Spectrum, Etiology and Clinical Features of Intracranial Hypertension Presenting in Neuro-Ophthalmology Clinic At AFIO Rawalpindi

Umar Ijaz, Kamran Saeed*, Imran Sarwar**, Noman Nazir, Hassan Sajjad Rathore, Muhammad Shahid

Armed Forces Institute of Ophthalmology/National University of Medical Sciences (NUMS) Rawalpindi Pakistan, *Combined Military Hospital Lahore/National University of Medical Sciences (NUMS) Pakistan, **Combined Military Hospital Abbottabad/National University of Medical Sciences (NUMS) Pakistan

ABSTRACT

Objective: To assess the clinical spectrum and various etiological factors of idiopathic intracranial hypertension in Pakistani patients.

Study Design: Cross-sectional study.

Place and Duration: Neuro-Ophthalmology Clinic, Armed Forces Institute of Ophthalmology, Rawalpindi Pakistan, from Jan 2017 to Jul 2018.

Methodology: A total of 64 individuals with intracranial hypertension (IH) were screened. On the basis of inclusion criteria 32 patients were selected for the study. The demographics, clinical features and etiologies were recorded for each patient.

Results: High frequency was noted among females, affecting 26 (84.37%) as compared to males who were 6 (18.75%) in number. Half of the patients had body weight lying in range of overweight, 16 (50%), however, only 5 (15.6%) patients fell into the category of obesity. Among clinical features, headache was the most prominent symptom seen in 17 (53.12%) patients and severe disc swelling seen in 19 (59.3%) was the most common sign. Most prevalent systematic association was anemia, 11 (34.3%), and the most prominent etiology was idiopathic intracranial hypertension, being 26 (81.25%).

Conclusion: There was high frequency of intracranial hypertension in the females with obesity, pregnancy and in the age group of 20-30 years.

Keywords: Etiology, Intracranial hypertension, Papilledema.

How to Cite This Article: Ijaz U, Saeed K, Sarwar I, Nazir N, Rathore HS, Shahid M. Spectrum, Etiology and Clinical Features of Intracranial Hypertension Presenting in Neuro-Ophthalmology Clinic at AFIO Rawalpindi. Pak Armed Forces Med J 2022; 72(1): 12-15. Doi: https://doi.org/10.51253/pafmj.v72i1.6868

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (https://creativecommons.org/licenses/by-nc/4.0/), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

INTRODUCTION

Intracranial hypertension (IH) comprises of neurological disorders in which cerebrospinal fluid (CSF) pressure is increased within the skull.¹ The pressure of CSF above 20cm H₂O in children and above 25cm H₂O in adults indicates increased intracranial pressure (ICP).² IH usually results from a neurological injury or insult but it may be idiopathic. Variety of clinical features are associated with IH including nausea, vomiting, headache, altered consciousness level, ocular palsies, papilledema and backache.³ If remained unchecked, papilledema can result in optic atrophy, visual disturbance and blindness.⁴

Increasing rate of obesity around the world is leading towards some life-threatening disorders like idiopathic IH.⁵ It is a rare disorder with no clinical, radiographic and laboratory evidence in relation to infection, vascular abnormality, lesion and even with hydrocephalus.^{6,7} The prevalence of idiopathic IH among female population is 3.5 per 100,000 in the age group of 15-44 years. However, this ratio amongst the

Correspondence: Dr Umar Ijaz, Classified Eye Specialist, Armed Forces Institute of Ophthalmology, Rawalpindi Pakistan males is 0.9 per 100,000. Chances of idiopathic IH are high when body mass index (BMI) increases to 26.⁸ One mechanistic theory claimed that obesity predisposes patients to raised intra-abdominal pressure, which leads to raised intrathoracic pressure, central venous pressure and finally to idiopathic IH. However, results of other studies were in contrast with this theory and observed that lower body (gynecoid) and increased level of estrogenicity causes IH instead of central venous pressure or intrabdominal pressure.⁹

Despite the emerging prevalence of IH in various countries, Pakistan has lagged behind in exploring different aspects of this disease. Rationale of conducting this study was the absence of adequate local data of our population pertaining to IH. Owing to the increasing obesity throughout the world, it is expected that IH will continue to increase in coming years.

METHODOLOGY

This cross-sectional study was conducted to assess the etiologies and clinical features of patients labeled with IH. For this purpose, all the 64 patients, referred to neuro-ophthalmology OPD of the Armed Forces Institute of Ophthalmology (AFIO), Rawalpindi from January 2017 to July 2018 were included in the

Received: 13 Apr 2021; revision received: 17 May 2021; accepted: 19 May 2021

study. Patients were selected through universal sampling, irrespective of the age, gender, ethnicity and residence.

Inclusion Criteria: Patients having active IH at the time of enrollment with headache and optic disc swelling were included in the study.

Exclusion Criteria: Patients with co-existing ocular diseases such as glaucoma, uveitis, ocular infection and corneal diseases were excluded from the study. Patients with papilledema secondary to space occupying lesion of brain were also not included in the study.

The study proposal was approved by the Ethical Committee of AFIO, Rawalpindi (221/ERC/AFIO).

Thirty-two patients, who were diagnosed with IH, were inducted in the study. Written informed consent was obtained from the selected participants. Detailed ophthalmic history and clinical examination of the selected participants was done. Their visual field charting, MRI brain, MR/CT venography and cerebrospinal fluid analysis were performed. Laboratory tests comprising complete blood count, urine routine examination, anti-nuclear antibody levels, thyroid hormone levels, and calcium/vitamin D levels were carried out.

The demographics of patients were noted, including age, gender and weight. Clinical features such as headache, tinnitus, diplopia, and transient visual obscurations, were recorded. Possible etiologies including cerebral venous sinus thrombosis (CVST), vasculitis, uveitis, juvenile and adult idiopathic intracranial hypertension were registered.

Statistical Package for Social Sciences (SPSS) version 21 was used for the data analysis. Quantitative Variable were summarized as Mean \pm SD, while qualitative variables were summarized as frequency and percentages.

RESULTS

Thirty-two patients, who were diagnosed with idiopathic intracranial hypertension (IIH), were selected after applying inclusion and exclusion criteria. The mean age of participants was 21.5 ± 3.4 years, (range: 13-56 years). Higher frequency of IIH was noted among females 26 (84.37%) as compared to males 6 (18.75%). The demographic data of participants was shown in Table-I. Half of the patients had weight lying in range of overweight 16 (50%). However, only 5 (15.6%) patients were obese. The mean weight was 78.88 \pm 2.54 kg.

Table-II depicted clinical features of patients. In total, 17 (53.12%) patients reported headaches. Tinnitus

was present in 4 (12.5%) patients, diplopia in 3 (9.3%), and transient visual obscurations in 7 (21.8%) patients. However, 1 (3.1%) patient was asymptomatic. Mean intracranial pressure was 32.1 ± 2.3 cm H₂O.

Table-I: Demographic characteristics of the participants.

Characteristics	Frequency (%)
Gender	
Male	6 (18.75%)
Female	26 (84.37%)
Age (years)	
10-20	4 (12.5%)
21-30	12 (37.5%)
31-40	9 (28.1%)
41-50	5 (15.6%)
51-60	2 (6.2%)
Weight (kg)	
Underweight	1 (3.1%)
Normal	10 (31.2%)
Overweight	16 (50.0%)
Obese	5 (15.6%)

Table-II: Clinical features of participants.

Clinical Features	Frequency (%)
Headache	17 (53.1%)
Tinnitus	4 (12.5%)
Diplopia	3 (9.3%)
Transient visual obscurations	7 (21.8%)
Asymptomatic	1 (3.1%)

The signs observed for patients were; severe visual loss 3 (9.3%), moderate visual loss 2 (6.2%), mild disc swelling 5 (15.6%), severe disc swelling 19 (59.3%), swelling with hemorrhages 4 (12.5%), optic atrophy unilateral 3 (9.3%), periphlebitis 1 (3.1%), and uveitis 1 (3.1%).

The risks factors for IH were found to be systematic associations and use of medications. Systematic associations in some patients were reported with pregnancy 1 (3.1%), anemia 11 (34.3%), sleep apnea 3 (9.3%), and hypertension 6 (18.7%). Other systematic associations found in the patients were; polycystic ovary disease 1 (3.1%), increased PTH level 3 (9.3%); parathyroid adenoma 1 (3.1%), hypothyroidism 2 (6.25%), celiac disease 2 (6.25) and menstrual irregularities 3 (9.3%). Some patients had a history of medication use. These medications included hormonal supplements 6 (18.75%), vitamin A 2 (6.25%) and oral steroids 2 (6.25%). Only one case required surgical intervention.

The etiologies of participants have been shown in the Table-III. The most prominent etiology was IIH (81.25%). This was followed by cerebral venous sinus thrombosis (12.5%), followed by uveitis (3.1%) and juvenile IIH (3.1%).

Etiologies	Frequency (%)
Juvenile Idiopathic Intracranial	1 (2 1 %)
Hypertension	1 (3.170)
Cerebral Venous Sinus Thrombosis	4 (12.5%)
Idiopathic intracranial hypertension	26 (81.25%)
Uveitis	1 (3.1%)

Table-III: Etiologies of raised intracranial hypertension.

DISCUSSION

The findings of our study revealed that obesity was the leading factor of IH in Pakistan. Majority of the participants were in the age group of 21-40 years. There was higher frequency of IIH among females as compared to males. This observation was supported by previous works of Radhakirshnan *et al*,¹⁰ Raoof *et al*.¹¹ In previous studies, researchers observed severe IH in the females of age group 20-30 years. With obesity and pregnancy patients developed severe complications in their vision. In our study, 3 (9.3%) patients had obesity and 16 (50%) were reported to be overweight. This was in accordance with the study by Sugerman *et al*.¹² Another study by Ottridge *et al*,¹³ found that IH occurs mostly in females between ages 20-44 years with weight above the normal.

Clinical features of the patients had variation. Headache was observed in majority of cases 22 (68.75%). This indicated association of headache with IH. This was followed by transient visual obscurations (31.25%), tinnitus (12.5%), and diplopia (9.3%). Previous research work of Hoffman *et al*,¹⁴ has shown association of headache (100%) with IH, which supports findings of the present study.

Other important systematic associations included anemia and hypertension. Ananth *et al*,¹⁵ previously indicated linkage of IH with anemia, whereas, association of hypertension with IH has previously been highlighted by Pal *et al*.⁴ Association of preganancy has been reproted by previous research work of Stevens *et al*,¹⁶ our study reported pregnancy in only one individual (3.1%).

Use of different medications can also be associated with IH. This has been reported by Tan *et al*,¹⁷ and Uldall *et al*.¹⁸ Similar findings were deduced from the present research. The patients had history of using various medications such as oral steroids, thyroxine, isotrenoin. However, the most prominant medications among patients were hormonal supplements.

Numerous research works, such as Fiyaso *et al*,¹⁹ Aojula *et al*,²⁰ and Zhang *et al*,²¹ had highlighted the high prevalence of IIH. Our study was in accordance with these research works as IIH accounted for 26 % of

the cases. Although our study has some limitations like small sample size and lack of diversity, the importance of our study lies in the inclusion of both subjective and objective data. It is also one of the first local studies encompassing neuro-ophthalmological dimension of IH in Pakistani population. However, we do need more studies with a more diverse and large sample in future.

The aggressive course of IH can result in blindness over a short period of time. High frequency of IH in females with age range of 20-30 years, obesity and pregnancy indicated high susceptibility of this population. Thus, individuals showing relevant clinical features and risk factors should be considered for early diagnosis in order to avoid adverse outcomes.

CONCLUSION

There was high frequency of IH in the females with obesity, pregnancy and in the age group 20-30 years.

Conflict of Interest: None.

Authors' Contribution

UI: Conception manuscript writing data collection, KS: Manuscript writing data collection, IS: Manuscript writing data collection, NN: Manuscript writing statistical analysis, HSR: Design proof reading referencing, MS: Data collection interpretation of data.

REFERENCES

- 1. Zanon E, Pasca S. Intracranial haemorrhage in children and adults with haemophilia A and B: a literature review of the last 20 years. Blood Transfus 2019; 17(5): 378-384.
- Mondragon J, Klovenski V. Pseudotumor Cerebri. In: StatPearls. Treasure Island (FL): StatPearls Publishing; July 10, 2020, Available at: https://www.ncbi.nlm.nih.gov/books/NBK536924/
- Godoy DA, Núñez-Patiño RA, Zorrilla-Vaca A, Ziai WC, Hemphill JC. Intracranial hypertension after spontaneous intracerebral hemorrhage: a systematic review and meta-analysis of prevalence and mortality rate. Neurocrit Care 2019; 31(1): 176-187.
- 4. Pal A, Sengupta P, Biswas D, Sen C, Mukherjee A, Pal S. Pattern of Idiopathic Intracranial Hypertension in Indian Population. Ann Ind Acad Neurol 2019; 22(1): 47-51.
- 5. Traber GL, Weber KP, Sabah M, Keane PA, Plant GT. Enhanced depth imaging optical coherence tomography of optic nerve head drusen: a comparison of cases with and without visual field loss. Ophthalmol 2017; 124(1): 66-73.
- Mangalore S, Rakshith S, Srinivasa R. Solving the riddle of "Idiopathic" in idiopathic intracranial hypertension and normal pressure hydrocephalus: an imaging study of the possible Mechanisms - Monro-Kellie 3.0. Asian J Neurosurg 2019; 14(2): 440-452.
- Mollan SP, Davies B, Silver NC, Shaw S, Mallucci CL, Wakerley, et al. Idiopathic intracranial hypertension: consensus guidelines on management. J Neurol Neurosurg Psychiat 2018; 89(10): 1088-1100.
- Markey KA, Mollan SP, Jensen RH, Sinclair AJ. Understanding idiopathic intracranial hypertension: mechanisms, management, and future directions. Lancet Neurol 2016; 15(1): 78-91.

- Mollan SP, Ali F, Hassan-Smith G, Botfield H, Friedman DI, Sinclair AJ. Evolving evidence in adult idiopathic intracranial hypertension: pathophysiology and management. J Neurol Neurosurg Psychiat 2016; 87(9): 982-992.
- Radhakrishnan K, Ahlskog JE, Cross SA, Kurland LT, O'Fallon WM. Idiopathic intracranial hypertension (pseudotumor cerebri). Descriptive epidemiology in Rochester, Minn, 1976 to 1990. Arch Neurol 1993; 50(1): 78-80.
- 11. Raoof N, Sharrack B, Pepper IM, Hickman SJ. The incidence and prevalence of idiopathic intracranial hypertension in Sheffield, UK. Eur J Neurol 2011; 18(10): 1266-1268.
- Sugerman HJ, De-Maria EJ, Felton WL, Nakatsuka M, Sismanis A. Increased intra-abdominal pressure and cardiac filling pressures in obesity-associated pseudotumor cerebri. Neurol 1997; 49(2): 507–511.
- 13. Ottridge R, Mollan SP, Botfield H, Frew E, Ives NJ, Matthews T, et al. Randomised controlled trial of bariatric surgery versus a community weight loss programme for the sustained treatment of idiopathic intracranial hypertension: the Idiopathic Intracranial Hypertension Weight Trial (IIH:WT) protocol. BMJ Open 2017; 7(9): e017426.
- Hoffmann J, Mollan SP, Paemeleire K, Lampl C, Jensen RH, Sinclair AJ. European headache federation guideline on idiopathic intracranial hypertension. J Headache Pain 2018; 19(1): 93-95.

- Ananth JV, Sudharshan S, Selvakumar A, Devaleenal BJ, Kalaivani K, Biswas J. Idiopathic intracranial hypertension associated with anemia, secondary to antiretroviral drug in a human immunodeficiency virus positive patient. Ind J Ophthalmol 2018; 66(1): 168-169.
- Stevens SM, McClelland CM, Chen JJ, Lee MS. Idiopathic Intracranial Hypertension in a Mother and Pre-pubertal Twins. Neuroophthalmol 2019; 43(1): 49-52.
- 17. Tan ACS, Tan GS, Denniston AK, Keane PA, Ang M, Milea D, et al. An overview of the clinical applications of optical coherence tomography angiography. Eye (Lond) 2018; 32(2): 262-286.
- Scotton WJ, Botfield HF, Westgate CS, Mitchell JL, Yiangou A, Uldall MS, et al. Topiramate is more effective than acetazolamide at lowering intracranial pressure. Cephalal 2019; 39(2): 209-218.
- Fisayo A, Bruce BB, Newman NJ, Biousse V. Overdiagnosis of idiopathic intracranial hypertension. Neurol 2016; 86(4): 341-350.
- Aojula A, Mollan SP, Horsburgh J, Yiangou A, Markey KA, Mitchell JL, et al. Segmentation error in spectral domain optical coherence tomography measures of the retinal nerve fibre layer thickness in idiopathic intracranial hypertension. BMC Ophthalmol 2018; 17(1): 257.
- 21. Zhang X, Medow JE, Iskandar BJ, Wang F, Shokoueinejad M. Invasive and noninvasive means of measuring intracranial pressure: a review. Physiol Meas 2017; 38(8): R143-R182.