CASE REPORT

A teachable moment: Identifying hyperostosis frontalis interna in a gross anatomy cadaver laboratory

Julia N. DiSalle¹, Alexis L. Novak², Prasad S. Dalvi³, Elisa M. Konieczko²


INTRODUCTION

Hyperostosis frontalis interna (HFI) is a condition with limited bone remodeling of the neurocranium. In HFI, newly formed cancellous bone is deposited on the inner lamina of the cranium forming irregular thickening on the internal surface of the frontal bone. Such thickening has been long documented in the literature, beginning in the early 1700’s with books written by Giovanni Batistta Morgagni, the father of anatomical pathology. This disease was further documented and detailed in the early 20th century [1-3]. HFI is usually not the primary cause of death for an individual and is often found through scans ordered for other possible diseases [4]. Sometimes, however, the hypertrophy of the frontal bone can be extensive and may result in compression of brain tissue [5].

Despite the wide-ranging literature on HFI, the etiology of this disease remains largely unknown [6]. Some authors have suggested insanity or dementia may cause this physical change in the cranium [1,5] while others have suggested hormonal causes [7-10] or metabolic disorders [4,11]. The literature is clear that there is marked female predominance in the epidemiology of this disorder, especially in post-menopausal elderly women [12,13]. However, HFI has also been reported in males with severe hypogonadism and testicular atrophy [9]. HFI is distinguished from other conditions, including acromegaly, Paget’s disease and bone cancer of the skull [4,10].

Gannon University teaches cadaver-based human gross anatomy to undergraduate college students in biology, chemistry, pre-physical therapy, and physician assistant majors [16]. These high volume high intensity courses involve both prosected cadavers and cadavers dissected by enrolled students [17]. One of the challenges of teaching cadaver-based anatomy is the extensive memorization students must achieve in order to be successful. Using pathologies discovered during dissection of the cadavers to supplement and enhance lecture material is one method used by course instructors to engage students in anatomical and clinical discussions. These discussions enhance what is often a learning experience dominated by extensive memorization.

CASE REPORT

An 80 year-old woman who was diagnosed with Alzheimer’s disease/vascular dementia at the time of death was found to have HFI with dural fusion to the calvarium during dissection in the Gannon University human gross anatomy laboratory (Figure 1). The examination of the internal side of the skull cap revealed that the inner table (lamina) of the calvarium was covered with large, irregular nodular bony thickening that was predominantly localized to the region of the frontal bones (Figure 1). Consequently, the frontal lobes were found to be compressed and subdural space was decreased by the nodular thickness of the frontal bones. The bony nodular thickening was not present on any part of the parietal or occipital bones. The thickness of the frontal bones was more marked and exceedingly dense than that of parietal or occipital bones, being apparently due to overgrowth of cancellous bony tissue (Figure 1). The diploe of the frontal bones was slightly reduced in amount. The dura mater was thickened and was adhered to the inner table of the skull and fused with, in particular, the frontal bone (Figure 2). The anterior end of falk cerebri contained a large bony plate, nearly a quarter of an inch thick and two inches long. The brain weightless less than the age-matching typical normal brain from another decedents without HFI (Table 1). The coronal sections of the brain described in this study revealed that both cerebral hemispheres were filled with a relatively larger amount of white matter composed of shallow sulci and atrophied gyri (Figure 3). In addition, the decedent’s brain showed atrophy of the frontal gyri, and the cerebral ventricles were found to be reduced in size (Figure 4).

DISCUSSION

HFI was first described by Morgagni more than 300 years ago in the early 1700’s, and since then HFI has been associated with frontal headaches, psychoneurosis, obesity, pregnancy, acromegaly, virilism, hypertrophisis, diabetes, and other hormonal and metabolic disorders [19,22]. Based on these case reports and studies, HFI has been included in Morgagni’s

1Department of General Medicine, Gannon University, Erie, PA, USA; 2Department of Biology, Gannon University, Erie, PA, USA

Correspondence: Prasad S. Dalvi, Department of Biology, Gannon University, Erie, PA, USA, Tel: 01814871-7651, E-mail: konieczko001@gannon.edu

Elisa M. Konieczko, Department of Biology, Gannon University, Erie, PA, USA, Tel: 01814871-5963, Email: dalvi001@gannon.edu

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syndrome (HFI, obesity, virilism), Stewart-Morel syndrome (HFI, obesity, mental disturbances), and Troell-Junet syndrome (HFI, acromegaly, toxic goiter, and diabetes mellitus) [19,22]. However, at present, HFI is viewed as an independent condition representing a phenomenon itself rather than related to any specific syndrome as suggested in the past [12]. HFI has been reported in 5-12% of the general population, but the magnitude of manifestation and frequency of HFI are much higher in the female population [12,23]. It is now accepted that HFI by itself does not cause a significant clinical disease and is usually an incidental finding, however, the bone growth can be exuberant and cause compression of the underlying brain tissue [5]. Past archeological investigations have shown that HFI was rarely present in historic populations regardless of geographical origin; however, on the other hand, the skulls from modern times have been found to develop HFI, predominantly in the post-menopausal female population [12]. Currently, the etiology of HFI remains uncertain, and therefore, several hypotheses have been proposed to explain HFI. One hypothesis proposes that during human evolution, a wider availability of food favored an increased metabolic rate which may have caused a higher incidence of HFI via increased levels of leptin, a satiety peptide that increases sympathetic tone and energy expenditure [24]. Another explanation for detecting HFI more

**TABLE 1**

<table>
<thead>
<tr>
<th>Brain</th>
<th>Weight (g)</th>
<th>Presence of Dura Mater (+ or -)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Case study brain</td>
<td>1046.31</td>
<td>+</td>
</tr>
<tr>
<td>Case study brain</td>
<td>975.77</td>
<td>-</td>
</tr>
<tr>
<td>#1</td>
<td>1147.26</td>
<td>+</td>
</tr>
<tr>
<td>#2</td>
<td>1435.43</td>
<td>+</td>
</tr>
<tr>
<td>#4</td>
<td>1141.90</td>
<td>+</td>
</tr>
<tr>
<td>#5</td>
<td>1417.67</td>
<td>+</td>
</tr>
<tr>
<td>#6</td>
<td>1007.12</td>
<td>*</td>
</tr>
<tr>
<td>#7</td>
<td>1121.97</td>
<td>**</td>
</tr>
<tr>
<td>#8</td>
<td>1401.24</td>
<td>***</td>
</tr>
<tr>
<td>#9</td>
<td>1315.59</td>
<td>-</td>
</tr>
<tr>
<td>#10</td>
<td>1367.86</td>
<td>-</td>
</tr>
<tr>
<td>#11</td>
<td>1458.64</td>
<td>-</td>
</tr>
<tr>
<td>#12</td>
<td>1219.85</td>
<td>-</td>
</tr>
<tr>
<td>#13</td>
<td>1216.73</td>
<td>-</td>
</tr>
</tbody>
</table>

* Did not have a cerebellum; ** half of the dura mater present; *** with only falx cerebri
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frequently in elderly population is that greater longevity has allowed HFI to become more prevalent and detectable in the aging population [15]. Another hypothesis is that prolonged high levels of estrogen during the reproductive period of life may be the primary triggering factor contributing to the greater frequencies of HFI seen in post-menopausal females [25], and therefore, the incidence in the female population has been reported to be much higher than in male population [12]. While HFI is found predominantly in females, it is not a purely female phenomenon. Males with hormonal irregularities, such as atrophic testis and/or feminism were found to have HFI of variable severity [9,12]. On the other hand, it has been shown that HFI is related to elevated androgen levels in pre-menopausal women [26]. At present, the most acceptable hypothesis regarding HFI pathogenesis is hormonal imbalance of the gonads.

HFI is usually not the primary cause of death for an individual and is often found through scans ordered for other possible diseases, and therefore, upon recognition of prominent HFI in imaging, it is important to distinguish it from other pathologic processes, such as leptomeningeal metastasis, hemorrhage, or infection [5]. While HFI is a benign process, prominent HFI may cause compression of the underlying soft tissues, inflammation and irritation of the meninges and pressure atrophy of the brain, for which surgical decompression is the treatment [27]. Cases of HFI, with or without neuropsychiatric disorders, have been reported by several authors [1,4,5,9,11]. These studies report that the obesity, diabetes, dyslipidemia and other metabolic disturbances, hormonal imbalance, post-menopausal state, and female gender are considered to be the risk factors of HFI. The only known risk

TABLE 2

Review of literature for the brain weight observed in the cases of HFI with neurological symptoms.

<table>
<thead>
<tr>
<th>Gender</th>
<th>Age (Years)</th>
<th>Pathology</th>
<th>Post-mortem brain weight (g)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Female</td>
<td>48</td>
<td>Melancholia, Loss of Memory</td>
<td>1150</td>
</tr>
<tr>
<td>Female</td>
<td>47</td>
<td>Dementia, Disorientation in time and place</td>
<td>1247</td>
</tr>
<tr>
<td>Female</td>
<td>62</td>
<td>Recurrent melancholia, dementia</td>
<td>1169</td>
</tr>
</tbody>
</table>

Figure 3) (A, B) Comparison of the serial coronal sections of the decedent’s brain on the left side of both images to a typical normal brain on the right side. The sections of the decedent’s brain show relatively larger amount of white matter.

Figure 4) (A-D) Comparison of the serial coronal sections of the decedent’s brain on the left side to a typical brain on the right side. (A, B) The decedent’s brain shows reduced volume of the lateral ventricles (black arrows) compared to the typical normal brain (white arrows). (C) The decedent’s brain shows decreased size of the third ventricle (black arrow) compared to the typical normal brain (white arrow). (D) The decedent’s brain shows reduced volume of the fourth ventricle (black arrow) compared to the typical normal brain (white arrow).
The present case of HFI was the first case found at the human gross anatomy laboratory at Gannon University within past 20 years, and therefore, this provides information about the brain weight and associated structural changes (Table 2) [1]. In the present case study, we measured the brain weight (Table 1) and compared it with another typical normal age-matching brain without HFI (Figures 3 and 4). The brain weight of the decedent was found to be decreased (Table 1). The presence of decreased ventricular volume suggests the compression of the brain tissue and a decrease in brain volume that could have a major clinical significance as it may indicate the beginning of degenerative processes of the brain. Indeed, the decedent was diagnosed with Alzheimer’s disease/vascular dementia at the time of death. Among other changes that confirm the neurodegenerative process, we observed that the frontal gyri were atrophied, and both cerebral hemispheres had increased white matter (Figures 3 and 4).

The first human dissection was recorded around 300 BC [28], and human cadavers have been used as an educational tool since the 16th or 17th century [29]. However, there is an ongoing debate about the practicality and usefulness of cadavers as an educational tool [30]. With the advancement in technology, the way anatomy is taught is changing, and the use of cadavers may become obsolete [31,32]. Computer technology has provided an alternative to both cadavers and textbook images, and is becoming popular [31,32]. Despite this, many academic institutions continue to believe that cadaveric dissection is still the best way to educate students about the complexity of the human body and continue to use the human cadavers in the classroom [33]. Supporters for the use of cadavers argue that human cadavers not only can provide the visual and tactile experiences necessary to learn and apply anatomical knowledge, but also provide in situ information and details about the complex human body structures with or without any existing pathology [34]. At Gannon University, cadaver-based human gross anatomy is taught to undergraduate college students in biology, chemistry, pre-professional therapy, and physician assistant majors [16]. Quite often we come across cadavers with variability in anatomical structures and pathologies. The enrolled students are very interested in these anomalies and diseases, and ask questions about them. Often, the undergraduate students are quite passive in their learning. This opportunity clearly engages the students and makes them active learners. It allows the instructors and teaching assistants to provide additional information about the anomalies and pathologies seen, and we incorporate this material in course assessments.

The present case of HFI was the first case found at the human gross anatomy laboratory at Gannon University within past 20 years, and therefore, this finding was quite novel to the instructors, teaching assistants and students. Gross anatomy laboratory instructors can use similar cases to engage students in anatomical and clinical discussions that can promote long-term retention of pathological information useful for the future clinical and health professionals. Medical professionals of all expertise believe that cadaveric dissection enables learning anatomy with relevant clinical correlates, and complemented with other innovative learning methods lays strong foundation critical to development of clinical skills [34].

CONCLUSION

Endocrine imbalance, metabolic disorders, and/or neuropsychiatric abnormalities may be responsible for the development of HFI in some cases. The epidemiology of HFI indicates marked female predominance. HFI is usually not the primary cause of death for an individual and is often found through scans ordered for other possible diseases. Although the etiopathogenesis of HFI is not quite clear, it is important to correlate its development to an underlying manifestation of an endocrine, metabolic and neuropsychiatric condition. Gross anatomy laboratory instructors can use similar cases to engage students in anatomical and clinical discussions through active learning strategies that can promote long-term retention of pathological information useful for the future clinical and health professionals. These discussions enhance what is often a learning experience dominated by extensive memorization.

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REFERENCES

23. Nikolic S, Djonic D, Zivcovic V, et al. Rate of occurrence, gross
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