Prevalence of glomerular diseases in Ain Shams University Hospitals

Thesis

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Contents

		Page
List of Abbre	eviations	i
List of Table	es	ii
List of Figur	es	iii
List of Grapl	nics	iv
Introduction		1
Aim of the w	/ork	3
Chapter I:	Epidemiology, updated classification and	
	pathogenesis of glomerulonephritis	4
Chapter II :	Pathological evaluation of renal biopsy	33
Chapter III	: Management of GN	40
Subjects and	Methods	63
Results		67
Discussion .		98
Summary		111
Conclusion		116
Recommend	ations	117
References .		118
Arabic Sumr	nary	



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List of Abbreviations

ACE : Angiotensin-converting enzyme

AGN : Acute endocapillary glomerulonephritis

AKI : Acute kidney injury

ANCA : Antineutrophil Cytoplasmic Antibodies Anti-GBM : Anti glomerular basement membrane

Anti-PLA2R: Anti phospholipase A2 receptor. ARBs : Angiotensin II receptor blockers

AP : Alternative pathway

AZA : Azathioprine

CFB : Complement factor B
CKD : Chronic kidney disease

CP : Classical pathwayCYC : cyclophosphamideDDD : Dense deposit disease

DM : Diabetes mellitus

eGFR : Estimated glomerular filtration rate

EM : Electron microscopy

ESRD : End-stage renal disease

FSGS : Focal segmental glomerulosclerosis
GBM : Glomerular basement membrane

GC : Glucocorticoids

Gd-IgA1 : Galactose deficient IgA1

GN : Glomerulonephritis
GP : Good pastures disease

HBV : Hepatitis B virus
HCV : Hepatitis C virus
HCV : Hepatitis C virus
HTN : Hypertension

HTN : Hypertension

IF : Immunofluorescence microscopy

List of Abbreviations (Cont.)

IFN : Interferone

Ig : Immunoglobulin

IgAN : Immunoglobulin A nephropathy

IP : Immunoperoxidase

KDIGO : Kidney Disease Improving Global OutcomesKDOQI : Kidney Disease Outcomes Quality Initiative

LM : Light microscopy
LN : Lupus nephritis

MAC : Membrane attack complexMCD : Minimal change disease

MGN : Membranous glomerulonephritisMIg : Monoclonal immunoglobulins

MMF : Mycophenolate mofetil

MPGN : Membranoproliferative glomerulonephritis

MPO : myeloperoxidase

NFKB : Nuclear factor Kappa light chain of activated B cells

ANCA : anti-neutrophil cytoplasmic antibody

NETs : Neutrophil extracellular traps

PR3 : Proteinase 3

PLA2R :Phosoholipase A2 receptor

PSGN : Post-Infectious GN

RPGN : Rapidly progressive glomerulonephritis

SLE : Systemic lupus erythematosus

SRNS : Steroid resistant NS

List of tables

Table	Title	Page
1	The recent classification of GN	12
2	Serum complement profiles and auto-	22
	immune features of the major forms of GN	
3	Genetic cause of FSGS and Nephrotic syndrome	24
4	Description of the Demographic data and co morbidities	64
5	Description of the clinical presentation of patients at time of biopsy	68
6	Description of the pathological Diagnosis of patients	70
7	Description of age groups and pathological diagnosis	80
8	Description of sex in relation to diagnosis	83
9	Description of viral hepatitis in relation to clinical diagnosis	85
10	Description of HTN in relation to the pathological diagnosis	86
11	Description of mean CR, BUN and 24H protein and pathological diagnosis at time of biopsy	87
12	Description of clinical presentation and pathological diagnosis	88
13	Description of pattern of IF and EM among FSGS cases	92
14	Laboratory Data among FSGS cases at the time of presentation	93

List of tables (Cont.)

Table	Title	Page
15	Description of ISN/RPS class, NIH activity and chronicity among lupus nephritis	93
16	Description of pattern of IF and EM among lupus nephritis cases	94
17	Laboratory data among lupus nephritis cases at time of biopsy	95
18	Description of types, IF and EM pattern among membranoprotliferative	96
19	Laboratory Data among Membrano- proliferative	97

List of figures

Fig.	Title	Page
1	Classification of GN	10
2	Proposed classification for membrano- proliferative glomerulonephritis based on the presence or absence of Igs	16
3	Structure of glomerulus	20
4	The clinical syndromes associated with podocyte dysfunction of various types	21
5	Proposed work-up of complement mediated membrano-proliferative glomerulonephritis	29
6	Diagram showing examples of correct and incorrect placement of needle biopsy devices	34
7	Class LN III/IV induction therapy	59
8	Treatment of class V	60
9	A case of FSGS showing a glomerulus with segmental sclerosis (arrow) and mild mesangial hypercellularity with neutrophil infiltrate.	72
10	An electron micrograph of FSGS showing focal thickening of glomerular basement membrane with diffuse effacement of podocytes.	73
11	A case of Lupus nephritis ISN/RPS class IV (G/A) in a 25 years old female presented by nephrotic range proteinuria.	74
12	An electron microgrpah of the same case. Subendothelial electron dense deposits (SED) are seen in relation to all parts of the GBM.	75
13	A case of IgA nephropathy with fibrocellular crescent (arrow), lobulated tuft and moderate global hypercellularity with mesangial expansion.	76
14	An electron micrograph for a case of IgA nephropathy showing mesangial and	77

Fig.	Title	Page
	paramesangial electron dense deposits (arrows).	
15	A case of membranoproliferative glomerulo- nephritis showing global thickening capillary walls and endocapillary proliferation with mild neutrophil infiltration.	78
16	an electron micrograph of a case of membrano- proliferative glomerulonephritis showing extensive subendothelial, subepithelial, and mesangial electron dense deposits (original magnification: 3800)	79

List of graphics

Graph.	Title	Page
1	Clinical presentation at time of biopsy	69
2	Pathological diagnosis in renal biopsies	71
3	Glomerular diseases among children	81
4	Glomerulonephritis among adults	82
5	Glomerulonephritis among elderly	83
6	Glomerulonephritis among males and	84
	females	
7	Pathological diagnosis which presented	90
	by nephrotic syndrome	
8	Pathological diagnosis which presented	90
	by CKD	
9	Pathological diagnosis which presented	91
	by AKI	

Prevalence of glomerular diseases in Ain Shams University Hospitals Abstract

Introduction: Glomerulonephritis (GN), is a mass term used for a substantial number of diseases with the common denominator of histological renal inflammation emerging from the glomerular tuft, it is the second most common cause of chronic renal failure. GN is a major cause of morbidity and mortality from renal disease in many parts of the world, particularly in the tropical and subtropical regions. According to several local registries and sporadic publications, it is responsible for about23.2 to 58.4% of patients on regular dialysis in the tropics, compared to contemporary areas of around 16-18% in the United States and 9-15% in Europe. It is responsible for 16.6% which has been reported in 1998 in Egypt.

Aim of the Work: To study the Histologic spectrum of glomerular diseases among Egyptian patients in Ain Shams university hospitals for a period of 3 years from 2011 to 2013 with its correlation with the clinical and laboratory parameters (Single center).

Subjects and Methods: A cross sectional retrospective study was done in pathology lab and EM lab of Ain Shams University specialized Hospitals referral center from different governorates in Egypt. We studied all clinical cases with glomerular diseases diagnosed with renal biopsies over a period of 3 years from 2011 to 2013. The study included 857 renal biopsies evaluated by LM, EM, IF.

Results: This is a retrospective study aimed at data collection in three years from 2011 to 2013 in Ain Shams university hospitals. The study included 857 renal biopsies, cases 2011: 163 (19%), 2012: 349 (40.7%) and 2013:345 (40.3%) with mean of age 31.35±18.43 years and range (3-81) years. Children <16 years: 241 (28.3%), adults (17-60years): 548 (64.2%), elderly >60 years: 64 (7.5%). 386 (45%) of total cases were females and 471 (55%) were males, co morbidities as hypertension in (22.9%) of cases followed by DM (5.6%) of studied cases.

We recorded that 196 patients were presented by hypertension and the most common pathological diagnosis associated with HTN is FSGS in (18.4 %) followed by membranoproliferative glomerulonephritis in (16.8 %): The most common pathological diagnosis in nephrotic syndrome is minimal change disease followed by membranous glomerulonephritis; In asymptomatic proteinuria is lupus nephritis followed by membranous glomerulonephritis; In CKD is FSGS followed by global glomerulo-sclerosis; In hematuria is IGA followed by Alport syndrome; In nephritic is membranoproliferative followed by FSGS; In RPGN in crescentic followed by lupus nephritis; AKI in lupus nephritis followed by membranoproliferative; Nephrotic nephritic in diffuse proliferative followed by membranoproliferative.

Keywords: FSGS: Focal segmental glomerulosclerosis; GN: Glomerulonephritis; MPGN: Membranoproliferative glomerulonephritis

Introduction

Glomerulonephritis (GN), is a mass term used for a substantial number of diseases with the common denominator of histological renal inflammation emerging from the glomerular tuft, it is the second most common cause of chronic renal failure (Segelmark and Hellmark, 2010).

GN is a major cause of morbidity and mortality from renal disease in many parts of the world, particularly in the tropical and subtropical regions. According to several local registries and sporadic publications, it is responsible for about 23.2 to 58.4% of patients on regular dialysis in the tropics, compared to contemporary areas of around 16-18% in the United States and 9-15% in Europe. It is responsible for 16.6% which has been reported in 1998 in Egypt (*Barsoum and Francis*, 2000).

In Syria, Moukeh et al reported that the prevalence of GN as a cause of ESRD is 20.5%, in Kwait GN was the first cause of ESRD by 32% (*Zahran*, *2011*).

In United Kingdom GN is the first cause of ESRD by 16% in 2010 (*Steenkamp et al., 2011*) and 19% in 2014 (18th annual registry) (*Macneil et al., 2016*).

Glomerular disease (GD) is one of the most common forms of renal diseases and can have many different clinical presentations. It can present as nephrotic syndrome (NS), nephritic syndrome, rapidly progressive renal failure (RPRF), acute kidney injury (AKI), chronic kidney disease (CKD), macroscopic hematuria (MH), recurrent disease in the post-

transplant kidney, as well as isolated proteinuria or hematuria (Golay et al.,2013).

Immune mechanisms are responsible for glomerular injury in most cases of primary glomerulonephritis and many of the secondary GN (*Segelmark and Hellmark*, 2010).

Biopsy registries can give an idea about the regional variations in the spectrum of GD as well as the trend over time, however, there is a variation in the prevalence of the type of GD according to geographical location and race of the study population (*Jennette et al.*, 2013).

The pattern of glomerular disease is different in various countries, and is changing with time within the same country. For instance, minimal change disease (MCD) was the most common cause of glomerular disease in Northern India, primary IgA nephropathy (IgAN) was more common in young adults from Western India, and mesangioproliferative glomerulonephritis (MesGN) was the predominant disease in South India (*Pesce and Schena*, 2010).

There is also a changing pattern of the incidence of GN in the different parts of the world. For instance, the incidence of end-stage renaldisease (ESRD) as a result of focal segmental glomerulosclerosis (FSGS) has increased 11-fold in the past two decades in a recent US study (*Ibrahim et al.*, 2012).

Aim of the Work

To study the Histologic spectrum of glomerular diseases among Egyptian patients in Ain Shams university hospitals for a period of 3 years from 2011 to 2013 with its correlation with the clinical and laboratory parameters (Single center).

Chapter 1

Epidemiology, updated classifications and pathogenesis of glomerulonephritis

Epidimiology:

Identification of the profile of glomerular disease in a certain geographical region is very important academically, clinically and epidemiologically. It helps in the identification of specific risk factors and for adequate prevention (*Barsoum and Francis*, 2000).

Racial factors is also very important, not only in the incidence, but also it define the pattern, severity and progression of the glomerular disease, environmental factors may also be involved as modifiers of the glomerular pathology in different geographical regions, of particular interest is the role of heavy metal and hydrocarbon pollution, the epidemiological significance of which remains to be elucidated. For these reasons, it is of considerable interest to identify the patterns of glomerular disease in specific regions (*Barsoum and Francis*, 2000).

GN varies in incidence among the different geographical areas due to socioeconomic conditions, ethnicity, genetic variability and environmental factors. Recent studies suggested a changing pattern of incidence of GN in the different parts of the world (*Ibrahim et al.*, 2012).

IgAN was the most common primary GN in young adults Caucasians (Europe and USA) (*Zaza et al., 2013*), and some countries in Asia as described in reports from China, Japan and Korea, and is the most common cause of end-stage renal disease