

Original Research Article

To Evaluate the Efficacy of DOTS Chemotherapy in Extra Spinal Osteoarticular Tuberculosis

Pradeep Kumar Saini¹, Sanat Singh², V. K. Goyal³, Sumant Sinha⁴, Lalit Kumar⁵, Gaurav Banoriya⁶

¹Consultant, Kasturba hospital BHEL, Bhopal, Madhya Pradesh, India

²Associate Professor, People's College of Medical science and Research Centre, Bhopal, Madhya Pradesh, India

³DDU Hospital, Hari Nagar, New Delhi, India

⁴Senior Consultant, DDU Hospital New Delhi, India

⁵Consultant, Patna, Bihar, India

⁶Consultant, Jhansi, Uttar Pradesh, India

***Corresponding author**

Dr. Pradeep Kumar Saini

Email: drpradeepsaini@gmail.com

Abstract: Noncompliance is one of the major issues with the treatment of tuberculosis (TB), which can lead to treatment failure. A short-course of chemotherapy as recommended by World Health Organization under the umbrella term of Directly Observed Treatment Short Course (DOTS) therapy seems to be effective. The main aim in present study was done to evaluate the efficacy of short term intermittent chemotherapy in DOTS regimen in patients with extra spinal osteoarticular TB. A prospective study was performed including 20 patients with extra spinal osteoarticular TB between October 2008 to June 2010 in the Department of Orthopedics, DDU Hospital, and Delhi. All the patients were given the DOTS regime as recommended by WHO and followed up at intervals of one month during the treatment for assessing the clinical improvement and compliance of the patient. The results in present study, there were 85% males and 15% were females. 40 % were students followed by the labourer (20%). Hip (50%) was the most commonly involved joint followed by short bones of hand and foot (20%). All (100%) patients had an elevated ESR at the time of presentation. Eighty percent patients had shown increase in weight at the end of 6 month of treatment. All patients of extra spinal osteoarticular TB showed improvement. Treatment was found sufficient for 80% of the patients. In 20% cases treatment duration was extended. In discussion the Short term intermittent chemotherapy in DOTS regimen was optimum for the treatment of extra spinal osteoarticular TB and was associated with good compliance.

Keywords: short term intermittent chemotherapy, extra spinal tuberculosis, DOTS regimen.

INTRODUCTION

According to World Health Organization (WHO), every year approximately 8.4 million people develop tuberculosis and more than 3 million patients die due to it each year [1]. Young adults in developing countries are mostly affected. India accounts for about one third of the total tuberculosis population of the world [1].

Tuberculosis involving joint other than spine is a form of disease usually secondary to a concomitant or old pulmonary Koch's. An early diagnosis along with proper chemotherapy is the key for excellent outcome [1]. The WHO has recommended category based treatment regimens of various forms of tuberculosis also known as short course chemotherapy (SCC) [2].

According to WHO, spinal tuberculosis is a serious form of tuberculosis and hence enclosed with

category-I whereas extra spinal tuberculosis is less severe and is therefore included in category-III [3]. The present study was done to evaluate the efficacy of short term intermittent chemotherapy in DOTS regimen in extra spinal tuberculosis.

MATERIALS AND METHODS

The present prospective study included 24 patients of extra spinal osteoarticular TB selected based on inclusion criteria. The study was done between October 2008 to June 2010 in the Department of Orthopedics, DDU Hospital, New Delhi. During the course of study 4 patients were lost to follow-up.

A Written informed consent from all the included patients and Ethical Committee approval was obtained before starting the study. Patients with extra spinal osteoarticular TB with immuno-compromised status such as HIV, cancer, severe, protein energy malnutrition,

diabetes or renal failure, age group of more than 70 years, defaulter and treatment failure cases, tuberculosis patients taking immunosuppressive drugs and cases of osteoarticular tuberculosis during pregnancy were excluded from the present study.

Laboratory investigation such as hemoglobin, chest X- ray PA view and C-reactive protein was performed. MRI of affected part was done in selective patients where above investigations were failed to diagnose the disease. In case, patient's condition was not improved by intensive treatment, suspicion of multiple drug resistance (MDR) was made and then additional culture and sensitive of aspirate was done for diagnosis and exclusion.

The DOTS regime recommended by WHO for extra spinal tuberculosis category III patients was followed i.e. 2(HRZE) 3 +4(HR) 3. Extension of the intensive phase was done for one month depending upon the response of clinical condition. Surgery of the affected part was considered if a lesion was not responding favorably to the ATT or as advocated in the "Middle Path Regime" or if there was any doubt in diagnosis. The surgical intervention was done as per the conventional indications.

Regular follow up was done at intervals of one month during the treatment for assessing the clinical improvement and compliance of the patient. The improvement was assessed with radiographs and haemogram with ESR and C-reactive protein at the interval of one month till the treatment was completed. The other clinical parameters such as weight gain, absence of other constitutional symptoms was also taken into account. After completion of treatment patients were called at interval of every three month and assessed to find local recurrence of osteoarticular tuberculosis. All the data were analyzed using IBM SPSS- ver.20 software. Analysis

was performed using chi-square test and independent sample student t test. P values <0.05 was considered to be significant.

RESULTS

In present study, median age was 24 years. Thirty percent of the patients belong to age group of 11-20 years. Students were mostly affected [8 (40%)] of total extra spinal cases followed by the labourer. At the time of presentation, all 20 (100%) patients had an elevated ESR, median value being 61. Skiagram of the affected part and MRI was done in 18 (90%) and 2 (10%) patients respectively and showed positive results.

Histopathologic results showed that, 2 (10%) were FNAC positive for AFB bacilli. Out of 7 patients who went for diagnostic biopsy, all have shown positive pathological picture of necrotizing granulomatous inflammation with Langhans giant cells. One patient had an active pulmonary disease. Among the patients of extra spinal osteoarticular TB hip (50%) was the most commonly involved joint. Short bones of hand and foot were involved in 20% cases. Other involved joints were knee (20%), shoulder (5%), ankle (5%) and elbow (5%). Long bone was involved in 5% cases.

Out of total 20 patients of EST, 16(80%) were given 6 months treatment of antitubercular drugs (ATT), 3 (15%) cases were given 7 months of ATT and 1 (5%) patients received 8 month treatment of ATT. At initial stage, 10 (50%) of extra spinal osteoarticular TB patients were CRP positive. Out of 20 extra spinal osteoarticular TB patients, 16 (80%) patients had shown increase in weight at the end of 6 month treatment. Patients' body weight improved with duration and good response seen in first 2 month of treatment. No relapse was observed during the course of study.

Table 1: Distribution of patients according to different parameters

Parameters		EST
Age	<40	17 (85)
	≥40	3 (15)
Gender	Male	17 (85)
	Female	3 (15)
Occupation	Students	8 (40)
	Housewife	2 (10)
	Manual worker	6 (30)
	Service	2 (10)
	Pre-school	2 (10)
Clinical feature	Pain	19 (95)
	Constitutional symptoms	17 (85)
	Palpable abscess	14 (70)
	Neurological deficit	0 (0)

Data is expressed as no of patients (%), ST; spinal tuberculosis, EST; extra spinal tuberculosis

Table 2: Trend of ESR, CRP and weight gain in extra spinal osteoarticular TB

Month			0	1	2	3	4	5	6	T-3	T-6	T-9	T-12
EST (n=20)	ESR *	M	64.5	53.8	34	32.9	27.5	12.9	20.5	15.9	14.9	14.9	14.5
	CRP	+ ve	10	0	0	0	0	1	0	0	0	0	0
	WG	M	45.8	46.2	47.7	46.4	46.1	47.2	46.5	47.6	47.7	49.5	50.5

*6 month of ATT, M; mean, N; no of patients, +ve; positive cases, ESR; erythrocyte sedimentation rate, CRP; C-reactive protein, WG; weight gain, EST; extra spinal tuberculosis, ST; spinal tuberculosis

DISCUSSION

WHO has also recommended the use of SCC in patients with EST in developing countries [2], Bhardwaj *et al.*; did a study of 25 newly diagnosed cases of EST and reported that all patients had pain and tenderness at the time of presentation [4]. Similar to Bhardwaj *et al.*; in present study too, almost all patients (95%) were presented with pain as a clinical symptom[4]. Hosalkar *et al.*; also reported almost similar findings [5].

In present study, 80% of patients had shown increase (mean change of 4.7 kg) in weight at the end of treatment. Almost similar to Bhardwaj *et al.*, they reported that all patients in their study gained weight from 40.3 Kg to 42.80 after three months and 45.50 kg after 6 months of therapy [4]. A study done by Wang *et al* also reported that during the course of therapy, mean values of ESR decreased from 52.03 to 11.33 mm in the first hour, hence indicating that there was a consistent betterment in ESR values [6]. In present study also all patients were found with elevated level of ESR.

Wang *et al.*; did a similar study on 185 patients and reported that patients with EST needed chemotherapy for longer period of time [6]. Effective use of DOTS therapy as recommended by WHO in patients with EST, assure the success of chemotherapy treatment, also it is the most cost effective treatment available [7, 8]. Effectiveness of DOTS therapy in pulmonary tuberculosis is established [9] but present study we found that regime was also effective for EST but the treatment required extended duration of chemotherapy. Similar findings were reported by Wang *et al.*; [6]. Hosalkar *et al.*; performed a retrospective review on EST cases and reported that lesions were distributed to different parts including ulna, scapula, distal humerus, distal tibia, proximal tibia and acetabulum [5]. But in present study most of the patients were having hip joint involvement along with short bones of hand and foot as reported by Hosalkar *et al.*; [5].

Short term intermittent chemotherapy was sufficient for 80% of extra spinal osteoarticular TB patients. Duration of treatment was extended in 20% cases for complete cure. From the present study data it is evident that short term intermittent chemotherapy (STIC) i.e. DOTS category-III efficacy was found

sufficient in (80%) EST, Parthasarathy *et al.*; in their study also recommended the use of short-course chemotherapy for the patients with EST [10]. The present study had few limitations like small sample size. Follow-up period was short. A prolonged follow-up of 5-10 years is essential to ascertain the actual rate of relapse.

CONCLUSION

DOTS treatment needs to be titrated depending on the clinical, lab finding and radiological evidence. As per our observation extra spinal tuberculosis patient should be given at least 8 months of DOTS treatment.

REFERENCES

1. Park K; Text Book of Preventive and Social Medicine; 18th Ed. 2005: 146-61.
2. Treatment of tuberculosis. Guidelines for National Programme, 3rd ed. 2003: 35 WHO.
3. Vaughn KD; Extra spinal osteoarticular tuberculosis. A forgotten entity? West Indian Med J 2005; 54(3): 202.
4. Bhardwaj V, Agrawal M, Suri T, Sural S, Kashyap R, Dhal A; Evaluation of adequacy of short-course chemotherapy for extra spinal osteoarticular tuberculosis using 99mTc ciprofloxacin scan. International Orthopaedics (SICOT) 2011; 35:1869-74.
5. Hosalkar HS, Agrawal N, Reddy S, Sehgal K, Fox EJ, Hill RA; Skeletal tuberculosis in children in the Western world: 18 new cases with a review of the literature. J Child Orthop 2009; 3:319-24.
6. Wang Z, Shi J, Geng G, Qiu H; Ultra short-course chemotherapy for spinal tuberculosis: five years of observation. Eur Spine J 2013; 22:274-81.
7. Hazra A, Laha B; Chemotherapy of osteoarticular tuberculosis. Indian J Pharmacol 2005; 37 (1): 5-12.
8. Teklali Y, El Alami ZF, El Madhi T, Gourinda H, Miri A; Peripheral osteoarticular tuberculosis in children: 106 case-reports. Joint Bone Spine 2003; 70:282-6.
9. Steffen R, Menzies D, Oxlade O; Patients' costs and cost-effectiveness of tuberculosis treatment in DOTS and non-DOTS facilities in Rio de Janeiro, Brazil. PLoS ONE 2010; 5(11): e14014.
10. Parthasarathy R, Sriram K, Santha T, Prabhakar R, Somasundaram PR, Sivasubramanian S; Short-course chemotherapy for tuberculosis of the spine. J Bone Joint Surg 1999; 81-B: 464-71.