A Study of Cognitive Impairment in HIV-TB Co-Infection

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Abstract: There is an increase in HIV-Tuberculosis co-infection cases which might affect the severity of the cognitive impairment in HIV positive individuals due to the impact of tuberculosis on HIV. This study is designed to evaluate and compare the cognitive function in HIV seropositive individuals with and without Tuberculosis.50 HIV seropositive individuals with Pulmonary Tuberculosis and 50 HIV seropositive individuals with Pulmonary Tuberculosis and 50 HIV seropositive individuals without tuberculosis are the study subjects.Cognitive function was evaluated using International HIV Dementia Scale.There is significant difference in recall and psychomotor speed score and total score in IHDS among two groups.The total score of <10 was found in 76% of cases and 58% of controls which is significant for cognitive impairment. Early and periodic neuropsychological screening of HIV positive asymptomatic individuals is a must in future to spot the cognitive deficits at an early stage.The current study shows the impact of Tuberculosis on HIV and its cognitive function. Hence the assessment and treatment strategies should consider the Tuberculosis co-infection.

Key Words: cognitive function, HIV infection, psychomotor speed, recall, treatment, Tuberculosis.

I. Introduction

Human Immune Deficiency Virus (HIV) enters the central nervous system (CNS) early during the infection^[1]. Neurocognitive impairments in overall HIV population appear to be nearly 50% ^[2]. The wide range of neurocognitive complications of HIV are grouped under the name of HAND(HIV associated neurocognitive disorders) and depending on the degree of the impairment, there are three categories; progressing in disabilities from asymptomatic neurocognitive impairment (ANI) to HIV associated mild neurocognitive impairment (MND) and to HIV associated dementia (HAD). Since the introduction of HAART in 1996, the incidence of moderate or severe dementia fell from about 7% in 1989 to only 1% in 2000^[3].

HIV - TB co infection: According to World Health Organisation (WHO), there were an estimated 1.1 million HIV positive new Tuberculosis (TB) cases globally in 2011^[4]. HIV and TB affect each other's clinical course and increase each-others progression ^{[5-9].} The higher mortality is due to the progression of AIDS rather than TB probably due to the fact that MTB increases viral replication ^[24]. There are inadequate studies on cognitive function in HIV –TB co-infection. There is an increase in HIV-TB co-infection cases which might affect the severity of the cognitive impairment in HIV positive individuals due to the impact of tuberculosis on HIV.Hence this study is designed to evaluate and compare the cognitive function in HIV seropositive individuals with and without Tuberculosis.

HIV-Associated Neurocognitive Disorders In India: In studies done by Satischandra et al., ^[10] in 2000; Wadia et al., ^[11] in 2001; the prevalence of HIV-Associated Dementia (HAD) is estimated to be less than 6%. In a study done by Riedel et al., ^[12] in 2006, 35% of HIV individuals scored in the impaired range on the International HIV Dementia Scale vs.15% of controls. Neurocognitive impairment rates > 55% in participants with advanced disease in a study done by Yepthomi et al., ^[13] in 2006 and across disease stages in a study by Gupta JD et al., ^[14] in 2007.

II. Objective

The objective of the study is to compare the cognitive function in HIV seropositive individuals with and without Pulmonary Tuberculosis.

III. Methodology

50 HIV seropositive individuals who are diagnosed of Pulmonary Tuberculosis (maintaining the inclusion and exclusion criteria) are the Cases in the study. Controls for the study are 50 HIV seropositive individuals without tuberculosis and matched with respect to age, sex and education to the study subjects.

Inclusion criteria: 1) Patients with Diagnoses of HIV and Pulmonary Tuberculosis. Those who were newly diagnosed and started on ART and DOTS treatment from 1 month,2) Patients with minimum of primary school

education,3) Both male and female patients,4) Age group between 18 and 50 years,5) Those who gave the consent.

Exclusion criteria:1) Patients with past history of psychiatric illness not attributable to either HIV or Tuberculosis, 2) Patients having other medical disorders like Diabetes mellitus (DM), Hypertension (HTN), thyroid and other endocrine disorders and renal disorders and other chronic debilitating medical conditions known to cause cognitive impairment, 3) Patients with substance dependence, 4) Individuals suffering from any sensory impairment that is visual or hearing impairment or learning disability which may serve as hindrance in performing the test,5) Patients who refused to give consent.

Cognitive function was evaluated using International HIV Dementia Scale (IHDS) after assessing inclusion and exclusion criteria. IHDS consists of three subtests (i) timed finger tapping (ii) timed alternating hand sequence test, and (iii) recall of four items in 2 minutes. The IHDS is now used in many studies as a screening test to detect dementia in HIV patients. Total International HIV Dementia Scale Score. This is the sum of the scores on items 1-3. The maximum possible score is 12. A patient with a score of ≤ 10 should be evaluated further for possible dementia^[15].

IV. Results

Motor Spood	Groups	Sample Size	Min	Max	Means	SD	t value	df	P value		
Motor Speed	Cases	50	03	04	3.84	0.37			P > 0.05		
	Controls	50	03	04	3.92	0.27	1.23	98			
	Cases	50	02	04	3.32	0.62					
Recall	Controls	50	02	04	2.92	0.72	2.97	98	P < 0.01		
Psychomotor	Cases	50	01	04	1.96	0.81					
Speed	Controls	50	01	04	2.5	1.05	2.84	98	P < 0.01		
	Cases	50	08	12	9.16	1.13					
Total Score	Controls	50	08	12	9.68	1.30	2.26	98	P< 0.05		

TABLE 1. International Hiv Dementia Scale

There is significant difference in recall and psychomotor speed score (p<0.01) and total score (P<0.05) in IHDS among two groups. The mean total score is <10 among both the groups.

TABLE 2: Comparison of International HIV Dementia Scale- total sc
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Groups	Score <10	Score >10	Total
cases	38	12	50
	(76.00)	(24.00)	(100.0)
controls	29	21	50
	(58.00)	(42.00)	(100.0)
Total	67	33	100

 x^2 Test value is 7.18 with 1 d.f. P < 0.01.The total score of <10 was found in 76% of cases and 58% of controls which is significant for cognitive impairment.



Figure 1: Ihds Total Score

V. Discussion

The human immunodeficiency virus (HIV) is frequently accompanied by progressive and severe cognitive loss referred to as the acquired immune deficiency syndrome (AIDS) dementia complex ^[16].Cognitive impairment is believed to be due to the direct effects of the retrovirus on the brain rather than to secondary complications of immunosuppression ^[17].The presence of dementia will interfere with ART treatment. ^[18].

This study has revealed that there is statistically significant difference in cognitive impairment in HIV-TB co-infected individuals as compared to HIV seropositive individuals without tuberculosis.

The International HIV Dementia scale(IHDS) is the sensitive test used to asses cognitive function in HIV positive individuals in many studies^[19,20]. In the present study there was statistically significant difference (p<0.01) in recall and psychomotor speed score in IHDS among two groups. The total score <10 is significant for cognitive impairment. This was found in 76% individuals of cases and 58% of controls. There was statistically significant difference (p<0.05) in the mean total score of IHDS among two groups. The mean total score in cases is 9.16 and in control group is 9.68.

These results were similar to other studies conducted in India. KarthigaipriyaMuniyandi et.al, ^[21] study showed the International HIV Dementia Scale was the most sensitive instrument and 63.6% the patients had abnormal scores in this scale.Riedel ^[12] has administered the IHDS scale to the asymptomatic HIV positive population in Pune and found that 35% of them have abnormal scores in this test.

The significant difference in impairment of cognitive function in HIV-TB co-infected individuals to HIV seropositive individuals might be because of the impact of TB on HIV infection. Studies show that TB causes increase in viral replication and impaired immunity status in HIV –TB co-infected individuals.

Early and periodic neuropsychological screening of HIV positive asymptomatic individuals is a must in future to spot the cognitive deficits at an early stage. This may help us in arresting the progression of the disease by early institution of ART therapy as HIV dementia is an indication for ART. The current study shows the impact of Tuberculosis on HIV and its cognitive function. Hence the assessment and treatment strategies should consider the Tuberculosis co-infection.

VI. Conclusion

Cognitive impairment was higher in patients of HIV with Pulmonary Tuberculosis than in patients with HIV infection alone. The limitations of the study are sample size is small and it is a cross sectional study. A longitudinal study would enable to determine stability of the cognitive deficits Hence, it is imperative that cognitive impairment should be recognized early in HIV positive patients to enhance drug compliance to ART. This will improve quality of life, increase longevity of life, and arrest further deterioration of brain function.

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