

بسم الله الرحمن الرحيم





HOSSAM MAGHRABY





شبكة المعلومات الجامعية التوثيق الالكتروني والميكروفيلم



HOSSAM MAGHRABY



جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم قسم

نقسم بالله العظيم أن المادة التي تم توثيقها وتسجيلها علي هذه الأقراص المدمجة قد أعدت دون أية تغيرات



يجب أن

تحفظ هذه الأقراص المدمجة بعيدا عن الغيار



HOSSAM MAGHRABY



Analysis of Outcome in Recipients with Different Graft-Recipient-Weight-Ratio (GRWR) Post Adult to Adult Living Donor Liver Transplant (LDLT)

Thesis

Submitted for Partial Fulfillment of Master Degree in General Surgery

By

Marolla Maher Iskander

(M B BcH) Cairo University

Under supervision of

Prof. Dr. Amr Ahmed Abd ElAal

Professor of General Surgery Faculty of Medicine, Ain Shams University

Prof. Dr. Ahmed AbdFlRazek

Assistant Professor of General Surgery Faculty of Medicine, Ain Shams University

Dr. Mahmoud Talaat

Lecturer of General Surgery Faculty of Medicine, Ain Shams University

> Faculty of Medicine Ain Shams University 2021

Acknowledgment

First and foremost, I feel always indebted to **God**, the Most Kind and Most Merciful.

I'd like to express my respectful thanks and profound gratitude to **Prof. Dr. Amr Ahmed Abd ElAal,** Professor of General Surgery, Faculty of Medicine, Ain Shams University for his keen guidance, kind supervision, valuable advice and continuous encouragement, which made possible the completion of this work.

I am also delighted to express my deepest gratitude and thanks to **Prof. Dr. Ahmed AbdElRazek**, Assistant Professor of General Surgery, Faculty of Medicine, Ain Shams University, for his kind care, continuous supervision, valuable instructions, constant help and great assistance throughout this work.

I am deeply thankful to **Dr. Mahmoud Talaat,** Lecturer of General Surgery, Faculty of Medicine, Ain Shams University, for his great help, active participation and guidance.

Marolla Maher

List of Contents

Title	Page No.
List of Tables	i
List of Figures	iii
List of Abbreviations	vi
Introduction	1
Aim of the Work	5
Review of Literature	
Anatomy	6
Liver Transplantation	21
Preoperative Assessment of the Recepients an Selection	
Post Operative Care and LT Complications	
Small For Size Syndrome	
Patients and Methods	
Results	
Discussion	111
Summery	121
Conclusion	
References	125
Arabic Summary	

List of Tables

Table No.	Title	Page No.
Table (1): Table (2): Table (3):	Indications for liver transplant Show MELD exceptions Show the preoperative workup patients pre LDLTx	30 for
Table (4):	Evaluation protocol for potential liveliver donors at the University of Ess Germany	ring sen,
Table (5):	The definition of small-for-syndrome* reported by various author	
Table (6):	Show the causes of SFSS and strategies to prevent SFSS	
Table (7):	Descriptive for demographic data of recipient.	
Table (8):	Showing intraoperative details of recipient.	
Table (9):	Comparison between patients w GRWR ≤ 0.8 and those with GRWR > regarding donors demographic data	0.8
Table (10):	Comparison between patients w GRWR ≤ 0.8 and those with GRWR > regarding recipient demographic d and hepatpathy	-0.8 lata
Table (11):	Comparison between patients w GRWR ≤ 0.8 and those with GRWR > regarding recipient preoperar assessment	-0.8 tive
Table (12):	Comparison between patients w GRWR ≤ 0.8 and those with GRWR > regarding Intra-operative assessment.	0.8

List of Tables Cont...

Table No.	Title	Page No.
Table (13):	Showing Comparison between patiewith GRWR ≤0.8 &>0.8 regarding suffer size syndrome manifestatic incidence and early post operations.	mall ons, tive
Table (14):	Comparison between cases with GR ≤ 0.8 and those with GRWR regarding overall survival (months)	>0.8
Table (15):	Showing comparison betw demographic data of Donors For patie who developed small for size and the who didn't	ents hose
Table (16):	Showing comparison betw demographic data & different characteristics of patient with SFSS those without	rent and
Table (17):	Comparing between patients who small for size syndrome & those didn't have with regard to S manifestations	who FSS
Table (18):	Showing comparison between patients with small for size syndrome & the without regarding intraopera assessment	nose ative
Table (19):	Comparing between patients who small for size syndrome & those didn't have with regard to S manifestations	who FSS
Table (20):	Comparison between the two students regarding overall surve (months)	died vival

List of Figures

Fig. No.	Title	Page No.
Figure (1):	Couinaud 8-segment (I–VIII) and division of the liver.(N).	
Figure (2):	Common variations of h	
Figure (3):	Maximal intensity projection view portal vein bifurcation	
Figure (4):	Ct scan showing the common open all three hepatic viens and in accessory hepatic vien(N)	nferior
Figure (5):	Hjortsjo's crook	18
Figure (6):	Extra hepatic and intra hepatic be system	•
Figure (7):	Anatomical variations of the hepatic biliary system	
Figure (8):	Three dimensions reconstruing of the liver and its vascular	
Figure (9):	Intraoperative cholangiogram resection to assure adequate lengthe bile duct with good stump	gth of
Figure (10):	Liver after completing f parenchymal resection	
Figure (11):	Irrigation of the graft with HTK so and assesmt of the veins t anastomsed.	o be
Figure (12):	Reconstruction of V5 using nature graft of the explanted liver on the table	e back
Figure (13):	The explanted liver	74

List of Figures Cont...

Fig. No.	Title	Page No.
Figure (14):	Rt lobe graft showing reconstruct. V5 into the confluence of MHV and of the recipient using synthetic graf	l LHV
Figure (15):	The left lobe graft	76
Figure (16):	Hepatopathy types of recipient grou	ıp81
Figure (17):	CHILD grade among recipient grou	p81
Figure (18):	The percentage of GRWR $\leq 0.8 \& >$	0.884
Figure (19):	Showing different types of grafts us	ed84
Figure (20):	Comparison between patients GRWR ≤0.8 &>0.8 regarding diff types of grafts	ferent
Figure (21):	Comparison between patients GRWR ≤0.8 &>0.8 regarding reconstruction.	MHV
Figure (22):	Comparison between patients GRWR ≤0.8 &>0.8 regarding early operative mortality	y post
Figure (23):	Overall survival for all the st patients	
Figure (24):	Kaplan–Meier curves for graft su in the first year after living donor transplantation for the 2 group patients	· liver ps of
Figure (25):	Comparison between patients with for size syndrome & those who have with regard to autoim hepatitis	didn't mune

List of Figures Cont...

Fig. No.	Title	Page No.
Figure (26):	Comparison between patients who small for size syndrome & those didn't have with regard to resistive of hepatic artery.	who index
Figure (27):	Comparison between patients who small for size syndrome & those didn't have with regard to GRWR	who
Figure (28):	Comparison between patients with and those with no SFSS with regardly of INR normalization	ard to
Figure (29):	Comparison between patients with and those with no SFSS with regarday of normalization of bilirubin	ard to
Figure (30):	Comparison between patients with with and those with no SFSS regard to day of normalization platelets.	with on of
Figure (31):	Comparison between patients with with and those with no SFSS regard to manifestations of SfSS	with
Figure (32):	Kaplan–Meier curves regarding or survival (in months) between the groups	e two

List of Abbreviations

Abb.	Full term
ATP	. Adenosine Triphosphate
	. Body mass index
	. Blood urea nitrogen
	. Complete Blood Count
	. Cholangiocarcinoma
	. Cytomegalovirus
	. C-reactive protein
	. Computed tomography
DDLT	. Deceased donor liver transplantation
EBV	. Epstein-Barr virus
ECG	. Electrocardiography
ESR	. Erythrocyte sedimentation rate
GIM	Graft Inflow Modulation
GRWR	. Graft to recipient weight ratio
GV/SLV	Graft volume standard liver volume ratio
HABR	. Hepatic arterial buffer response
HAT	. Hepatic artery thrombosis
HCC	. Hepatocellular carcinoma
HCV	. Hepatitis C virus
HGF	. Hepatocyte growth factor
HGFA	. Hepatocyte growth factor activator
HIV	Human immunodeficiency virus
HPCS	Hemi-portocaval shunt
HTK	Histidine-tryptophan-ketoglutarate solution
IHPBA	International hepato-pancreaticobiliary association
INR	. International normalized ratio
IVC	. Inferior vena cava
LDLT	. Living donor liver transplantation

List of Abbreviations Cont...

Abb.	Full term
L/Tx	. Liver transplantation
	. Model of end-stage liver disease
	. Middle hepatic vein
	. Middle hepatic vein
	Overwhelming post-splenectomy sepsis
	Primary non-function
PSS	. Porto-systemic shunts
	. Prothrombin time
PTT	. Partial thromboplastin time
PVF	. Portal venous flow
PVP	. Portal vein pressure
PVT	. Portal vein thrombosis
RPR	. Rapid plasma reagin
SAL	. Splenic artery ligation
SFSG	. Small-for-size graft
SFSS	. Small-for-size syndrome
SMV	. Splanchnic venous inflow
VEGF	. Vascular endothelial growth factor

ABSTRACT

Background: The problem of graft size is one of the critical factors limiting the expansion of adult-to-adult living donor liver transplantation (LDLT). Graft-to-recipient weight ratio (GRWR) > 0.8% is perceived as the critical graft size to meet the metabolic demand of the recipient. Small-for-size graft (SFSG) is the graft with GRWR < 0.8 and when its unable to meet the recipients metabolic demands, small-for-size syndrome (SFSS) occurs which is a life-threatening condition with rapidly progressive liver failure. This lower limit of GRWR (0.8%) has been challenged over the last decade perhaps due to better understanding of the pathophysiology, coupled with technical refinements, particularly related to venous outflow reconstruction.

Aim of Work: Our aim was to analyse the different outcomes of patients undergoing Living donor liver transplantation using grafts with $GRWR \le 0.8\%$ with those >0.8% to evaluate the factors that affect the post-operative outcome and the overall one year Graft survival.

Methods: This is a retrospective cohort study on 100 adult patients with ESLD or HCC (mean age 54.16 ± 10.53 years old) who underwent adult LDLT in the period between 2018 and 2020 and follow up the patients for one year post transplant, Patients were divided into two groups, patients who received Graft with GRWR ≤ 0.8 (N= 22 patients with lower limit of GRWR = 0.58 and there is 7 patients with GRWR ≤ 0.7) and those with GRWR >0.8 (N= 78 patients). We compared the the Donor factors, preoperative patient factors, intraoperative factors, development of small for size syndrome and graft survival in patients received small-for-size grafts (GRWR ≤ 0.8) with patients received GRWR <0.8.

Results: We retrospectively evaluated the donor factors, recipient factors and operative factors through the medical records. Small-for-size syndrome (SFSS) occurred in 2 of 22 patients (9.1%) in patients with GRWR \leq 0.8 and in 1 of 78 patients (1.3%) in patients with GRWR >0.8 which was statistically insignificant between two groups. There was No significant difference in the Donor age, Preoperative MELD, CHILD, Portal hypertension, cold and warm ischemia, operative time, presence of PVT and HCC in patients who received GRWR \leq 0.8 and patients with GRWR >0.8. the mean overall survival for all the studied cases was found 43.35 months and the overall survival at 6 months and 1 year was 96.0% but there was statistically significant increase in the overall survival of cases with GRWR >0.8 than those with GRWR \leq 0.8 with p-value = 0.009, as the overall 6 months and 1 year survival for patients with GRWR >0.8 was 98% and 98% compared to 86% and 86% in thoses with GRWR \leq 0.8

Conclusion: There is no difference in the outcome in form of development of Small for size syndrome when we use Grafts with GRWR >0.8 or with GRWR ≤ 0.8 , However the venous Outflow of SFSG (GRWR < 0.8) is very critical to maintain good graft function in the recipient.

Keywords: Living donor liver transplantation, graft-to-recipient weight ratio



Introduction

gypt is a heavily populated country, with a strikingly high prevelance of hepatitis C virus (HCV) infection, 26%. That has led to increasing numbers of Egyptian patients suffering from end-stage liver disease (El-Elemi and El-Gazzaz, 2010). Thus, liver diseases are amongst the national health problems that have a great impact on health insurance programs, national and financial resources. Hence, the remarkable continuous growth of liver transplantation programs and the increase of the number of transplanted patients (Sholkamy, 2014).

Indications for liver transplantation can be classified into end-stage liver disease, acute liver failure and certain benign and malignant liver tumors. LT should be considered for any patient in whom anticipated overall survival exceeds life expectancy of the underlying disease or where significant increase in quality of life can be achieved (Hackl et al., 2014).

Living donor liver transplantation (LDLT) has been a well-recognized alternative to whole graft transplantation from deceased donor in face of organ shortage in the past two decades. Since the first successful LDLT from adult to child reported by Strong et al. in 1989 in Australia, the operation has been rapidly taken up by various centers (Miyagawa et al., 2001).