# Effect of Omega 3 Fatty Acids Supplementation on Patency of Arteriovenous Access in Hemodialysis Patients

Thesis
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Presented By:

Ahmad Mohamed Alsawy (MB.B.Ch)

Under Supervision of

# Prof. Dr. Howayda Abd Elhameed Elshinnawy

Professor of Internal Medicine and Nephrology Faculty of Medicine Ain Shams University

#### Dr. Walid Ahmed Bichari

Lecturer of Internal Medicine and Nephrology Faculty of Medicine Ain Shams University

> Faculty of Medicine Ain Shams University 2014

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#### **List of Abbreviations**

AA ..... Arachidonic Acid **ACE**......Angiotensin-Converting Enzyme **ACR**.....Albumin to Creatinine Ratio AHA......American Heart Association **AKI**.....Acute Kidney Injury ALA ..... a-Linolenic Acid ARBs ......Angiotensin Receptor Blockers AusDiab......Australian Diabetes AV ......Arteriovenus AVF ...... Arteriovenous Fistula AVG......Arteriovenous Graft CARI.....The Australian Caring for Australasians with Renal Impairment CHD..... Coronary Heart Disease **CKD**......Chronic kidney Disease **COX**.....Cyclooxygenase **Cr** ...... Creatinine CS ...... Cross-Sectional **CSN**......Canadian Society of Nephrology CVD......Cardiovascular Disease DHA......Docosahexanoic Acid **DOQI**.....The National Kidney Fondation Dialysis **Outcomes Quality Initiatives ECM** ..... Extracellular Matrix eGFR ..... Estimated Glomerular Filtration Rate **EPA** ..... Eicosapentanoic Acid **EPIC**.....European Prospective Investigation into

Cancer Study

#### **List of Abbreviations (Cont.)**

**EPO** ..... Erythropoietin ePTFE ..... Expanded Polytetrafluoroethylene **ESRD**..... End Stage Renal Disease **F**......Female **F**A.....Fatty Acid **GFR**.....Glomerular Filtration Rate GISSI......Gruppo Italiano per lo Studio della Sopravvivenza nell'Infarto miocardico **HDL**.....High Density Lipoprotein INR......International Normalised Ratio K/DOQI.....The Kidney Disease Outcomes Quality **Initiative** KDIGO......The Kidney Disease: Improving Global **Outcomes** L.....Longitudinal **LDL** .....Low Density Lipoprotein **LOX**.....Lipooxygenase LT.....Leukotriene **M**......Male MA..... Microalbuminuria MDRD..... The Modification of Diet in Renal Disease N......Number of Participants n-3 PUFAs ...... Omega 3 Polyunsaturated Fatty Acids **NEJM** ...... The New England Journal of Medicine **NEOERICA** ...... New Opportunities for Early Renal Intervention by Computerised Assessment NFkB ...... Nuclear Factor Kappa Light Chain NHANES ...... National Health and Nutrition Evaluation Survey

#### **List of Abbreviations (Cont.)**

NICE......National Institute of Health and Clinical Excellence **NP** ......Nonpredictive **NSAIDs**......Non Steroidal Anti-Inflammatory Drugs **PDGF**.....Platelet-Derived Growth Factor **PET** ......Polyethylene Terephthalate **PKD**.....Polycystic Kidney Disease PR ..... Predictive **PREVEND**.....Prevention of Vascular Renal and **Endstage Disease** PTFE .....Polytetrafluoroethylene **PTH**.....Parathyroid Hormone **RAAS**.....The Renin-Angiotensin-Aldosterone System RCM ...... Radiocontrast Material RCT.....Randomized Controlled Trial **RR**.....Relative Risk RRT.....Renal Replacement Therapy **T3**.....Tri Iodothyronine **T4**......Thyroxine The pl-TFE.....The Plasma Tetrafluoroethylene **TxA**.....Thromboxane U.K..... The United Kingdom US ......United State

**USRDS**.....The United States Renal Data System

#### Introduction

Maintaining vascular access in hemodialysis patient is considered a challenge since the portal is vulnerable to infection, stenosis, and thrombosis. Vascular access options for hemodialysis patients include the placement of arteriovenous fistulas, arteriovenous grafts, and double lumen, cuffed central venous catheters (*Frederick 2010*).

Catheter use is generally associated with higher rates of infection and could compromise the adequacy of hemodialysis (*Quarello et al.*, 2006). Other risk factors linked to catheter use include increased thrombosis, unreliable blood flows, central venous stenosis and patient cosmetic concern (*Quarello et al.*, 2006).

In United States, 55% of patients are allocated towards AV fistula, 21% towards AV grafts and 24% double lumen dialysis catheter (*Charmaine et al.*, 2012).

Thrombosis occurs in more than 50% of all arteriovenous grafts within 1 year after placement, necessitating a salvage procedure in more than 75% (*Charmaine et al.*, 2012).

Risk factors for access failure include increased age, female gender, hypertension, diabetes mellitus, and positive HIV status (*Schild 2004*).

Various studies have attempted to discover a pharmacological approach to minimize vascular access failure induced by thrombosis most of them were equivocal and needed further time to reassess (*Frederick 2010*).

Novel strategies to prevent dialysis access thrombosis are needed to reduce the cost and morbidity of maintenance hemodialysis. Diets enriched with  $\omega$ -3 fatty acids, derived from fish oil, may offer such an opportunity. Such diets may favorably impact the vascular perturbations that could contribute to synthetic graft thrombosis.

**Lok 2007,** in his study reported the value of  $\omega$ -3 fatty acids in protecting the vascular access against thrombosis.

Other pharmacological agents such as warfarin were deemed ineffective and the trial was prematurely terminated due to increased bleeding episodes in hemodialysis patients (*Crowther 2002*).

## Aim of the Work

Our study is a prospective study aiming to evaluate the role of  $\omega$ -3 fatty acids in preserving the patency of arteriovascular fistulas and grafts in hemodialysis patients.

#### Chapter (I): Epidemiology and Pathophysiology of Chronic Kidney Disease

#### **Definition**

Chronic kidney disease (CKD) is defined as kidney damage or glomerular filtration rate (GFR) below 60 ml/min per 1.73 m2 for 3 months or more irrespective of the cause. The Kidney Disease Outcomes Quality Initiative (KDOQI) guidelines have classified CKD into five stages (*K/DOQI*, 2002).

This classification, although useful in simplifying the categorization of CKD, has its limitations, which include classifying people with isolated microalbuminuria as suffering from CKD, labeling mild and stable kidney damage as CKD, and not differentiating between age-related impaired kidney function and progressive disease-induced CKD (*Glassock and Winearls*, 2008).

In 2005, the Kidney Disease Improving Global Outcomes (KDIGO) group suggested clarifications including the addition of the suffix T for patients with renal allografts and D to identify CKD stage 5 patients on dialysis (*Levey et al.*, 2007).

The U.K. National Institute of Health and Clinical Excellence (NICE) has modified, in 2008, the KDOQI CKD classification by subdividing CKD stage 3 into 3A and 3B,