Viral infections among volunteer blood donors in the national blood transfusion centre in Egypt

Thesis

Submitted for partial fulfillment of master degree in Clinical Pathology

By

Hossam El-Din Fikry Ahmed Sabaa

M. B. B. Ch Faculty of medicine Cairo University

Supervised by

Prof.Dr.: Naglaa Gamal Mady Abo-El-azaem

Professor of clinical pathology

Faculty of medicine, Al- Azhar University

Prof.Dr.: Wael Refaat Abd-El-Hamid Hablas

Professor of clinical pathology

Faculty of medicine, Al- Azhar University

Dr.: Abd-El-Raouf Abd-El-Raouf Abo-Nar

Lecturer of clinical pathology

Faculty of medicine, Al- Azhar University

Faculty of medicine

Al- Azhar University

2013

بسم الله الرحمن الرحيم

(و علمك ما لم تكن تعلم و كان فضل الله عليك عظيما)

صدق الله العظيم

(النساء-١١٣)

ACKNOWLEDGMENT

Before all, thanks to "ALLAH" who granted me the ability to perform the work of this thesis.

I wish to introduce my respect and thanks to **Prof.Dr. Naglaa Gamal Mady Abo-El-azaem**, Professor of clinical pathology, Faculty of medicine, AlAzhar University, for her great support and help to complete this work.

I would like to thanks **Prof.Dr. Wael Refaat Abd-El-Hamid Hablas,** Professor of clinical pathology, Faculty of medicine, Al- Azhar University, for his great support and help to complete this work.

I would like to thanks **Dr. Abd-El-Raouf Abd-El-Raouf Abo-Nar**, lecturer of clinical pathology, Faculty of medicine, Al- Azhar University, for his continuous encouragement and valuable expert guidance and unlimited support during the entire course of the study.

LIST OF CONTENTS

Title	Page no.
I INTRODUCTION	
III REVIEW OF LITERATURE	4
1.Blood donation	4
2.Transfusion Transmitted viral infections	
IV SUBJECTS AND METHODS	66
V RESULTS	79
VI DISCUSSION	91
VII SUMMARY	98
VIII CONCLUSION AND RECOMMENDATION	100
IX REFERENCES	101
X ARABIC SUMMARY	

LIST OF TABLES

Table no.	Title	Page no.
1	Interpretation of serologic and molecular markers of hepatiti	is B virus
	during different stages of infection	29
2	Mode of transmission of HIV	41
3	Gender of the studied cases	82
4	Age of the studied cases	82
5	Weight of the studied cases	82
6	Residence of the studied cases	82
7	ELISA and NAT results	83
8	ELISA and NAT results in HCV donors	83
9	Relation between ELISA and NAT results in HCV screening	83
10	Age, sex, weight and residence among the HCV cases	84
11	Residence and sex among the positive HCV cases	84
12	Relation between ELISA and NAT results in HBV +ve cases	84
13	Age, sex, weight and residence among the HBV cases	85
14	The mean age of HCV positive cases and HBV positive cases.	86

LIST OF FIGURES

Figure no.	Title	Page no.
1	The genome organisation of HBV. The genes overlap	20
2	HIV structure	44
3	PROCLEIX TIGRIS System	74
4	The Centrifugation device	74
5	The Microplate (96 well plate)	74
6	The Microplate reader	75
7	The incubator	75
8	The Microplate washer	75
9	Reagent preparing incubator	76
10	Multiple units (MTUs)	76
11	Disposable Tips (DiTi)	76
12	A-T binding	77
13	Transcripted mediated amplification	77
14	Amplicons	78
15	Hybridization protection assay	78
16	The light signals	78
17	Male/Female ratio	86
18	Age of cases	86
19	Weight of cases	87
20	Residence of cases	87
21	ELISA results	88
22	NAT results	88
23	ELISA and NAT results	89
24	Male/Female ratio of HCV cases	89

Residence of HCV cases	.90
Male/Female ratio of HBV cases	.90
Residence of HBV cases	.90

LIST OF ABBREVIATIONS

AE Acridinium ester

AIDS Acquired immunodeficiency syndrome

anti-HBc IgM IgM antibodies to the hepatitis B core antigen

cccDNA Covalently closed circular Deoxyribonucleic acid

CDC Centers for Disease Control and Prevention

cDNA Complementary DNA

CMV Cytomegalovirus

DNA Deoxyribonucleic acid

EIAs Enzyme immunoassay

ELISA Enzyme-linked immunosorbent assay

HAART High active antiretroviral therapy

HBV Hepatitis B virus

HBcAg Hepatitis B core antigen

HBe Ab (anti-HBe) Hepatitis B envelope antibody

HBeAg Hepatitis B envelope antigen

HBIG Hepatitis B immunoglobulin

HBs Ab (anti-HBs)

Antibody to the hepatitis B surface antigen

HBsAg Hepatitis B surface antigen

HCV Hepatitis C virus

HDV Hepatitis D virus

HIV Human immunodeficiency virus

HPA Hybridization Protection Assay

HTLV Human T-Lymphotropic virus

IDT Individual donor testing

IDU Intravenous drug use

MGN Membranous glomerulonephritis

mRNAs Messenger ribonucleic acid

MTCT Vertical Transmission from mother to child

MTUs Loading Multi-Tube Units

NAT Nucleic Acid Testing

NBTC National blood transfusion centre

OBIs Occult HBV infections

OD Optical density

PCR Polymerase chain reaction

RNA Ribonucleic acid

STDs Sexually transmitted diseases

TCR Target Capture Reagent

TMA Transcription mediated amplification

TTIs Transfusion transmissible infections

VNRD Voluntary non remunerated repeat donations

WHO World Health Organization

WNV West Nile virus

I.INTRODUCTION

Blood transfusion is one of the life saving interventions but carries the risk of acute and delayed complications such as transfusion transmissible infections (TTIs). Blood donors are the cornerstone of a safe and adequate supply of blood and blood products and the safest blood donors are voluntary, non remunerated blood donors from low risk populations (**Anjali et al., 2012**).

Hepatitis B is one of most common infectious diseases of the world infecting two billion people including an estimated 400 million chronically infected cases. Individuals with chronic infection have a high risk of developing liver cirrhosis and hepatocellular carcinoma. Hepatitis C virus infection is another common chronic blood borne infection with an estimated 3.9 million persons infected with the virus and it has a high rate of development of liver cirrhosis. Infection by Hepatitis B virus (HBV) and Hepatitis C virus (HCV) cause serious mortality, morbidity and financial burden and are thus a major global health problem, so predonation screening of blood donors or screening of donated blood for HBV and HCV are thus a routine practice (Nkrumah et al., 2011).

Although blood transfusion safety has greatly improved over the last 15 years, the transfusion risk of HBV remains high in developing countries. The development of quality practice for blood donation based on the use of the most sensitive techniques for the detection of infectious risk of blood donation should be a priority of health authorities (**Noah et al., 2011**).

The current use of nucleic acid testing for detection of the human immunodeficiency virus (HIV) and hepatitis C virus (HCV) RNA and HBV DNA in a single triplex assay may provide additional safety (**Stramer et al., 2011**).

The availability of hepatitis B virus (HBV) nucleic acid testing (NAT) for donor blood screening led to its implementation in low prevalence and high prevalence countries. Genomic detection was a substantial addition to HBV surface protein (HBsAg) screening by detecting window period infections and occult HBV infections (OBIs) which is characterized by undetectable HBsAg, low viral load and presence of serological markers (anti-HBc and/or anti-HBs). OBIs are the result of multiple, poorly understood mechanisms including incomplete immune control mutations of the HBsAg antigenic determinants, abnormal expression of S gene, and inhibition of genome transcription. Infectivity for the recipient is high for window period blood and relatively low for OBIs (Williams & Wilkins., 2011).

II.AIM OF THE STUDY

This study aims to screen HCV, HBV, and HIV in blood donors in the national blood transfusion centre (NBTC) in Egypt for detection of prevalence of viral infections among volunteer blood donors.

III.REVIEW OF LITERATURE

1.Blood donation

A blood donation occurs when a person voluntarily has blood drawn and used for transfusions or made into medications by a process called fractionation (Goldman et al., 2007).

In the developed world, most blood donors are unpaid volunteers (voluntary non remunerated repeat donations, VNRD) who donate blood for a community supply. In the poorer countries, most blood donors are directed donors. Many donors donate as an act of charity, but some are paid and in some cases there are incentives other than money such as paid time off from work. Donation is relatively safe, but some donors have bruising where the needle is inserted or may feel faint (Goldman et al., 2007).

Potential donors are evaluated for anything that might make their blood unsafe to use. The screening includes testing for diseases that can be transmitted by the blood transfusion including HIV and viral hepatitis. The donor must also answer questions about medical history and take a short physical examination to make sure the donation is not hazardous to his or her health. How often a donor can give varies from days to months based on what he or she donates and the laws of the country where the donation takes place. For example in the United States, donors must wait eight weeks (56 days) between whole blood donations but only three days between platelet pheresis donations (**Riley et al., 2007**).

Types of blood donation

Blood donations are divided into groups based on who will receive the collected blood. An allogeneic (also called homologous) donation is when a donor gives blood for storage at a blood bank for transfusion to an unknown recipient. A directed donation is when a person, often a family member, donates blood for transfusion to a specific individual. Directed donations are relatively rare when an established supply exists. A replacement donation is a hybrid of the two and is common in developing countries such as Ghana. In this case, a friend or family member of the recipient donates blood to replace the stored blood used in a transfusion, ensuring a consistent supply. When a person has blood stored that will be transfused back to the donor at a later date, usually after surgery, that is called an autologous donation. Blood that is used to make medications can be made from allogeneic donations or from donations exclusively used for manufacturing (Riley et al., 2007).

Blood is sometimes collected using similar methods for therapeutic phlebotomy, which is used to treat conditions such as hereditary hemochromatosis or polycythemia vera. This blood is sometimes treated as a blood donation, but may be immediately discarded if it cannot be used for transfusion or further manufacturing (Goldman et al., 2007).

The actual process varies according to the laws of the country, and recommendations to donors vary according to the collecting organization. The World Health Organization gives recommendations for blood donation policies, but in developing countries many of these are not followed. For example, the recommended testing requires laboratory facilities, trained staff, and specialized

reagents, all of which may not be available or too expensive in developing countries (WHO., 2008).

An event where donors come to donate allogeneic blood is sometimes called a blood drive or a blood donor session. These can occur at a blood bank, but they are often set up at a location in the community such as a shopping center, work place, or school (Gómez-Simón et al., 2007).

Donor safety

The donor is also examined and asked specific questions about their medical history to make sure that donating blood is not hazardous to their health. The donor's hematocrit or hemoglobin level is tested to make sure that the loss of blood will not make them anemic and this check is the most common reason that a donor is ineligible (**Gómez-Simón et al., 2007**).

Pulse, blood pressure and body temperature are evaluated before donation. Elderly donors are sometimes also deferred on age alone because of health concerns. The safety of donating blood during pregnancy has not been studied thoroughly and pregnant women are usually deferred (**Goldman et al., 2007**).

Methods of blood donation

There are two main methods of obtaining blood from a donor. The most frequent is to simply take the blood from a vein as whole blood. This blood is typically separated into parts, usually red blood cells and plasma, since most recipients need only a specific component for transfusions. A typical donation is 450 millilitres of whole blood, though 500 millilitre donations are also common. The blood is usually stored in a flexible plastic bag that also contains sodium