cell types (e.g., fibrous, osseous, hematopoietic, vascular), explaining the variety of derived tissues referred to as meningiomas [4]. Meningiomas are generally slow-growing tumors derived from the membranous layers surrounding the brain and spinal cord [22]. Arising from arachnoid granulations, they cluster around venous sinuses and dural folds [44]. .

### 1.2. History

First reported by Felix Plater 1614 [58] and subsequently named by Harvey Cushing in 1922 [54], meningiomas have been variously been referred to as neoplastic (i.e., endotheliosis of the meninges, epithelioma, angioendothelioma, dural endothelioma, meningoblastoma, arachnoidal or meningeal fibroblastoma, mesothelioma of the meninges, dural sarcoma and fibrosarcoma), infectious (i.e., dura mater fungus or fungus tumors) and even as psammoma [2]. Most are actually benign, with only one to nine percent interpreted as malignant or atypical [33,46,61].

### 1.3. Prevalence

The prevalence of meningiomas is 7.7-8.4 per 100,000 adults [49,61,83]. Kostandy et al. [44] reported prevalence of 0.3/100,000 in childhood (males 2:1). This ratio in children contrasts with the 2-4-fold female predominance noted in adults [49,61,83], indistinguishable from that noted with hemangiomas [32,78,84,86]. Nakasu et al. [52] reported the presence of meningiomas in 2.3% of autopsies. They noted that 8.2% were multiple, but that none of their sample had neurofibromatosis. The latter is noteworthy as fifty percent of individuals with type 2 neurofibromatosis develop meningiomas [53]. Sosman [73] suggested the role of trauma in their development. Increased body fat also appears to be a risk factor for meningioma development [53].

### 1.4. Clinical symptoms

Only one in four individuals with meningiomas is symptomatic [61,83]. Clinical symptoms include headache, mental status (thought process) changes, ataxia (impaired balance), vertigo (dizziness), seizures, paresis (impaired muscle function), paresthesias (pins and needles feelings), aphasia (loss of speech), visual disturbance and hearing, dependent upon the brain region juxta-positional to the meningioma.

### 1.5. Character

Starr and Cha [74, p. 722] divided meningiomas into “expansile masses with wide dural attachment or ‘en plaque’ patterns of growth in a sheet-like pattern along the dura.” It is the macroscopically recognizable ectocranial appearance of the latter group that is the subject of the current analysis. Medically-confirmed cases are reported as having flat cranial accretions [42,57], which Kostandy et al. [44] referred to as the plaque variant. The thickened bone contains tumor cells [20,69].

### 1.6. Radiographic appearance

Pertinent radiologic findings include increased vascularity, calcification and bone destruction or hyperostosis in 25-50% [35,39,42,57]. They may present as predominantly osteolytic (e.g., Jónsdóttir et al., 2003) or as osteoblastic phenomena. Meningiomas have whorl-shaped inhomogeneous areas [6,76] and may have a nodular appearance [5]. A focal, diffuse, rim-like or punctate calcification may be noted [34] lobulated masses have also been reported [40,42,57]. Rohringer et al. [61] reported finding the mushroom (herein, referred to as nodular) form only in malignant cases. What seems to be absent from all medically-confirmed cases is a honeycomb appearance [16,51,68,84].

Additional diagnostic signs have been suggested, although have yet to be verified in individuals with independently-confirmed (e.g., medical) diagnoses. Campillo [14] suggested exaggerated middle meningeal artery imprinting as a possible sign of a meningioma, while Waldron [80] hypothesized that increased vascular channels might facilitate diagnosis. Sosman [73] suggested diagnostic significance of vascular channels radiating from the lesion.

#### *1.7. Controversy as to macroscopic appearance*

One of the terms (what might be referred to as ectocranial “protuberances”) often utilized in histologic descriptions of cranial alterations has caused confusion, as it is usually applied by radiologists to a very different phenomena. That term is spicules. Huggins [39] suggested that some meningiomas present with spikes parallel with the cranial surface, although neither Kim et al. [42] nor Phemister [56] identified any such findings. Arana et al. [6] referred to surface alterations as spicules, despite the smooth or multinodular ectocranial appearance, as Daffner et al. [24] noted. Rohringer et al. [61] suggested that the macroscopic appearance does not permit distinguishing malignant from benign meningiomas, although they reported finding the mushroom (what we refer to as nodular) form only in malignant cases. Rowbotham [65 p. 605] used the term spicules to describe “right-angled speculation had been laid down in layers parallel with the two tables of the skull.” The endocranial surface was spiculated or eroded, but the ectocranial surface was smooth on macroscopic examination.

Another challenge relates to use of the term “sponge-like” for the spaces (surface-visible holes) in a sponge or to its surrounding matrix. Sponge-like pattern seems to be a matter of perception of the pertinent attributes of a sponge: The holes or the matrix forms the search image we use for recognition of a structure as sponge-like. This has led to confusion related to distinguishing meningiomas and hemangiomas.

#### *1.8. Differential diagnosis*

The major differential diagnostic considerations related to macroscopic recognition of meningiomas include hemangiomas and bone marrow hyperplasia (e.g., thalassemia related/induced porotic hyperostosis). Also requiring consideration (Table 1) are fibrous dysplasia, Paget’s disease, osteoma, xanthomatous disease, histiocytosis, Proteus syndrome (characterized by asymmetrical body part hamartomatous overgrowth), haemangiopericytomas, lymphomas, schwannomas, fibrous tumors, chondrosarcomas, metastases, plasmacytomas and chondromas, astrocytomas, gliosarcomas, haemangioblastomas, giant cell tumors, osteomyelitis and fungal lesions [31,73,85]. In their series of 185 CT (computerized tomographic) studies, Arana and Martí-Bonmati [5] reported 18.9% histiocytosis, 15.1% osteoma, 12.9% epidermoid and dermoid cysts, 12.4% metastasis, 10.8% meningioma, 9.1% hemangioma, 6.4% fibrous dysplasia, with 14.4% miscellaneous diseases.

**Table 1.** Differential diagnosis of meningiomas [5,73,76,77,84]

Diagnosis/Finding	Pressure erosion	Interstices*	Fenestrated	Sclerosis	Honeycomb	Prominent vessel
Meningioma	Present	Parallel	Absent	Present	Absent	Present
Hemangioma	Present	Variable	Present	Absent	Present	Present
Marrow hyperplasia	Present	Variable	Present	Absent	Present	Absent
Fibrous dysplasia	Absent	Absent	Absent	Variable	Absent	Absent
Paget's disease	Absent	Absent	Absent	Cotton wool	Absent	Absent
Hamartoma	Absent	Absent	Absent	Present	Absent	Absent
Osteoma	Absent	Absent	Absent	Present	Absent	Absent
Epidermoid/dermoid cyst	Present	Absent	Absent	Absent	Absent	Absent
Xanthomatous	Present	Absent	Absent	Absent	Absent	Absent
Histiocytosis	Absent	Absent	Absent	Absent	Absent	Absent
Haemangiopericytoma	Absent	Absent	Absent	Absent	Absent	Absent
Hemangioblastoma	Absent	Absent	Absent	Absent	Absent	Absent
Giant cell tumor	Absent	Septated	Absent	Present	Multi-locular	Absent
Lymphoma	Absent	Absent	Absent	Absent	Absent	Absent
Schwannoma	Absent	Absent	Absent	Absent	Absent	Absent
Fibrous tumor	Absent	Absent	Absent	Absent	Absent	Absent
Chondrosarcoma	Absent	Calcifications	Absent	Calcifications	Absent	Absent
Metastasis	Absent	Absent	Absent	Variable	Absent	Absent
Multiple myeloma	Present	Absent	Absent	Absent	Absent	Absent
Chondroma	Present	Absent	Absent	Absent	Absent	Absent
Astrocytoma	Absent	Absent	Absent	Absent	Absent	Absent
Gliosarcoma	Present	Absent	Absent	Absent	Absent	Absent
Osteomyelitis	Present	Variable	Variable	Present	Absent	Absent
Fungal	Present	Absent	Absent	Absent	Absent	Absent
Actinomycosis	Present	Absent	Absent	Absent	grape-like	Absent

\*Striations

Hemangiomas occasionally presentation as “a hard, blue-domed lump on the skull, lying beneath the pericranium, which can be lifted off it.” They appear as a round or oval area of rarefaction with irregular borders, but never serpiginous [85]. What seems to be absent from all medically-confirmed cases of meningiomas is a honeycomb appearance, a phenomenon reported with hemangiomas [16,51,68,85]. The latter are characterized by an irregularly fenestrated meshwork with greatly variable interstices between bony trabeculae [85], which may be responsible for the honeycomb appearance.

Bone marrow hyperplasia (such as that related to thalassemia) alters the appearance of the outer cortex. It also presents as an irregularly fenestrated meshwork with greatly variable interstices between bony trabeculae [37,85]. The resultant radiating bone spicules give a granular osteoporosis, widening of the diploic space, and thinning with perforation of outer table of skull and subperiosteal proliferation producing a “hair on end” appearance on x-ray. The latter is observed in 8-12% of individuals with thalassemia [66,84]. Tyson and Alcauskas [79] reviewed Hrdlička's paleopathology collection at the San Diego Museum of man. Peruvian skulls 1915-2-145, 147, 151 and 154, diagnosed as having porotic hyperostosis, also have a sponge-like appearance, while 1915-2-158 had a sponge-like applicée composed of parallel spicules. They are characterized by an irregularly fenestrated meshwork with greatly variable interstices

between bony trabeculae (Wyke, 1949), which may be responsible for the honeycomb appearance

Arana et al. [6] and Kim et al. [42] noted that the irregular endocranial surface facilitated distinguishing meningiomas from fibrous dysplasia and from osteomas with their smooth inner surfaces and button appearance. The diffuse cranial thickening and cotton wool radiologic appearance of Paget's disease (see chapter 11) is distinguishable from meningiomatous ectocranial alterations [30]. Neither Schüller-Christian disease (xanthomata) have striations or honeycomb findings. Meningiomas lack the serpiginous shape characteristic of histiocytosis, epidermoid cysts and osteomyelitis [5,84]. Epidermoid and dermoid cysts cause pressure erosions with expansion of surrounding bone, but not new bone formation, although saponification does produce internal calcification in dermoid cysts [5].

One further differential consideration: The contemporary case reported by Huggins et al. [39] was a post-traumatic lump followed by swelling. Moth-eaten osteolysis was associated with remodeling. The possibility of a primary lesion aggravated by a hematoma seems likely.

### 1.9. Recognition, phylogenetic distribution and antiquity

The first clinical case of meningioma is apparently that of Platter in 1614, according to Bir et al. [11]. The oldest archaeological reports of a meningioma is that of Czarnetzki et al. (2003) in a 366,300 year old *Homo steinheimensis*.

Meningiomas are not limited to humans, but have also been reported in cats and dogs, especially, dolichocephalic dog breeds (e.g., collies, shepherds) of the latter [47].

The current study was pursued to identify macroscopically observable differences among the surface appearances of meningiomas, hemangiomas and bone marrow hyperplasia, as manifest by porotic hyperostosis

## 2. Materials and Methods

Medically (clinically)-documented cases of meningiomas, hemangiomas and cranial marrow hyperplasia in adults were examined to establish a macroscopic standard for confident identification of meningiomas and for distinguishing them from hemangiomas and marrow hyperplasia, manifest as porotic hyperostosis. Images and descriptions of archeologic site alleged meningioma cases (derived from Pub Med and Google searches and from 40 years of personal files) (Table 2), including those suggested by Cook and Danforth [18], were then examined for conformity with those criteria.

**Table 2.** Reassessment of archaeological site cranial pathology attributed to meningiomas.

Dating (Century)	Sex	Location	Atypical Features	Reassessment	Reference
3,653 <sup>rd</sup> BCE	?F	Germany	<i>Homo steinheimensis</i>	Likely	23
2,000 <sup>th</sup> BCE	?	France	Osteolytic with thick groove	Hemangioma	27,28,29
34 <sup>th</sup> BCE	F	Egypt	Honeycomb ectocranial, smooth endocranial expansion	Hemangioma	60
32 <sup>th</sup> BCE	M	Germany	Vascular impression as basis	Uncertain	81
45 <sup>th</sup> -10 <sup>th</sup> BCE	M	Denmark	Centripedal columns	Not meningioma	9
	M	France	Amorphous	Challenged	18, citing 29

			surface		
	M	Catalonia	Osteolytic with thick groove	Osteolytic	14
	F	Catalonia	Endocranial enostoma	Uncertain	14
	?	Catalonia	Endocranial hypervascularization	Uncertain	14
33 <sup>rd</sup> -21 <sup>st</sup> BCE	F	Austria	Hemangiomatous pattern	Hemangioma	70
12 <sup>th</sup> -11 <sup>th</sup> BCE	F	Egypt	Called honeycomb, but actually perpendicular, but uniform spicules	Sarcoma	12,60
8 <sup>th</sup> -5 <sup>th</sup> BCE	M	China	sponge-like	Hemangioma	56
8 <sup>th</sup> BCE-1 <sup>st</sup> CE	F	England	Focal "bump"	Possible	75
	M	Peru	Lysis with irregular trabeculae	Cancer	10
1 <sup>st</sup> -4 <sup>th</sup> CE	F	England	Low resolution image	Possible	12
3 <sup>rd</sup> -9 <sup>th</sup> CE	M	Scotland	honeycomb	Hemangioma	13
	?	Spain	Endocranial "enostoma"	Uncertain	14
	?	Spain	Endocranial enostosis	Uncertain	14
8 <sup>th</sup> -11 <sup>th</sup> CE	?	England		Likely	80
10 <sup>th</sup> -18 <sup>th</sup> CE	M	Alaska		Possible	41
11 <sup>th</sup> - 13 <sup>th</sup> CE	M	Germany	classic honeycomb	Hemangioma	82
12 <sup>th</sup> -13 <sup>th</sup>	M	Poland	No description	Uncertain	36
12 <sup>th</sup> -14 <sup>th</sup>	F	Illinois	Indefinite; no images or description	Uncertain	17
13 <sup>th</sup> -14 <sup>th</sup>	F	England	Sponge-like surface	Hemangioma	3
13 <sup>th</sup> -16 <sup>th</sup> CE	M	Czech Republic	No ectocranial alterations	Uncertain	72
14 <sup>th</sup> CE	F	Sweden	Focal "bump"	Possible	45
Pre 15 <sup>th</sup> CE	M	California	Honeycomb, "hair-on-end"	Sarcoma	1
	F	Peru		Likely	50
	M	Peru	Focal bump	Possible	50
Post 15 <sup>th</sup> CE		Mexico	Irregular intraocular mass	Unlikely	15
16 <sup>th</sup> -17 <sup>th</sup> CE	?F	Belize	Honeycomb	Hemangioma	18,25
18 <sup>th</sup> CE	M	Germany	Inadequate drawing quality	Uncertain	43
19 <sup>th</sup> -20 <sup>th</sup> CE	F	New York	No such presentation listed	Non-existent	71

Unspecified	M	England	Aberrant endocranial vessel	Uncertain	80
	F	England	Holes, endocranial vasculature	Uncertain	80
	M	England		Likely	80
	F	Sri Lanka	Multinodular	Likely	55
	F	Peru	Sponge-like applicée	Hemangioma	79
	M	Peru	Towering spicules	Osteosarcoma	48,55

\*Originally reported as thalassemia by Thomas (2016).

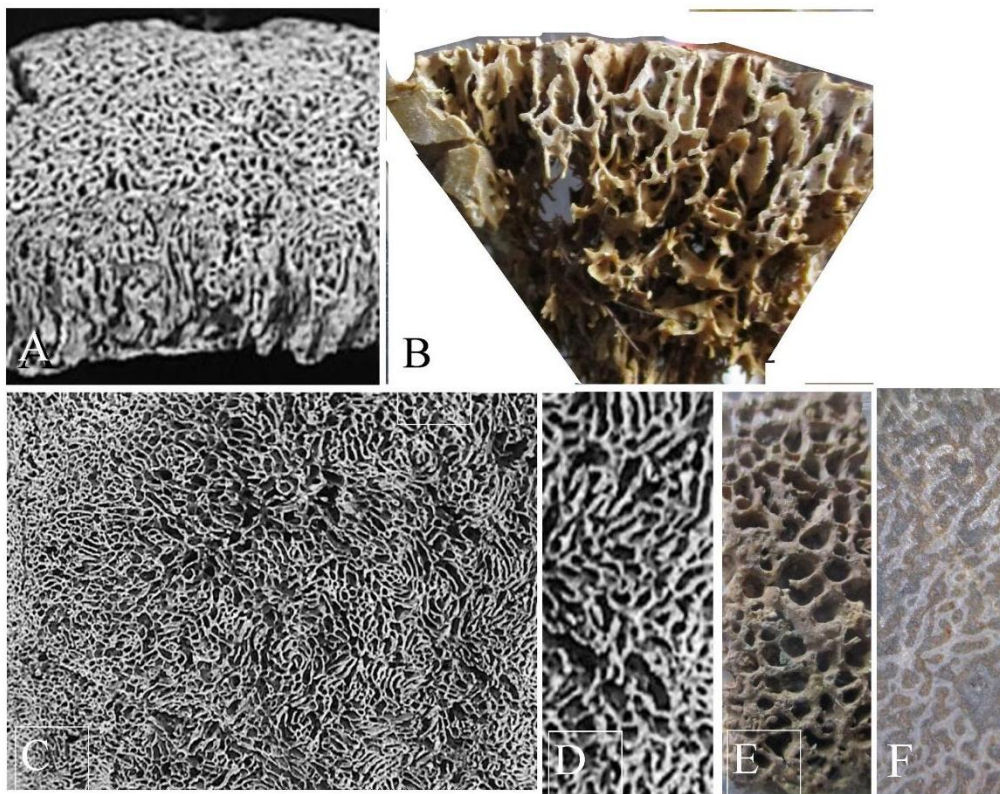
### 3. Results

#### **Establishment of criteria for macroscopically recognizing meningiomas and distinguishing them from hemangiomas and porotic hyperostosis**

A honeycomb appearance of the surface of the pathology was found only in hemangiomas and those cases of porotic hyperostosis in which the marrow and its trabeculae are actually exposed, in contrast to those in which only surface “pores” are recognizable. This is parsimonious with their previous characterization as an irregularly fenestrated meshwork with greatly variable interstices between bony trabeculae (Wyke, 1949)

The most pathognomonic surface sign of a meningioma appears to be a relatively uniform mesh (Figure 1D) with whorl patterns (Figure 1C). Figure 1A illustrates the plates of new bone that form the basis of this pattern. That contrasts with the trabeculae displaced by expansion of marrow spaces in porotic hyperostosis (Figure 1B). The latter gives rise to the non-uniform fenestrations seen in Figure 1E. This appears indistinguishable from the effect on trabeculae of hemangioma-related expansion of vasculature (Figure 1F). The trabeculae in meningiomas are parallel (Figure 1A), contrasted with the variably oriented components (Figure 1B).





**Figure 1.** Cranial pathology. (A) Oblique view of meningioma fragment. Mesh-like surface created by parallel plates; (B) Cross-section of porotic hyperostosis. Irregular trabeculae surrounding marrow spaces; (C) En face view of meningioma fragment shown in A. Whorled pattern of meshwork; (D) Enlarged view of C. Uniform meshwork; (E) En face view of porotic hyperostosis. Irregular meshwork; (F) En face view of hemangioma. Irregular meshwork.

#### Analysis of the accuracy of past archeologic cases attribution as meningiomas

Reassessment of past attributions revealed few cases that fulfilled criteria (Table 2) for meningiomas, identified cases actually representing hemangiomas and identified inadequate support for confident diagnosis in many others. Further, Cook and Danforth's ([18] citation of Siriani et al. [71] as reporting 19<sup>th</sup>-20<sup>th</sup> century meningioma in an individual from the Erie County Poor House apparently represents their unassessed utilization of a secondary or tertiary citation, as there is no record of any such presentation at the 2014 Paleopathology Association meeting. Excluding uncertain cases, those with alternative diagnoses and unconfirmed citations, the ratios women to men is 5:4 for meningiomas.

Danforth et al. [25] state that the Belize skull fragment resembles that reported by Schamall et al. [70], which actually appears to be a hemangioma, not a meningioma. Their diagnostic approach appeared somewhat convoluted and perhaps circular? They stated (p. 1046) that "virtually, no source discusses a case with as much vault thickness present in Burial 157 as being nutritional in origin." (That statement appears to be their attempt to rule out iron deficiency or other nutritional deficiencies and thus porotic hyperostosis). They continued "All scholars who have seen Burial 157 note that it far exceeds that of any proposed case of porotic hyperostosis that they had observed." That contrasts with figure 1, which clearly documents the extent of porotic hyperostosis-derived diploic space expansion. Cook and Danforth [18 p. 1047] also ruled out the latter because "it would be highly unlikely that any sort of genetic resistance would emerge in only a few generations," according to their limited differential considerations, failing to consider marrow hyperplasia induced by parasite-related blood loss. They



ruled out hemangioma because “the inner table” (of their fragment) “does not show the porosity characteristic of hemangiomas,” although porosity has never been documented as a “characteristic of hemangioma.” Absence of a honeycomb appearance does make the meningioma diagnosis more likely, but the possibility of porotic hyperostosis (e.g., from parasite-induced blood loss was not excluded.

Abbott and Courville [1] examined San Nicolas Island and Inuit skulls in the San Diego Museum collection, noting twenty with neoplastic lesions. Osteomas account for 11, cancer for seven and large hyperostoses were present in two. Number 158 was interpreted [18, p. 103] as “formed by the fused radiating spicules and the intervening openings having been compared by Moodie [50] to the structure of certain corals... had the appearance of honeycomb on its surface. It was only a few millimeters thick at the most and could easily be scraped away from the inner table of the skull.” The surface, however is relatively uniform, with no protruding spicules. While Moodie related it to a meningioma, the associated lytic area and “hair-on-end” radiologic appearance seem more suggestive of a sarcoma. The possibility of malignant, rather than benign meningioma could be entertained. Indeed, Rogers’ [59] report of a “diffuse honeycomb type of hyperostosis in a 20<sup>th</sup> dynasty Egyptian skull was attributed to a sarcomatous meningioma, or simply a sarcoma. Number 17661 was noted to have “spongy hyperostosis with an irregular, pitted and cratered surface. What Abbott and Courville [1] referred to as “vertical spicules” seemed contained within and not extending beyond the lesion surface. Derivation of such spicules from deposition along stretched periosteally-sourced vessels was suggested. The report by Bianco et al. [10] illustrates an exostosis with major lytic component and irregular trabeculae incompatible with a diagnosis of meningioma. Cancer is more likely. Kompanje’s [43] republication of Salzmann’s [67] 1730 drawing of an alleged meningioma in a 43 year old contained insufficient details for diagnosis.

The partially healed osteitis that Bennike [9, p. 201] suggested might be a meningioma is especially of interest. She illustrated a Mid to Late Neolithic male with a relatively large lytic skull lesion, surrounded by circumferential minimally elevated, flat centripedally-oriented “columns.” Unlike previously published images of skull pathology, the edges (not the columns) have almost a rosette appearance, similar to the edges noted with metastatic carcinoma that is hypothesized [62] to be uterine, ovarian or breast cancer-related. The male-attribution of the skull makes the latter unlikely. Ricci et al. [59] considered meningiomas in the differential for the multicentric cranial lucencies, but the ill-defined borders are more suggestive of metastases.

#### 4. Discussion

Examination of the purported occurrences of meningiomas reported by Cook and Danforth [18] and of additionally recognized cranial pathology (delineated in Table 2) suggested the need for reevaluation. The sex ratio reported by Cook and Danforth [18] is converse to that observed in clinical samples [49,61,83]. ..That may be explainable, as their report seems to conflate several diagnostic entities. Herein are established criteria derived from medically confirmed cases and refined documentation of the prevalence of meningiomas over time.

Structural organization seems a major macroscopic characteristic that distinguishes among meningiomas, hemangiomas and marrow hyperplasia (e.g., porotic hyperostosis). Unique and apparently pathognomonic for meningiomas among these entities is the whorl pattern. Superficially having a sponge-like appearance, the appearance is the result of tumor-derived thin plates of bone (directed perpendicular to the original cortex). This contrasts with the non-uniform, thick bony corridors which constitutes the osseous component of hemangiomas. It also contrasts with the irregular surface distribution and variable thickness of the osseous component of marrow hyperplasia.

Examination for fulfillment of criteria for meningiomas, hemangiomas and sarcomas seems to provide a picture more parsimonious with the clinical literature. Excluding uncertain cases and those otherwise diagnosed, a 4:2 female predominance of meningiomas in archaeological cases is indistinguishable from clinical reports [32,49,61,83,85]. It should be noted that the epidemiology of archeologically-recognized meningiomas will likely underestimate their prevalence, given the propensity of this tumor to occur in later life [19]. However, that “cut off,” reducing potential recognition of full population prevalence, does not appear to affect sex ratio assessment.

Criteria are suggested for identifying meningiomas and distinguishing them from hemangiomas, bone marrow hyperplasia (porotic hyperostosis) and the macroscopically-observable surface spicules characteristic of osteosarcomas. The parallel trabeculae of meningiomas may be the product of the contained tumor cells ([20,66]. One final observation: Confidence in recognizing porotic hyperostosis may be compromised because of apparent similar macroscopic alterations to those seen with hemangiomas. After all, hemangiomas are more commonly recognized at autopsy than the hair-on-end phenomena [5].

## 5. Conclusions

The findings in reports alleging meningiomas in the paleopathology literature were subjected to comparison with the macroscopic appearance medically documented cases of the phenomenon and of disorders in its diagnostic differential. Those reports often appeared to be at variance with anatomical/medical findings in scientifically-identified cases, especially related to distinguishing hemangiomas and bone marrow hyperplasia (recognized in the form of porotic hyperostosis). While analogy, often used to characterize findings in past reports, failed to delineate the component considered significant, resulting in misinterpretations.

A uniform mesh surface pattern with contained whorls appears pathognomonic for meningiomas, in contrast to the non-uniform marrow expansion displacement of trabeculae in porotic hyperostosis and the non-uniform vascular displacement of trabeculae in hemangiomas. Criteria suggested for identifying meningiomas permits distinguishing from hemangiomas, bone marrow hyperplasia (porotic hyperostosis) and from the macroscopically-observable surface spicules characteristic of osteosarcomas. Reassessment of past attributions revealed few cases of meningiomas that could be confidently diagnosed as well as recognizing potential for misdiagnosis of porotic hyperostosis. Most of previous claims of meningiomas are not supported by criteria-based review, noting that the diagnosis has been incorrectly applied, at least in some instances to porotic hyperostosis.

**Funding:** This research received no external funding.

**Institutional Review Board Statement:** Not applicable

**Informed Consent Statement:** Not applicable

**Data Availability Statement:** Available at museum sites

**Acknowledgments:** None

**Conflicts of Interest:** The authors declare no conflict of interest.

## References

1. Abbott KH, Courville CB. 1939. Historical notes on the meningiomas. I. A study of hyperostosis in prehistoric skulls. *Bulletin of the Los Angeles Neurologic Association* **4**, 101-113.
2. Al-Rohdan RF, Laws ER Jr. 1990. Meningioma. A historical study of the tumor and its surgical management. *Neurosurgery* **26**, 832-847.
3. Anderson T. 1992. An example of meningiomatous hyperostosis from Medieval Rochester. *Medical History* **36**, 207-213.
4. Anegawa S, Hayashi T, Torigoe R, Furukawa Y. 1999. Diffuse calvarial meningioma: Case report and review of the literature. *Journal of Neurosurgery* **90**, 970-973.

5. Arana E, Marti-Bonmatí L. 1999. CT and MR imaging of focal calvarial lesions. *American Journal of Roentgenology* , 1683-1688.
6. Arana E, Diaz C, Latorre FF, Menor F, Revert A, Beltrán A, Navarro M. 1996. Primary intraosseous meningiomas. *Acta Radiologica* **37**, 937-942.
7. Banerji D, Inao S, Sugita K, Kaur A, Chhabra DK. 1994. Primary intraosseous orbital hemangioma: A case report and review of the literature. *Neurosurgery* **35**. 1131-1134.
8. Bastug D, Ortiz O Schochet SS. 1995. Hemangiomas in the calvaria: Imaging findings. *American Journal of Roentgenology* **164**, 683-687.
9. Bennike P. 1985. *Paleopathology of Danish Skeletons*. Akademisk Forlag: Copenhagen, Denmark.
10. Bianco P, Corsi A, Gattini F, Porta D. 2008. Facial reconstruction and meningioma-related hyperostosis in a 2000 BP man from the Peruvian Andes. *Journal of Paleopathology* **20**, 5-19.
11. Bir SC, Msaiti TK, Bollam P, Nanda A. 2015. Felix Platter and a historical perspective of the meningioma. *Clinical Neurology and Neurosurgery* **134**, 75-78.
12. Brothwell D. 1967. The evidence of neoplasms. In: Brothwell DJ, Sandison AT, eds. *Diseases in Antiquity*. Charles C Thomas: Springfield, IL, 320-345.
13. Brothwell M, Brothwell D. 2016. Evidence for ancient meningiomas and a probable case from Medieval Tarbat, Scotland. *International Journal of Paleopathology* **13**, 65-69.
14. Campillo D. 1991. The possibility of diagnosing meningiomas in paleopathology. *International Journal of Osteoarchaeology* **1**, 225-230.
15. Campillo D, Salas-Cuesta ME. 1995. Signs of meningiomas in a skull of the Mexican colonial period. *International Journal of Osteoarchaeology* **5**, 144-150.
16. Choi JS, Bae YC, Kang GB, Choi K-U. 2018. Intraosseous hemangioma of the orbit. *Archives of Craniofacial Surgery* **19**, 68-71.
17. Cook DC. 1985. Three cranial tumors from Late Woodland sites: Diagnosis and cultural implications. *Proceedings of the Indiana Academy of Science* **84**, 94.
18. Cook DC, Danforth ME. 2022. Meningiomas in ancient human populations. *Cancer* **14**, 1058. Doi: 10.3390/cancers14041058
19. Cucu AL, Costea CF, Perciaccante A, Caruleanu A, Turliuc S, Costachescu B, Poeta I, Turliuc MD. 2019. The history of Archne through historic descriptions of meningiomas from prehistory to the present. *World Neurosurgery* **128**, 37-46.
20. Cushing H. 1922. The cranial hyperostosis produced by meningeal endothelios. *Archives of Neurology and Psychiatry* **8**, 139-154.
21. Cushing H. 1923. Surgical end results in general with a case of cavernous hemangioma of the skull inn particular. *Surgery, Gynecology, Obstetrics* **32**, 303-308.
22. Cushing H, Eisenhardt L. 1938. *Meningiomas: Their Classification, Regional Behavior, Life History and Surgical End Results*. Charles C Thomas: Springfield, IL.
23. Czarnetzki A, Schwaderer E, Pusch CM. 2003. Fossil record of meningioma. *Lancet* **9381**, 408.
24. Daffner RH, Yakulis R, Maroon JC. 1998. Intraosseous meningioma. *Skeletal Radiology* **27**, 108-111.
25. Danforth ME, Kramer K, Cook DC, Cohen MN. 2019. The youngest meningioma. *International Journal of Osteoarchaeology* **29**, 1042-1050.
26. De Lumley MA. 1962. Lesions osseuses de l'homme de Castellar. *Bulletin de la Muse Anthropol Préhistorique Monaco* **9**, 191-205.
27. De Lumley H. 1969. Une Cabane Acheuléenne dans la Grotte du Lazaret (Nice). *Mémoire de la Société préhistorique française*, **1962**, 223-232.
28. De Lumley M-A, Piveteau J. 1969, 2018. Les restes humains de la grotte du Lazaret (Nice, Alpes-Maritimes). In: De Lumey, ed. *Grotte du Lazaret. Les restes humains fossiles de la Grotte du Lazaret (Archéologie/Préhistoire)*. CNRS: Paris. 224-231.
29. De Lumley Becam G, Colard T, Duplay J, Paquis P, Quatrehomme G. 1969. Pathologie de l'homme du Lazaret. In: De Lumley MA, ed. *Les restes humains fossils de la Grotte du lazaret (Nice, Alpes-Maritimes, France): des Homo erectus Europeen evolues en voie de Neandertalisation*. CNRS: Paris. 469-480.
30. Dihlmann W. 1981. Computed tomography in typical hyperostosis cranii (THC). *European Journal of Radiology* **1**, 2-8.
31. El-Sobky A, Elsayed SM, El Mikkawy ME. 2015. Orthopaedic manifestations of Proteus syndrome in a child with literature update. *Bone Reports* **3**, 104-108.
32. Escoda AP, Baudin PN, Mora P, Cos M, Gañan JH, Narváez JA, Aguilera C, Majós C. 2020. Imaging of skull vault tumors in adults. *Insights into Imaging*. **11**(1), 1. Doi:10.1186/s13244-019-0820-9
33. Ettinger SJ. 1983. *Textbook of Veterinary Internal Medicine: Diseases of the Dog and Cat*. Saunders Philadelphia.
34. Garfinkle J, Melançon D, Cortes M, Tampieri D. 2011. Imaging pattern of calvarial lesions in adults. *Skeletal Radiology* **40**, 1261-1273.
35. Ginsberg L. 1996. Radiology of meningiomas. *Journal of Neuro- Oncology* **29**, 229-238.
36. Gladkowska-Rzeczycka J. 1988. Tumors in antiquity in east and middle Europe. In: Ortner DJ, Aufderheide AC, eds. *Human Paleopathology: Current Synthesis and Future Options*. Smithsonian Press: Washington DC, 251-256.

37. Hanakova H, Vyhnanek L. 1981. Palaeopathologische Befunde aus dem Gebit der Tchechoslovakei. *Sbornik Narodniho Muzea v Praze* **37B(1)**, 1-90.
38. Hitzrot JM. 1917. Hemangioma cavernosum of bone. *Annals of Surgery* **65**, 475-482.
39. Huggins TJ, Ragsdale BD, Schnapf DO, Madewell JE, Youngblood L. 1981. RPC from the AFIP. *Radiology* **141**, 709-713.
40. Jayaraj K, Martinez S, Freeman A, Lyles KW. 2001. Intraosseous meningioma – a mimicry of Paget’s disease? *Journal of Bone and Mineral Research* **16**, 1154-1156.
41. Jónsdóttir B, Ortner DJ, Frohlich B. 2003. Probable destructive meningioma in an archaeological adult male skull from Alaska. *American Journal of Physical Anthropology* **122**, 232-239.
42. Kim KS, Rogers LF, Goldblatt D. 1987. CT features of hyperostosing meningioma en plaque. *American Journal of Roentgenology* **149**, 1017-1023.
43. Kompanje EJ. 2004. A patient with a large intra- and extracranial tumor, most probably a primary extradural meningioma, described in 1730. *Journal of Neuro-Oncology* **67**, 123-125.
44. Kostandy G, Ottley R, Salama S, Ghaly M, Taha H, Sosler B, Maqbool S, Ashamalla H. 2001. Intracranial meningiomas: A clinical update. *Resident and Staff Physician* **47(7)**, 35-48.
45. Landtblom AM. 2004. Did St. Birgitta suffer from epilepsy? A neuropathography. *Seizure* **131**, 161-167.
46. Lee JH (2008-12-11). *Meningiomata: Diagnosis, Treatment, and Outcome*. Springer Science & Business Media, 3–13
47. Luginbuhl H, Frankhauser R, McGrath JT. 1968. Spontaneous neoplasms of the nervous system in animals. *Progress in Neurological Surgery* **23**, 85-164.
48. MacCurdy GG. 1923. Human skeletal remains from the highlands of Peru. *American Journal of Physical Anthropology* **6**, 217-329.
49. Mehta N, Bhagwati S, Parulekar G. 2009. Meningiomas in children: A study of 18 cases. *Journal of Pediatric Neuroscience* **4**, 61-65.
50. Moodie RL. 1926. Studies in paleopathology. XVIII. Tumors of the head among pre-Columbian Peruvians. *Annals of Medical History* **8**, 394-412.
51. Nair P, Srivastava AK, Kumar R, Jain K, Sahu RN, Vij M, Jain M. 2011. Giant primary intraosseous calvarial hemangioma of the occipital bone. *Neurology India* **59**, 775-776.
52. Nakasu S, Hirano A, Shimura T, Llana JF. 1987. Incidental meningiomas in autopsy study. *Surgical Neurology* **27**, 319-322.
53. Niedermaier T, Behrens G, Schmid D, Schlecht I, Fischer B, Leitzmann MF. 2015. Body mass index, physical activity, and risk of adult meningioma and glioma: A meta-analysis. *Neurology* **85**, 1342-1350.
54. Okonkwo DO, Laws ER (2009). "Meningiomata: Historical Perspective". *Meningiomata* **2009**, 3–10.
55. Ortner DJ, Putschar WG. 1981. *Identification of Pathological Conditions in Human Skeletal Remains*. Smithsonian Press: Washington DC, p. 378.
56. Pechenkina K, Wenquan F, Xiaodong L. 2019. What’s that big thing on your head? Diagnosis of a large lesion on an Eastern Zhou skull from Henan, China. *International Journal of Paleopathology* **26**, 84–92.
57. Phemister DB. 1923. The nature of cranial hyperostosis overlying endothelioma of the meninges. *Archives of Surgery* **6**, 554-572.
58. Platter F. 1968. Felix Platter (1536-1614) Basle Physician. *Journal of the American Medical Association* **203**, 357-358.
59. Ricci R, Lama R, Di Tota G, Pietrangelo F, Vecchio F, Evangelista A., Capelli A, Capasso L. 1994. Skull osteolytic lesions in a XV century child: a case of childhood malignancy. *Journal of Paleopathology* **6**, 151-159.
60. Rogers L. 1949. Meningiomas in pharaoh’s peoples: Hyperostosis in ancient Egyptian skulls. *British Journal of Surgery* **36**, 423-424.
61. Rohringer M, Sutherland GR, Louw DF, Sima AA. 1989. Incidence and clinicopathological features of meningioma. *Journal of Neurosurgery* **71**, 665-672.
62. Rothschild BM. Metastatic cancer and multiple myeloma: One ancient, the other a disease of modernity? *Anthropologischer Anzeiger* (in press).
63. Rothschild BM, Martin LD. 2006. *Skeletal Impact of Disease*. New Mexico Museum of Natural History Press: Albuquerque, NM.
64. Rothschild BM, Schultze H-P, Pelligrini R. 2012. *Herpetological Osteopathology: Annotated Bibliography of Amphibians and Reptiles*. Springer-Verlag: Heidelberg, Germany.
65. Rowbotham GF. 1939. The hyperostosis in relation with the meningiomas. *British Journal of Surgery* **26**, 593-623.
66. Roy RN, Banerjee D, Chakraborty KN, Basu SP. 1971. Observations on radiological changes of bones in thalassaemia syndrome. *Journal of the Indian Medical Association* **57(3)**, 90-95.
67. Salzman DJ. 1730. Tumor capitis a carne fungosa productus & cum carie crania notabili conjunctus. *Acta Physico-Medica Academiae Caesareae Leopoldino-Carolinae Naturae Curiosum* **2**, 225-228.
68. Sargent N, Reilly EB, Posnikoff J. 1965. Primary hemangioma of the skull. Case report of an unusual tumor. *American Journal of Roentgenology* **95**, 874-879.
69. Satter AM, Talha KA, Rashid F, Selina F, Khan MD, Hossain AT, Shaikh AK. 2011. Invasion of meningioma cell in bony hyperostosis- An observational study of 34 cases. *Bangladesh Journal of Neuroscience* **27**, 78-82.



70. Schamall D, Teshler-Nicola M, Hübisch P, Kneissel M, Plenk H Jr. 1999. Differential diagnosis on ancient skeletal remains: Conventional methods and novel application of BSE-mode in SEM on a skull of the early Bronze Age. *Collegium Antropologicum* **23**, 843-894.
71. Sirianni JE, Byrnes JF, Odien JE. 2014. An osteoblastic infracranial meningioma en plaque: A curious case from the Erie County Poorhouse Cemetery. *Proceedings of the Annual Paleopathology Association Meeting (Calgary, Alberta, Canada. 8-9 April 2014)*. [https://physanth.org/documents/44/2014\\_AAPA\\_meetings\\_supp.pdf](https://physanth.org/documents/44/2014_AAPA_meetings_supp.pdf), viewed 6 April 2022.
72. Smrčka V, Kuželka V, Melkva J. 2003. Meningioma probable reason for trephination. *International Journal of Osteoarchaeology* **13**, 325-330.
73. Sosman MC. 1927. Radiology as an aid in diagnosis of skull and intracranial lesions. *Radiology* **9**, 396-404.
74. Starr CJ, Cha S. 2017. Meningioma mimics: Five key imaging features to differentiate them from meningiomas. *Clinical Radiology* **72**, 722-728.
75. Stead IM. 1991. *Iron Age Cemeteries in East Yorkshire: Excavations at Burton Fleming, Rudston, Garton-on-the-Wolds, and Kirkburn*. English Heritage Archaeological Report 22. British Museum Press: London, England.
76. Terstegge K, Schorner W, Henkes H, Heye N, Hosten N, Lanksch WR. 1994. Hyperostosis in meningiomas; MR findings in patients with recurrent meningioma of the sphenoid wings. *American Journal of Neuroradiology* **15**, 555-560.
77. Thillaud PL. 1996. *Paléopathologie Humaine*. Sceaux: Kronos.
78. Toynbee J. 1845. An account of two vascular tumors developed in the substance of bone. *Lancet* **2**, 676.
79. Tyson RA, Alcauskas ES. *Catalogue of the Hrdlička Paleopathology Collection*. San Diego Museum of Man: San Diego, CA.
80. Waldron T. 1998. An unusual cluster of meningiomas? *International Journal of Osteoarchaeology* **8**, 213-217.
81. Weber J, Czarnetzki A. 2002. A primary interosseous meningioma in a skull of the medieval period of southwestern Germany. *International Journal of Osteoarchaeology* **12**, 385-392.
82. Weber J, Spring A, Czarnetzki A. 2002. Parasagittales meningeom bei einem 32500 jahre alten schädel aus dem südwesten von Deutschland. *Deutsche Medizinische Wochenschrift* **127**, 2757-2650.
83. Wiemels J, Wrensch M, Claus EB. 2010. Epidemiology and etiology of meningioma. *Journal of Neuro-Oncology* **99**, 307-314.
84. Wisetsin S. 1990. Cephalography in thalassemic patients. *Journal of the Dental Association of Thailand* **40(6)**, 260-268.
85. Wyke BD. 1949. Primary hemangioma of the skull: A rare cranial tumor. Review of the literature and report of a case, with special reference to the roentgenographic appearances. *American Journal of Roentgenology* **61**, 302-316.
86. Yang Y, Guan J, Ma W, Li Y, Xing B, Ren Z, Su C, Wang R. 2016. Primary intraosseous cavernous hemangioma in the skull. *Medicine* **95(11)**, e3069. Doi: 10.1097/MD.0000000000003069
87. Zhang Q, Zhang Q, Han T, Zhu H, Wang Q. 2019. An Iron Age skull with a bone neoplasm from Nilka County, Xinjiang, China. *International Journal of Osteoarchaeology* **29**, 1034-1041.