Prevalence of HCV Infection in Children with Arthritis

Thesis

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Abstract:

Background/Aim: Hepatitis C virus (HCV) infection can induce immunological extrahepatic manifestations including arthralgia and arthritis mimicing rheumatoid arthritis. Our aim is to report the prevalence of HCV infection in children presenting with arthritis or arthralgia.

Methods: The study included 40 children presenting with arthritis or arthralgia as a cases and another 40 age and sex matched children, not complaining of liver disease or arthritis as a control group. All cases and controls were subjected to full history taking, clinical examination and HCVab detection by rapid test. To all positive cases with rapid test a confirmatory HCVab ELISA test was done and HCV PCR as well.

Results: Cases were 45% males and 55% females, with a mean age of $10.5 \pm$ 3.7 years. Among cases, prevalence of HCVab as detected by rapid test and confirmed **ELISA** 5%. **HCV** RNA by was was positive in 2.5%. In the control group HCVab was 2.5% by rapid test and confirmed by ELISA and HCV RNA. Both arthritis and lower limb oedema were more prevalent in HCVab positive cases than negative but with no statistical significance difference. By examination; HCVab positive cases had a higher prevalence of presence of scratch marks and they also have a significantly higher AST and ALT levels than HCVab negative cases.

Conclusion: HCV infection is prevalent in children with arthritis or arthralgia.

Key words: HCV, arthritis, arthralgia, HCVab, HCV RNA, rapid test, children.

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List of Abbreviations

AA Amyloid A

ALT Alanine transaminase

AKA Anti Keratin Antibody

AST Aspartae transaminase

ANA Anti Nuclear Antibody

APF Anti Perinuclear Factor

APR Acute Phase Reactant

BMC Bone Mineral Content

BILAG British Isles Lupus Assessment Group

CRP C Reactive Protein

COMP Cartilage Oligometric Matrix Protein

CTLs Cytotoxic T Lymphocytes

CRF Chronic Renal Failure

CLA chemilminescence immunoassay

CNS Central Nervous System

CVD Coronary Vascular Disease

CD2BP1 CD-2 binding protein-1

CYC Cyclophosphamide

DEXA Dual Energy X-ray Absorptiometry

ESR Erythrocyte Sedimentation Rate

EDHS Egyptian Demographic Health Survey

ELISA Enzyme Linked Immunosorbont Assay

EVR End Viral Response

FMF Familial Mediterranean Fever

GN Glomureolo Nephritis

Hp Haptoglobin

HBV Hepatitis B Virus

HCV Hepatitis C Virus

HCVab Hepatitis C Virus antibody

HIV Human Immunodeficiency Virus

IDDM Insulin Dependent Diabetes Mellitus

IASL International Association for the Study of The Liver

IFN-α Interferon-α

IGg Immunoglobulin g

IGM Immunoglobulin M

ILD Interstitial Lung Disease

ITP Idiopathic Thrombocytopenia

JRA Juvenile Rheumatoid Arthritis

JIA Juvenile Idiopathic Arthritis

LEF leflunomide

LLN Lower Limit of Normal

MEFV Mediterranean Fever

MHC Major Histocompatibilty

MMF Mycophenolate mofetil

MRI Magnetic Resonance Image

MTX Methotrexate

NAFLD Non alcoholic Fatty Liver Disease

NHANES national health and nutrition examination survey

NP Neuropsychiatric

NSAIDs Non Steroidal Anti Inflammatory Drugs

Oligo JIA Oligoarticular juvenile idiopathic arthritis

PAPA pyogenic arthritis, pyoderma gangrenosum, and acne

PCR Polymerase Chain Reaction

PEG Polyethylene glycol

pSLE Pediatric Systemic Lupus Erythematosus

PSTPIP1 proline serine threonine phospatase-interacting protein 1

Poly JIA Polyarticular juvenile idiopathic arthritis

RA Rheumatoid Arthritis

RFs Rheumatoid Factors

RIBA Recombinant Immunoblot Assay

RVR Rapid Viral Response

SAARDs	slow-acting antirheumatic drugs
S-JIA	Systemic juvenile idiopathic arthritis
SLE	Systemic Lupus Erythematosus
SVR	Sustained Viral Responce
TMA	Transcription mediated amplification
TNF	Tumor Necrosis Factor

Introduction:

Hepatitis C virus (HCV) is an RNA flavivirus with six major genotypes and several subtypes, HCV is an important causative agent of liver diseases. However, HCV infection is more than just a liver disease and has been associated with numerous hematologic, renal, dermatologic, rheumatic and autoimmune disorders (Cacoub et al., 2000, Ramos-Casals and Font, 2005).

Recent studies have reported that more than half (54%) of the HCV infected Egyptian children had persistently elevated ALT levels (**El-Raziky et al., 2004**). The worldwide seroprevalence of HCV ranges between 0.2-2% (**Wilber, 1995**). In 2007, an HCV seroprevalence of 2% was reported in healthy Egyptian children (**El-Raziky et al., 2007**).

HCV infection is usually asymptomatic and may be detected incidentally; it also induces immunological extrahepatic manifestations including arthralgia and arthritis which can mimic rheumatoid arthritis (RA). HCV is associated with a variety of rheumatic disorders and autoimmune phenomena. This includes arthralgia, arthritis, vasculitis, sicca syndrome, myalgia and fibromyalgia (Palazzi et al., 2008). Arthralgia is the most common rheumatic manifestation. Arthritis is less common and may present as a rheumatoid arthritis-like, non deforming arthritis mainly involving small joints with RF and absence of rheumatoid nodules and less commonly a mono-oligoarthritis usually of large joints (Sterling and Bralow, 2006).

Associations have been reported between HCV infection and other autoimmune diseases including systemic lupus erythematosus, Sjogren's syndrome and antiphospholipid antibody syndrome (APS) (Lormeau et al., 2006; Ramos-Casals and Font, 2005).

Rosner and colleagues in 2004 summarized the clinical characteristics of HCV associated arthritis. Arthritis, not otherwise explained, has been noted in 2% to 20% of HCV patients, that arthritis was rheumatoid-like in two thirds of the cases and a waxing waning oligoarthritis in the rest. It was concluded that HCV arthropathy should be considered in the differential diagnosis of new onset arthritis

The pathogenesis of HCV-related arthritis is not entirely clear, but three possible mechanisms have been suggested, namely: synovial tissue damage by direct viral invasion, synovial autoimmune response induced by the virus and immune complexes or cryoglobulins deposition (**Zuckerman et al., 2001**).

Aim of work:
The aim of this study is to report the prevalence of hepatitis C virus infection in children and adolescence with arthritis or arthralgia.