

Epidemiology of CKD in Children



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CKD Course
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Aim & Plan

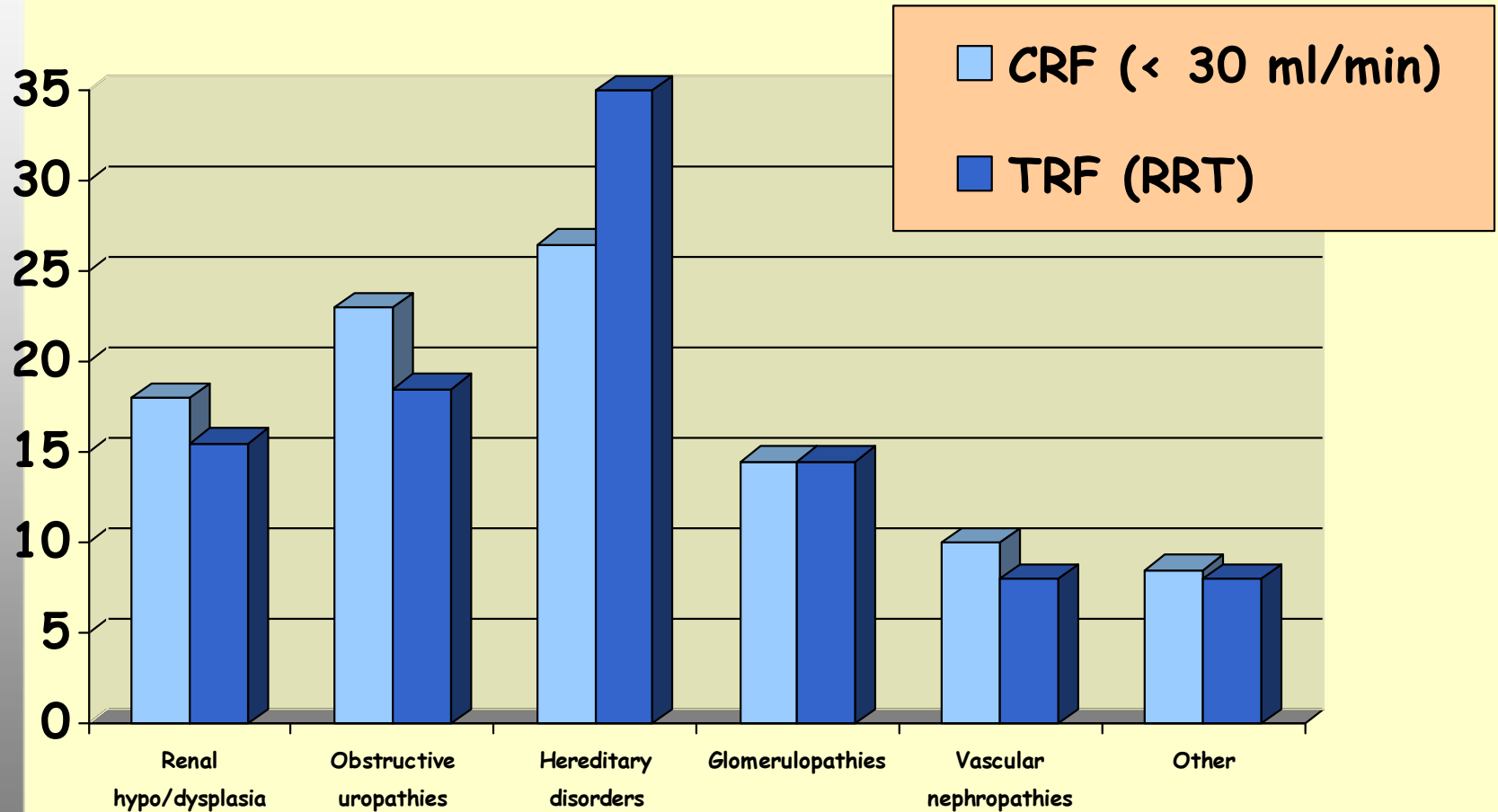
- Causes of CKD in children
- Incidence & prevalence & risks for progression
- Plan
 - 1990-2000
 - Italkid Project
 - ESPN Registry, NAPRTCS 2008 Annual reports
 - Data from Turkey

K/DOQI stages of CKD

Stage	Description	GFR (ml/min/1.73 m ²)
I	Kidney damage with normal or increased GFR	≥90
II	Kidney damage with mild decrease in GFR	60-89
III	Moderate decrease in GFR	30-59
IV	Severe decrease in GFR	15-29
V	Kidney failure	<15 (or dialysis)

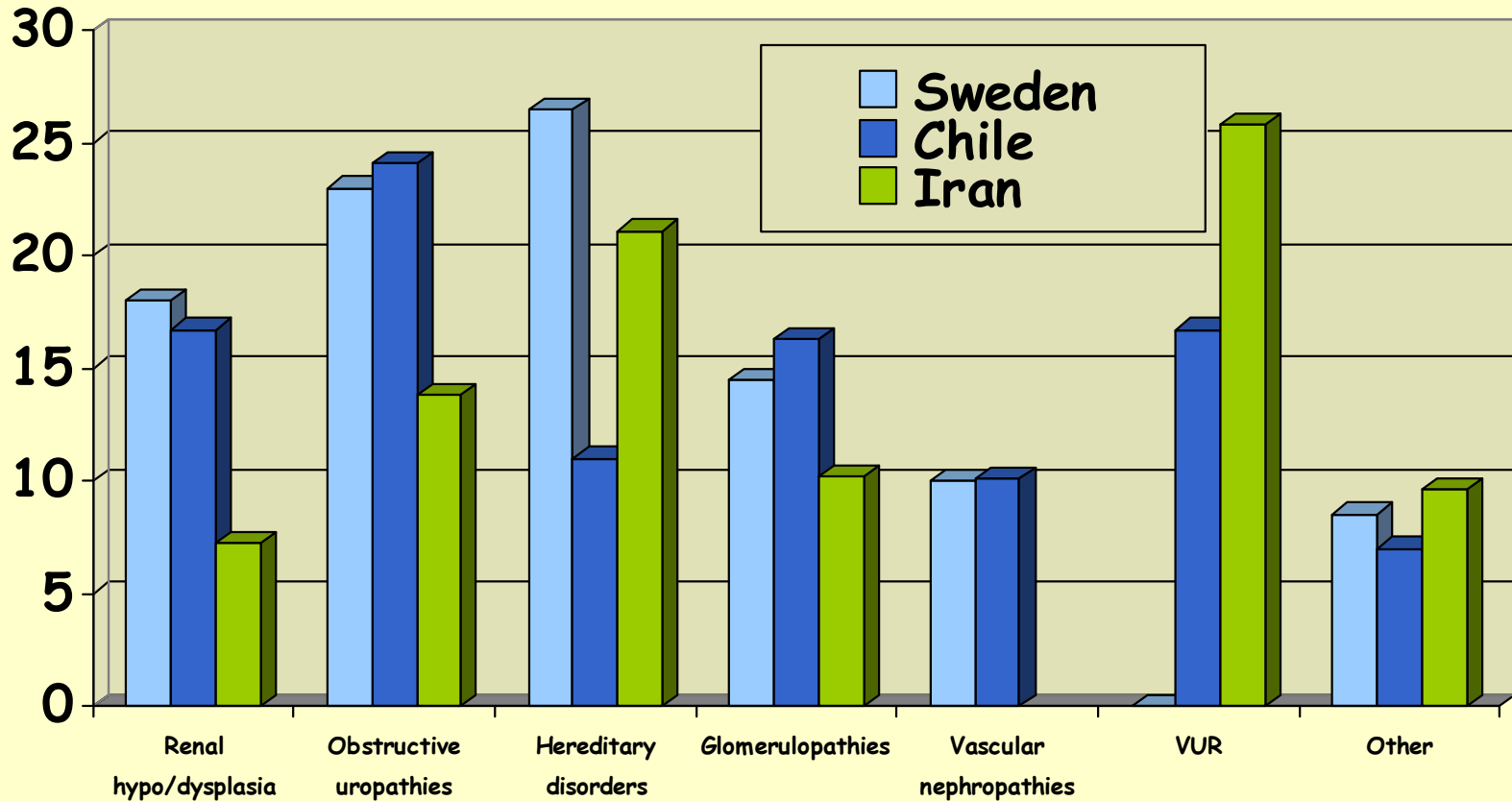
Year of publication	Country	Period	Study design	Definition	N	M/F
1995	Turkey	1979-1993	Single center	GFR <50	459	1.2
1997	Sweden	1986-994	Nationwide	GFR <30	118	1.6
1999	Chile	1996	Nationwide	GFR <30	227	1.0
1999	India	NA-1 year	Single center	Cre > 2 mg/dl	48	2.2
2001	Iran	1991-1999	Single center	GFR <30	166	1.3
2002	Japan	1998	Nationwide	ESRD	105	1.9
2003	Nigeria	1985-2000	Single center	GFR <30	45	1.6

Primary renal diseases in children; Sweden 1986-1994



■ Esbjörner et al. *Pediatr Nephrol* 1997; 11: 438-442.

Country	n	Incidence	Prevalence
Sweden (1986-1994)	118	7.7	25.0
Chile (1996)	227	5.7	42.5
Iran (1991-1999)	166	NA	NA



- Esbjörner et al. *Pediatr Nephrol* 1997; 11: 438-442.
- Lagomarsimo et al. *Pediatr Nephrol* 1999; 13: 288-291.
- Madani et al. *Pediatr Nephrol* 2001; 16: 140-144.

Epidemiology of Chronic Renal Failure in Children: Data From the Italkid Project

Gianluigi Ardissino, Valeria Daccò, Sara Testa, Roberto Bonaudo, Aldo Claris-Appiani, Emanuela Taioli, Giuseppina Marra, Alberto Edefonti and Fabio Sereni

Pediatrics 2003;111:e382-e387

DOI: 10.1542/peds.111.4.e382

The lack of population-based information means that the epidemiology of childhood CRF is often based on renal replacement therapy data,⁷ but the vast majority of children with renal impairment (particularly congenital abnormalities) reach ESRD when they are far beyond pediatric age and are therefore not included.



ItaKid Project

- Population based registry in 1990
- Covered the entire country, 16.8 million children
- Inclusion criteria
 - Creatinine clearance of $75 \text{ ml/min/1.73 m}^2$
 - for children < 1 year: Serum Cre > mean sCr + 3 SD;
 - Age younger than 20 years at the time of registration

ItaKid Project

- In 10 years, 1.197 pts. have been registered.
- M/F:
 - 2.03 for the population as a whole
 - 1.72 when the pts. with PUV were excluded.
- The mean age at registration: 6.9 ± 5.4 years
- The mean Ccr at registration: 41.7 ± 20.5 mL/min/ 1.73 m²
- The mean incidence (1995-2000): 12.1 cases pmarp (8.8-13.9).
- The point prevalence (January 1, 2001) 74.7 cases pmarp

TABLE 1. Primary Cause of CRF in Children*

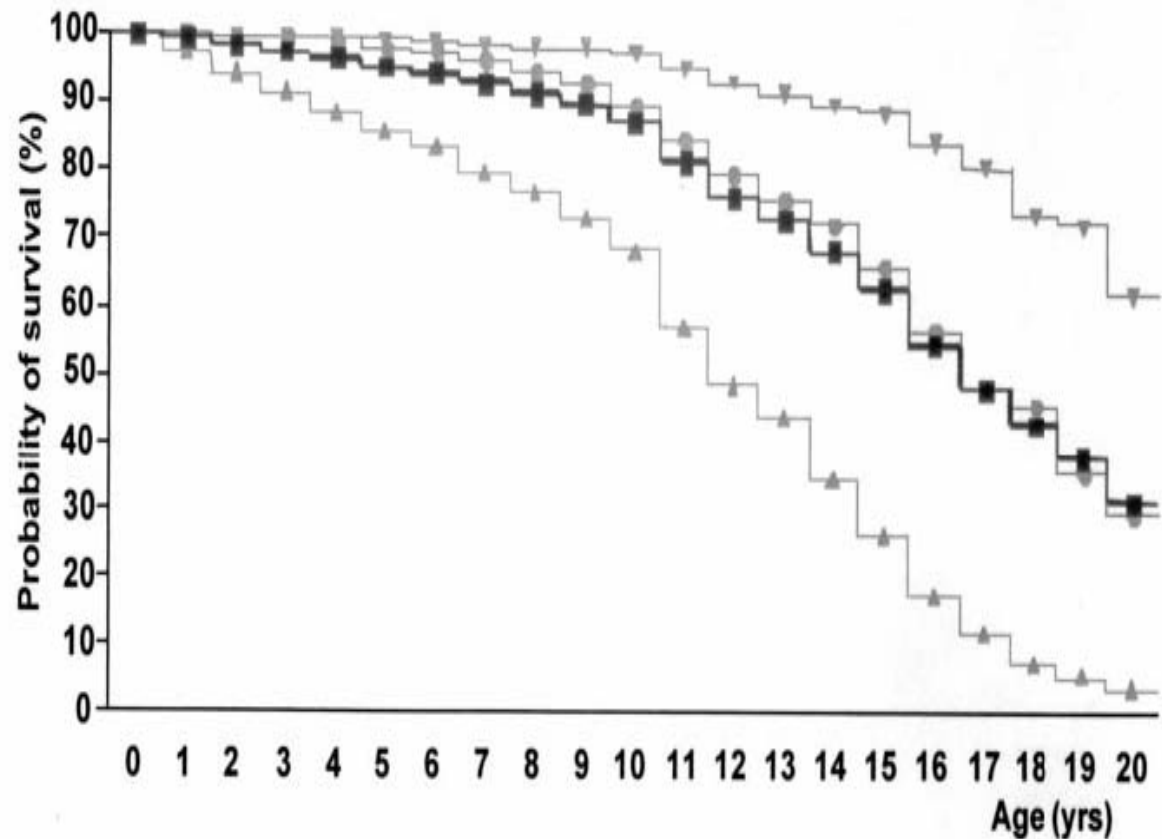
	All Registered Patients		Patients Reaching ESRD	
	No.	%	No.	%
Hypodysplasia				
With identified uropathy	522	43.6	71	27.1
Without urinary tract malformations	167	13.9	33	12.5
Neurogenic bladder	44	3.7	8	3.0
Chronic glomerulonephritis†	31	2.6	16	6.1
Focal glomerulosclerosis	21	1.8	12	4.6
Congenital nephrotic syndrome	13	1.1	5	1.9
Membranous nephropathy	3	0.3	2	0.8
Systemic lupus erythematosus	13	1.1	5	1.9
Hemolytic uremic syndrome	43	3.6	9	3.4
Polycystic kidney disease	60	5.0	9	3.4
Nephronophthisis	41	3.4	21	8.0
Alport's syndrome	18	1.5	9	3.4
Cystinosis	22	1.8	9	3.4
Hereditary nephropathies‡	45	3.8	13	4.9
Cortical necrosis (perinatal)	49	4.1	8	3.0
Medications	14	1.2	2	0.8
Idiopathic interstitial nephritis	24	2.0	9	3.4
Wilms' tumour	4	0.3	3	1.1
Miscellaneous non-hereditary diseases	23	1.9	8	3.0
Unknown	40	3.3	11	4.3

* Including monolateral renal agenesis (65), nephrectomy (30), and multicystic kidney (25).

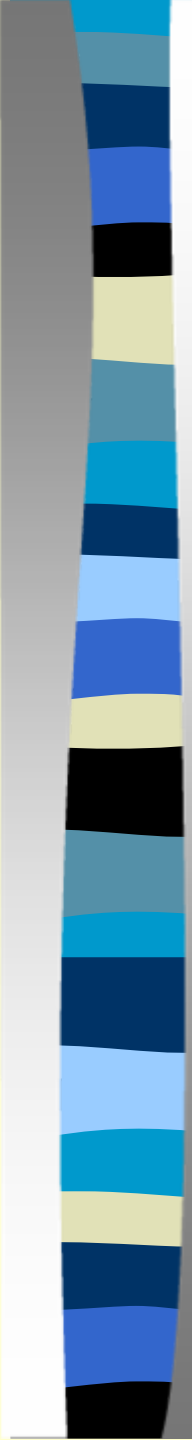
† Including unlisted systemic immunological diseases.

‡ Other than those individually listed.

Fig 2. Estimated kidney survival in children with CRF by age. Overall population ($n = 1197$; \blacksquare); patients with baseline creatinine clearance <25 mL/min ($n = 315$; \blacktriangle); 25-50 mL/min ($n = 419$; \bullet); 51-75 mL/min ($n = 463$; \blacktriangledown).



- The incidence of RRT was 7.3/year/100 patients
- The risk of developing ESRD by the age of 20 years was 68%



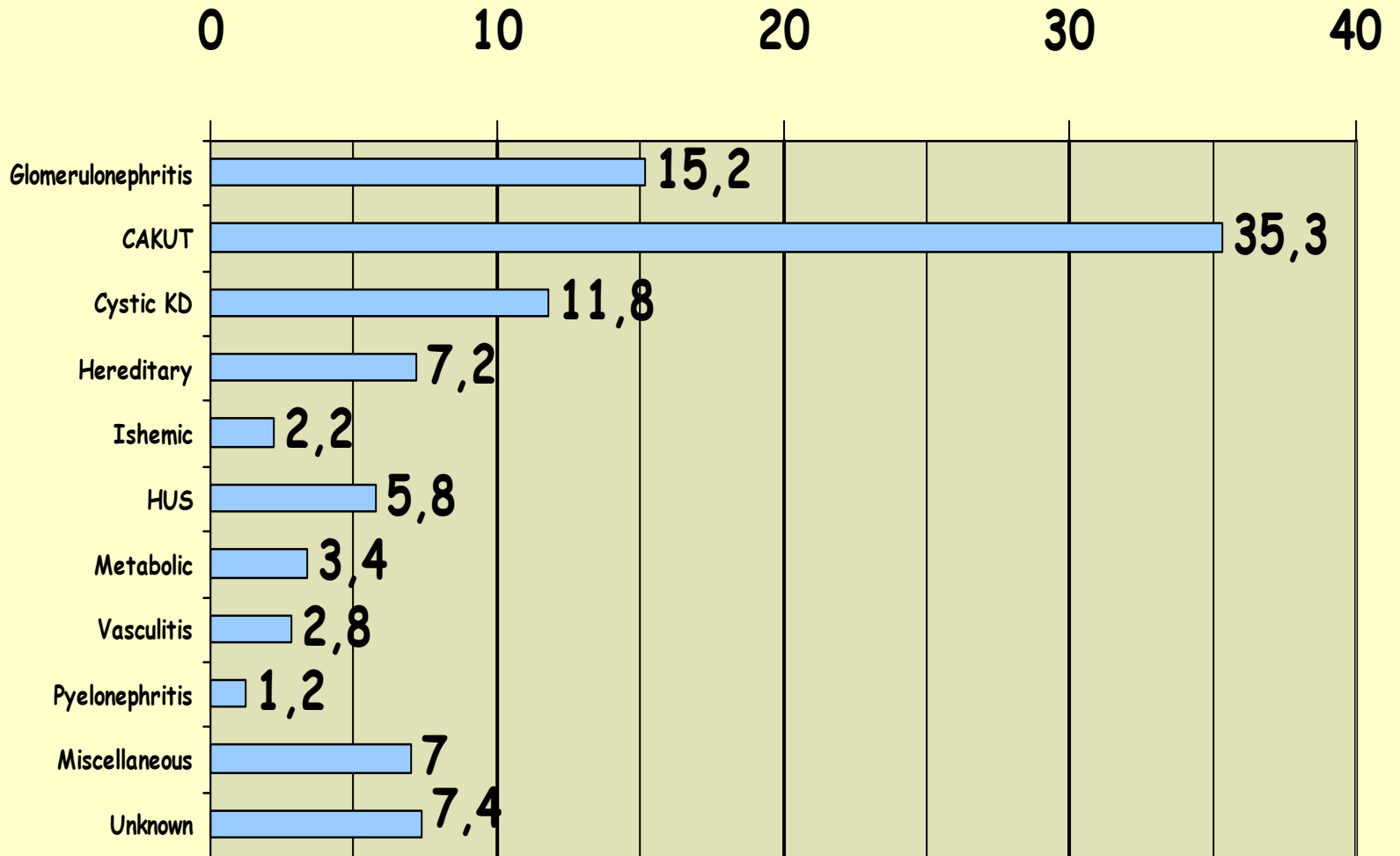
ESPN 2008 Report

- Total population: 628 million
- Children (0-14 years): 98 million (15.6%)

Patients accepted for RRT

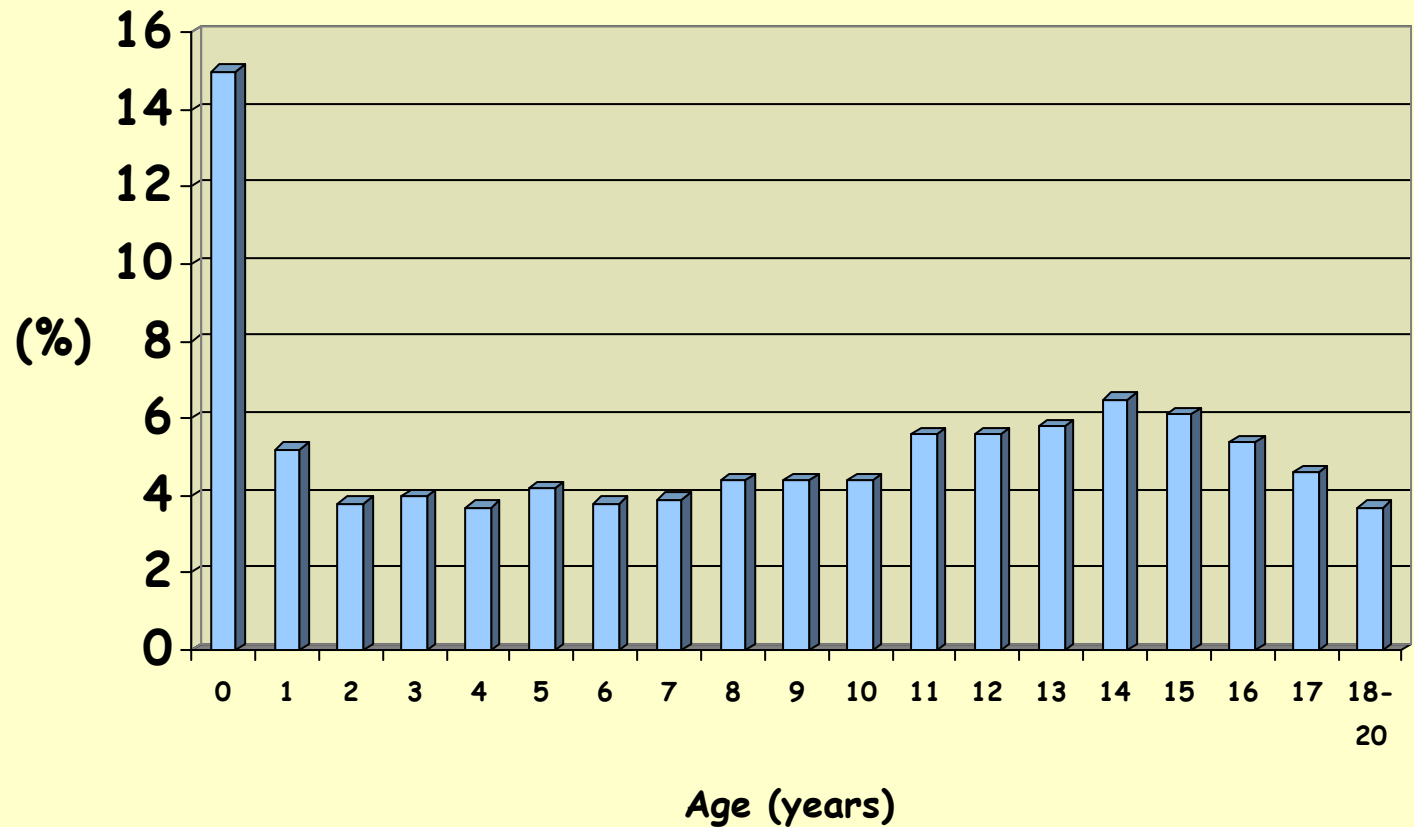
- Incidence 5.2 pmarp (0-24.5)
- Prevalence 28.6 pmarp (3.2-72.6)

ESPN 2008 Report



NAPRTCS 2008 (n: 7036)

(1994; GFR < 75; Canada, Mexico, Costa Rica, USA)

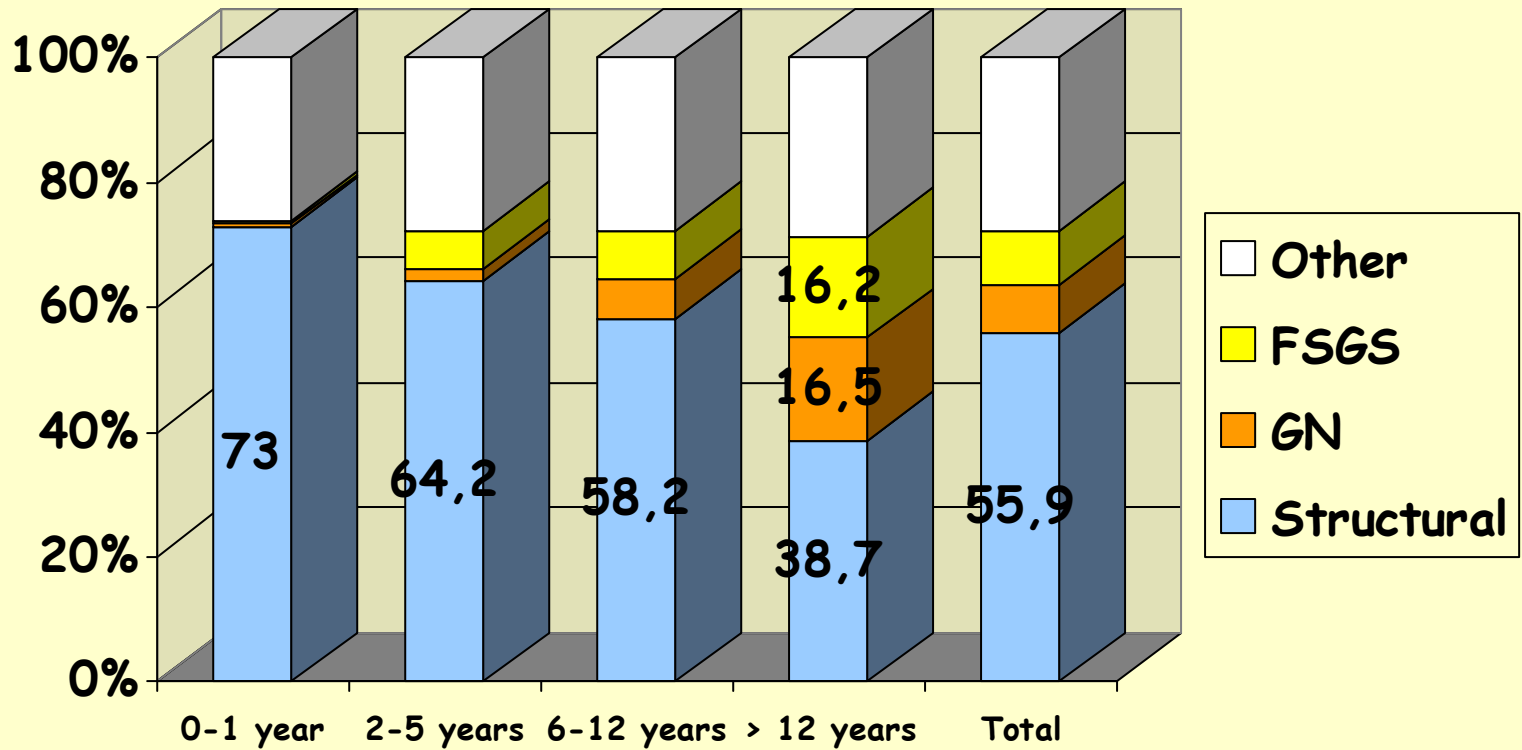




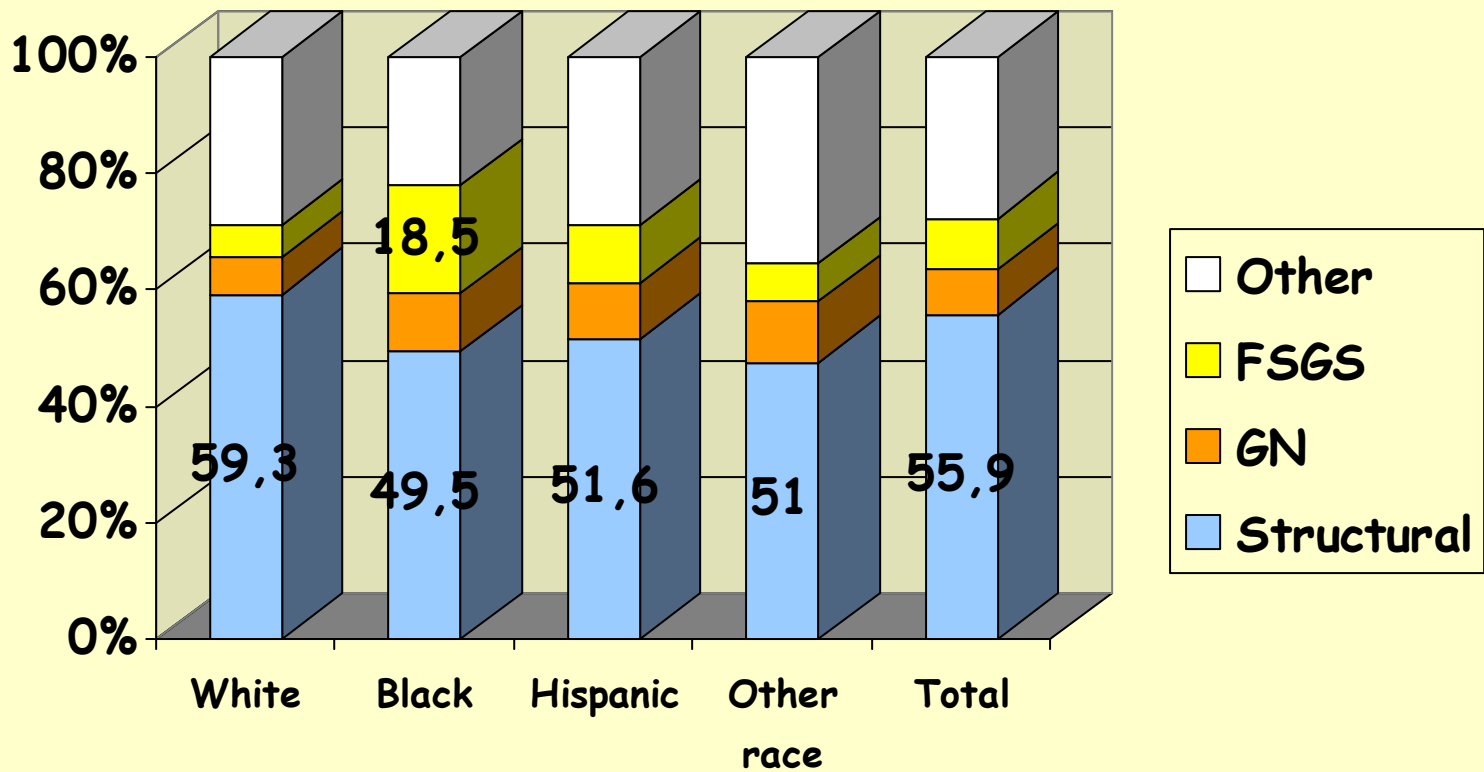
NAPRTCS 2008

- Structural
 - Prune belly, reflux nephropathy and aplasia/hypoplasia/displasias
- Glomerulonephritis
 - Chronic GN, idiopathic crescentic GN
 - MPGN, MGN, SLE, HSP, IgAN
 - Wegener's granulomatosis
- FSGS
- Other

NAPRTCS 2008 (n: 7026)



NAPRTCS 2008



**EXHIBIT 14.4
PROGRESSION TO ESRD**

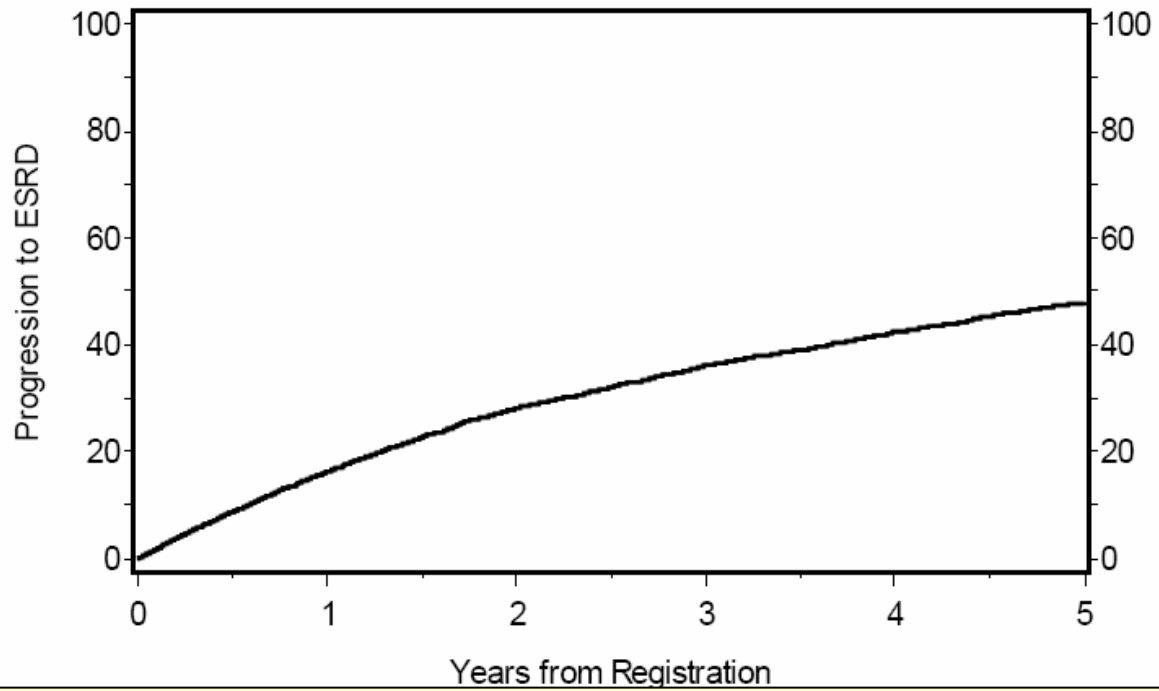


EXHIBIT 14.5
PROGRESSION TO ESRD
BY BASELINE CALCULATED CREATININE CLEARANCE (mL/min/1.73 m²)

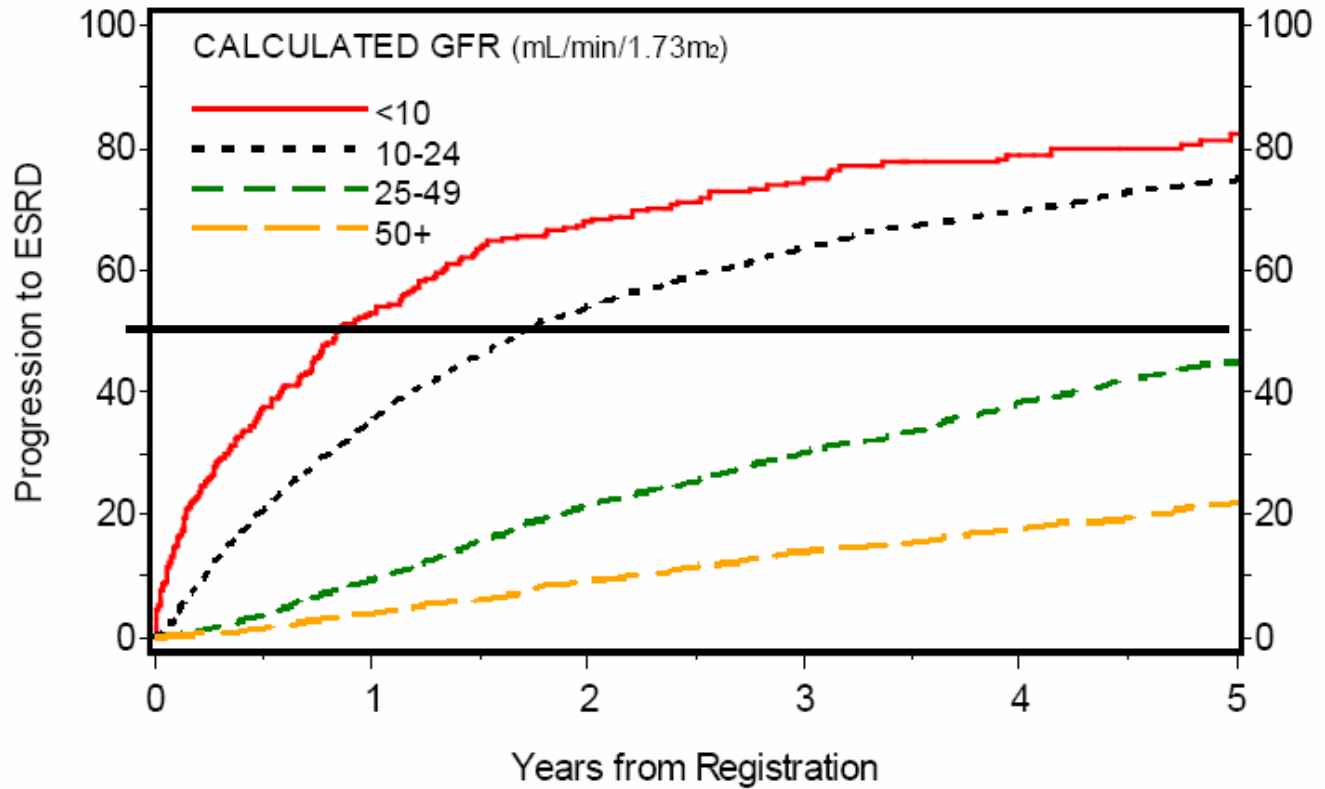
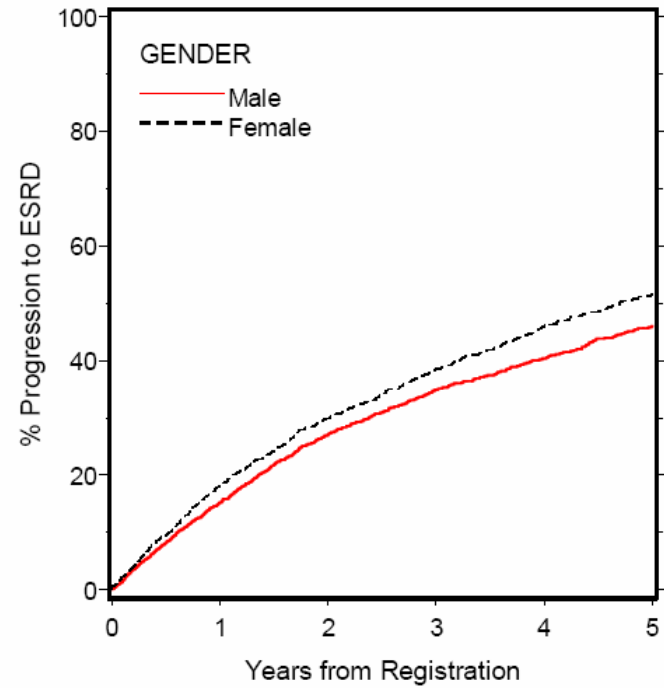
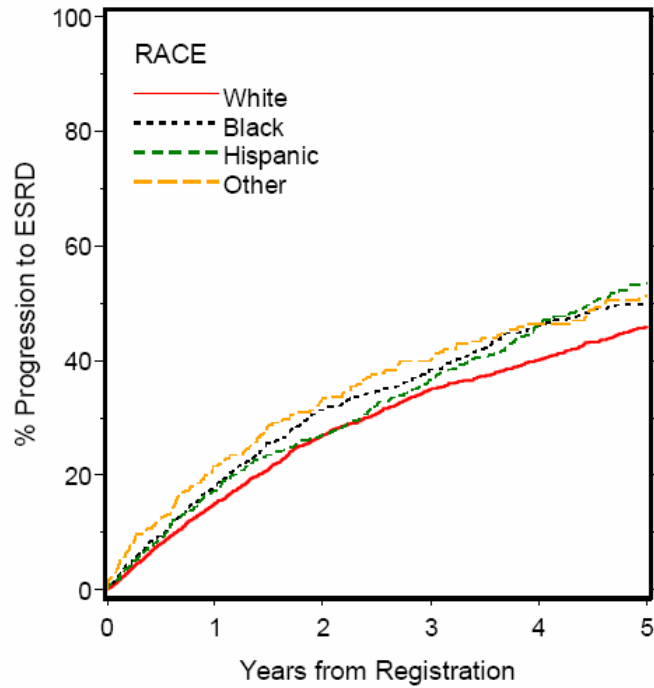
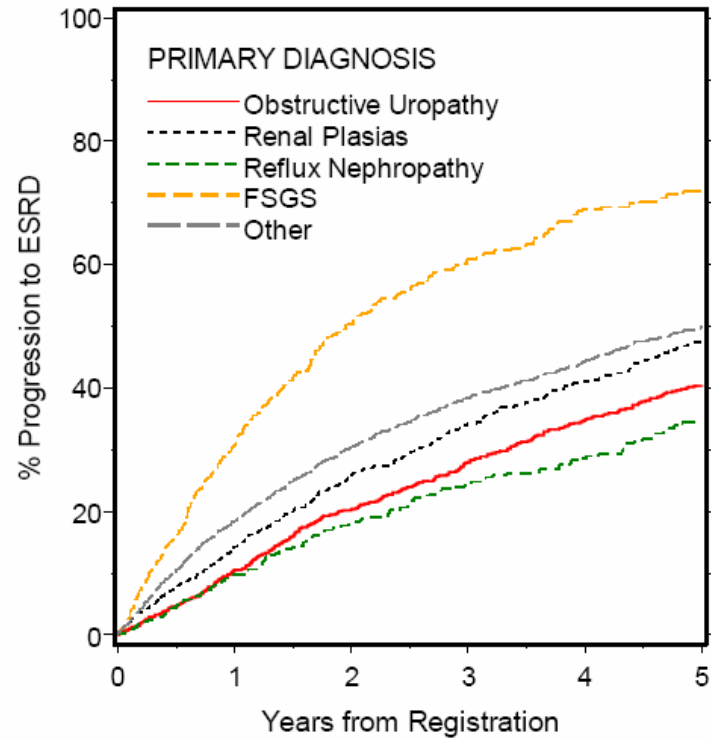
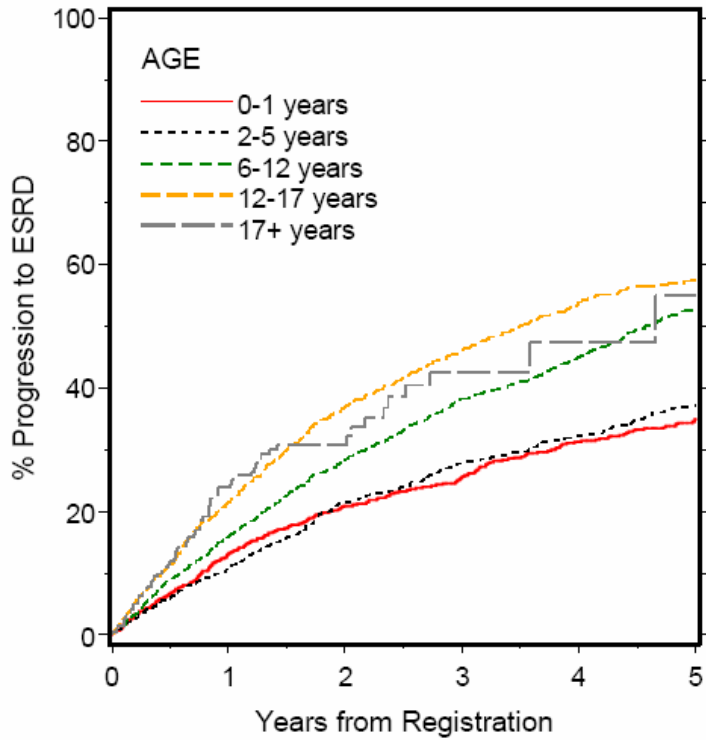
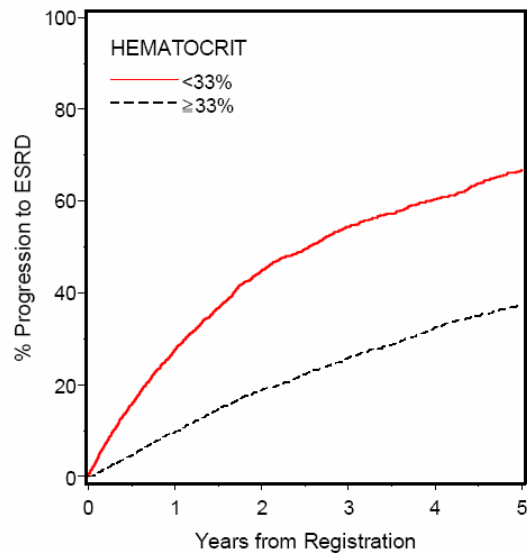
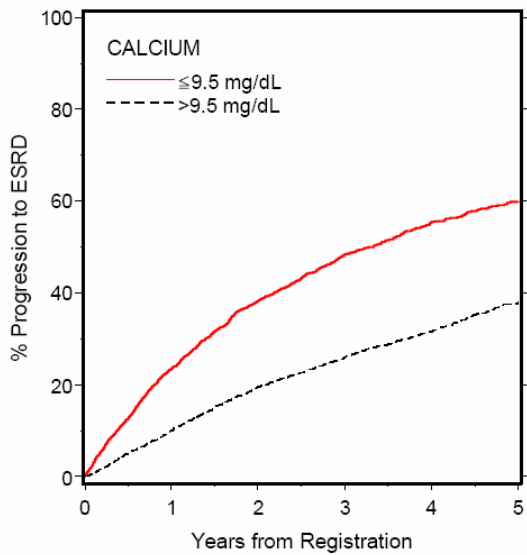
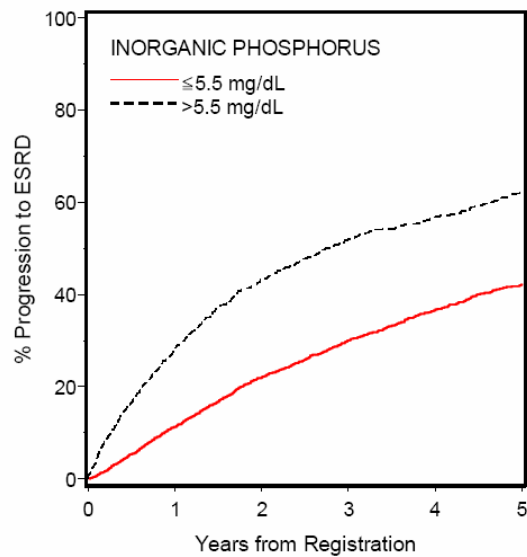
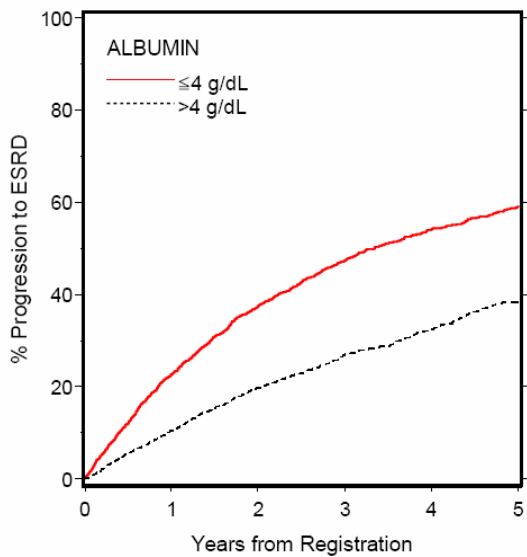


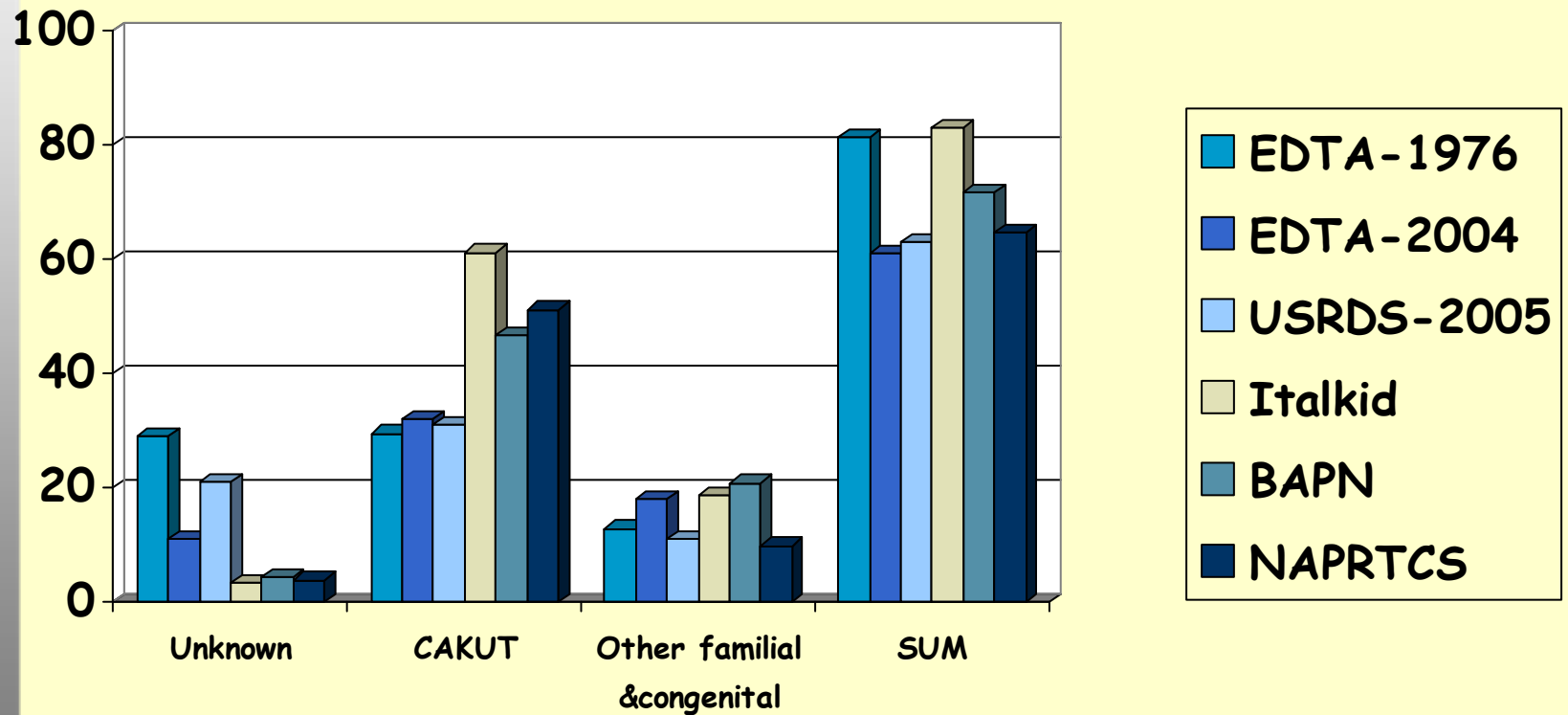
EXHIBIT 14.6 PROGRESSION TO ESRD BY SELECTED PATIENT CHARACTERISTICS







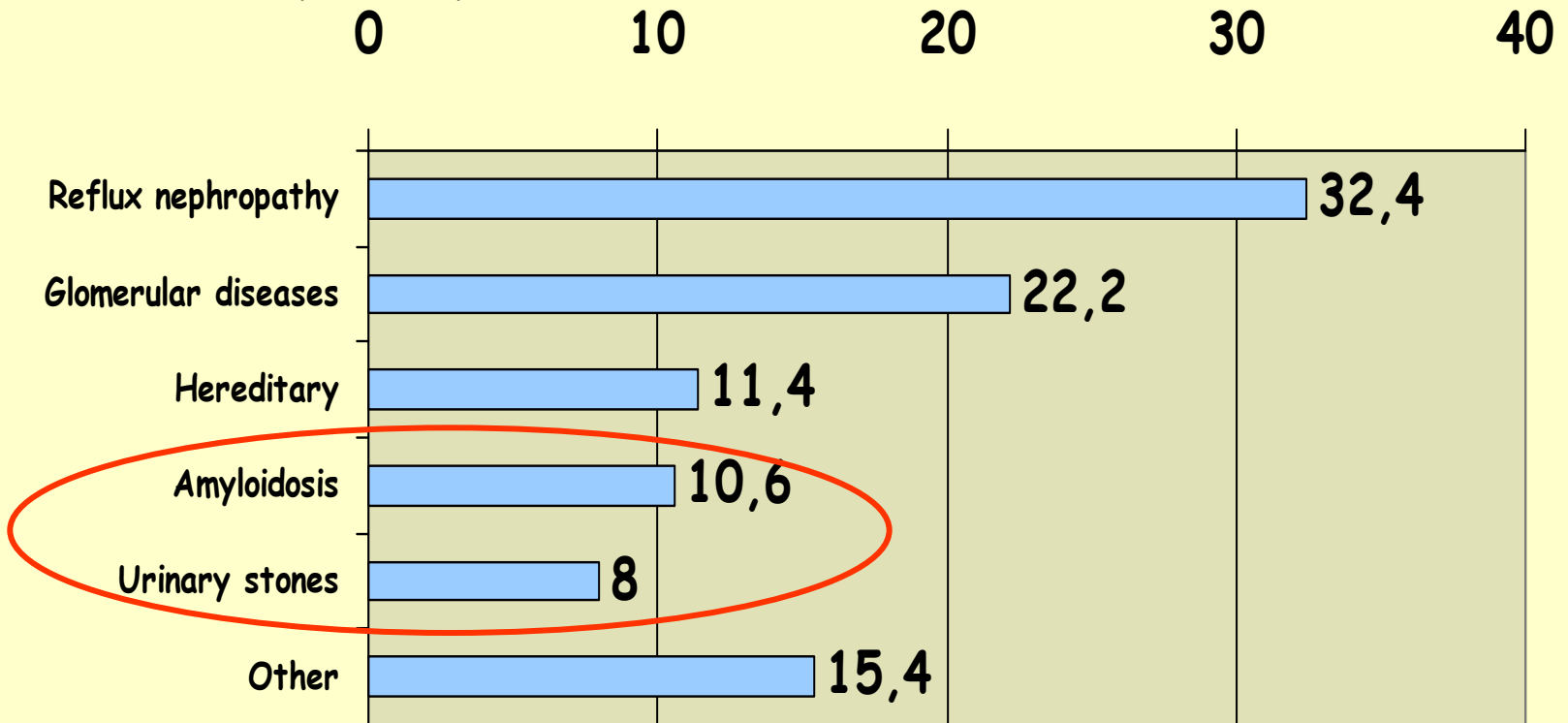
Causes of CKD



Adapted from: Neild GH. *Pediatr Nephrol* 2009; 24: 1913-1919.

Etiology of chronic renal failure in Turkey

- Study period: 1979-1993
- GFR < 50
- 459 children
- M/F: 1.23 (254/205)



■ Sirin A et al. *Pediatr Nephrol* 1995; 9: 549-552.

Pediatr Nephrol (2009) 24:797–806

DOI 10.1007/s00467-008-0998-4

ORIGINAL ARTICLE

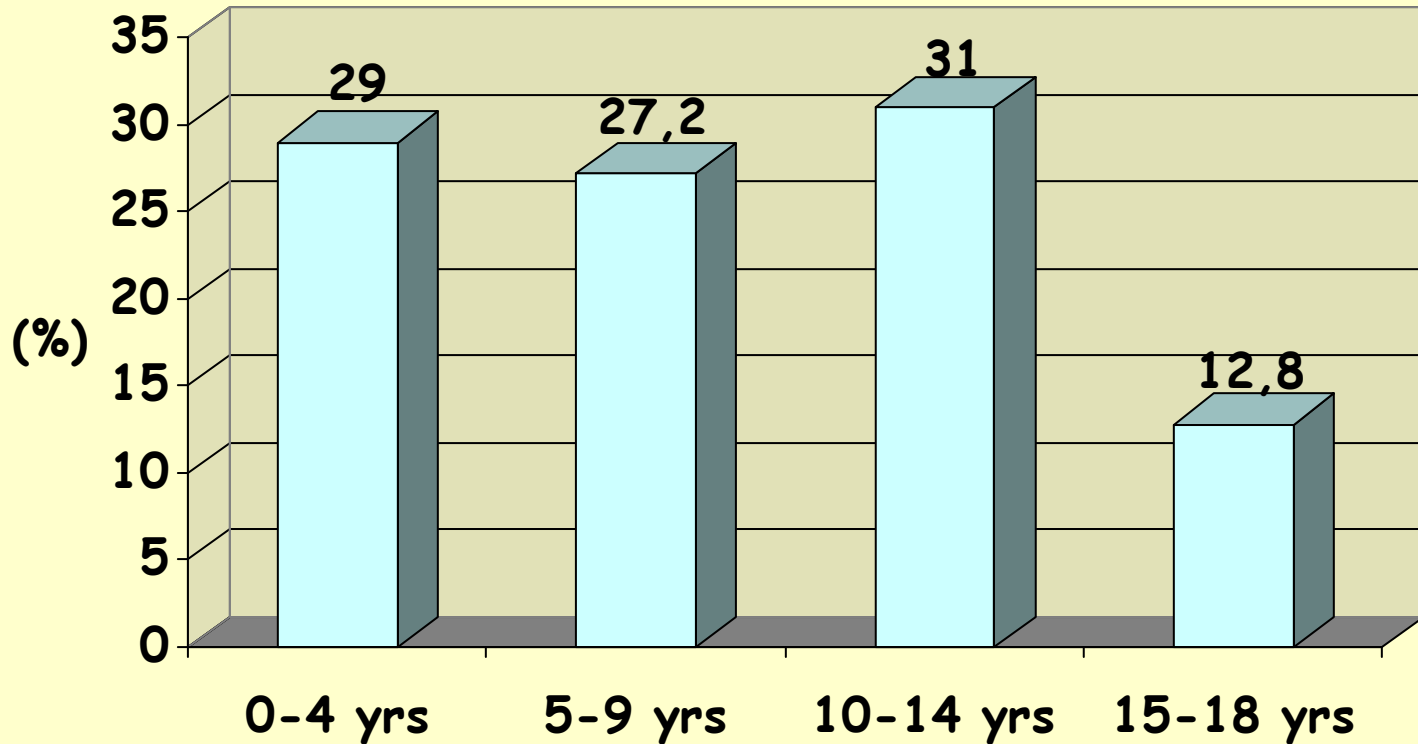
Chronic kidney disease in children in Turkey

Kenan Bek • Sema Akman • Ilmay Bilge •
Rezan Topaloğlu • Salim Çalışkan • Harun Peru •
Nurcan Cengiz • Oğuz Söylemezoğlu

- Turkish Society for Pediatric Nephrology-CKD Working Group
- 29 pediatric nephrology centers in Turkey
- $eGFR \leq 75$ ml/min per 1.73 m^2
- < 19 years, diagnosed in 2005

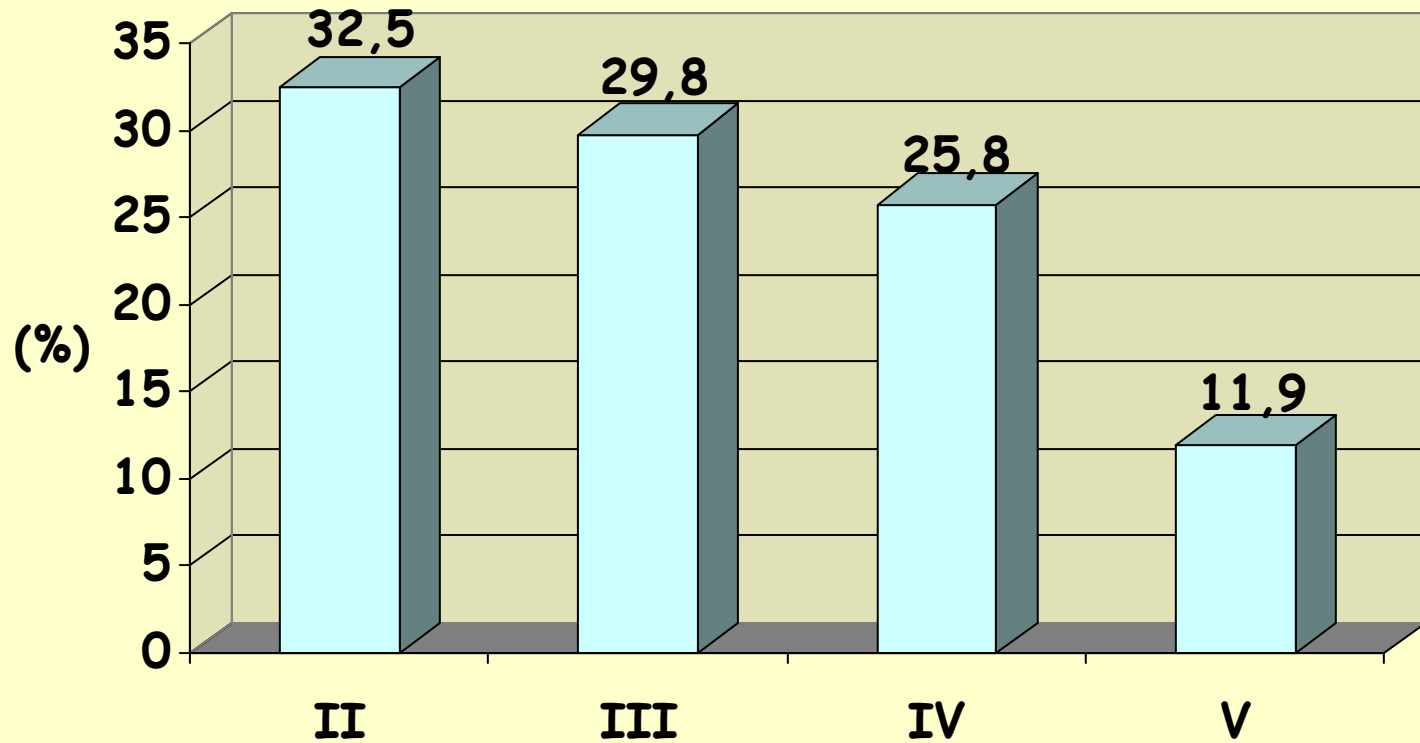
- 282 patients
 - M/F 1.29 (159/123)
 - mean age 8.05 ± 5.25 years

Age distribution of patients at diagnosis



■ Bek et al. Turkish SPN CKDWG. *Pediatr Nephrol* 2009; 24:797-806.

CKD stages of the patients at diagnosis



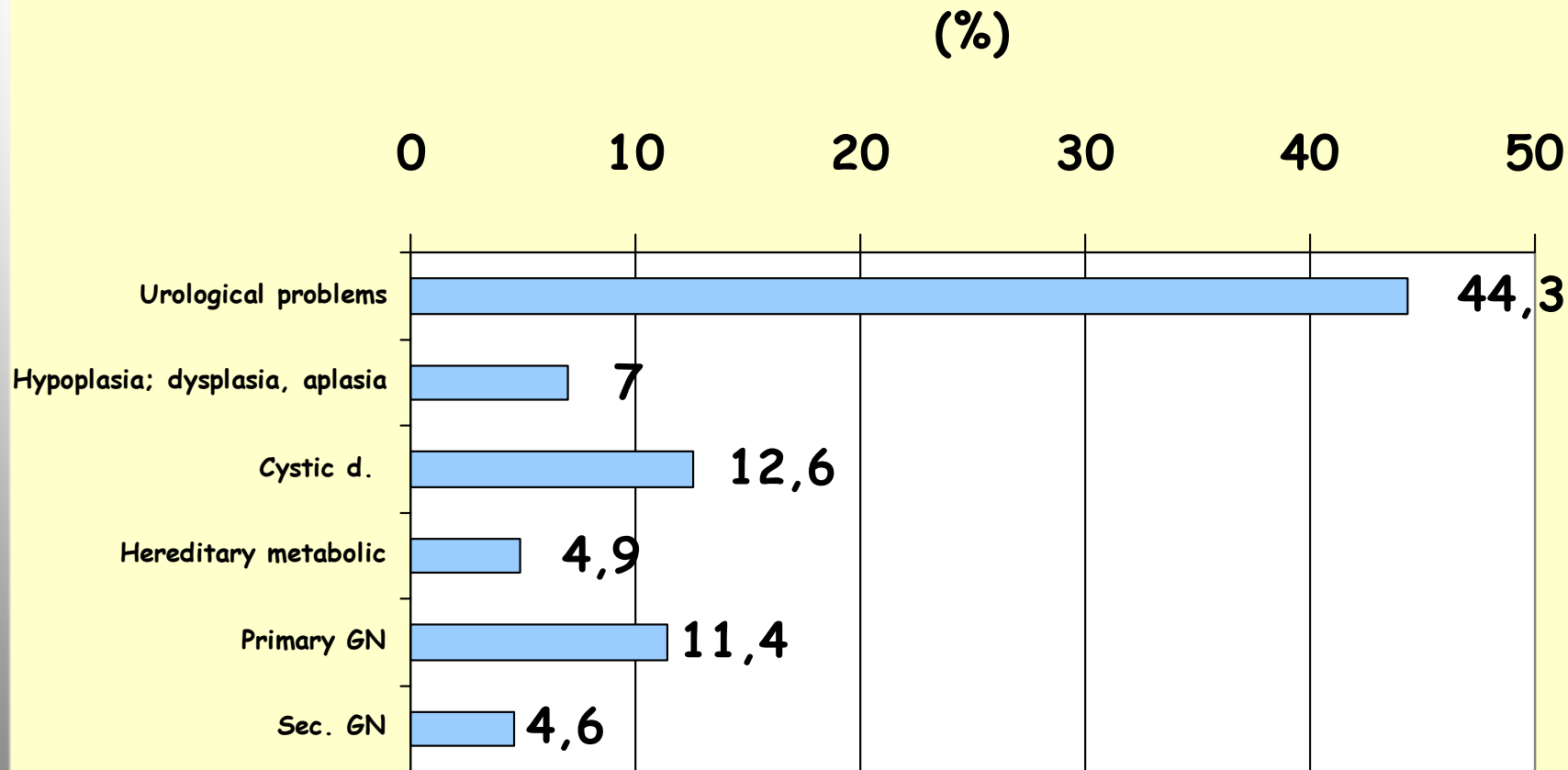
■ Bek et al. Turkish SPN CKDWG. *Pediatr Nephrol* 2009; 24:797-806.

Table 3 Etiologic classification of the patients (*FSGS* focal segmental glomerulosclerosis, *PKD* polycystic kidney disease, *SLE* systemic lupus erythematosus, *HUS* hemolytic uremic syndrome, *HSP* Henoch–Schönlein purpura)

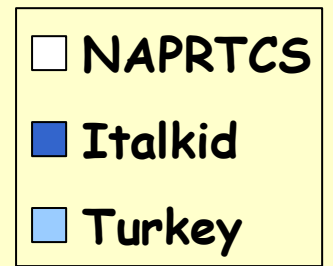
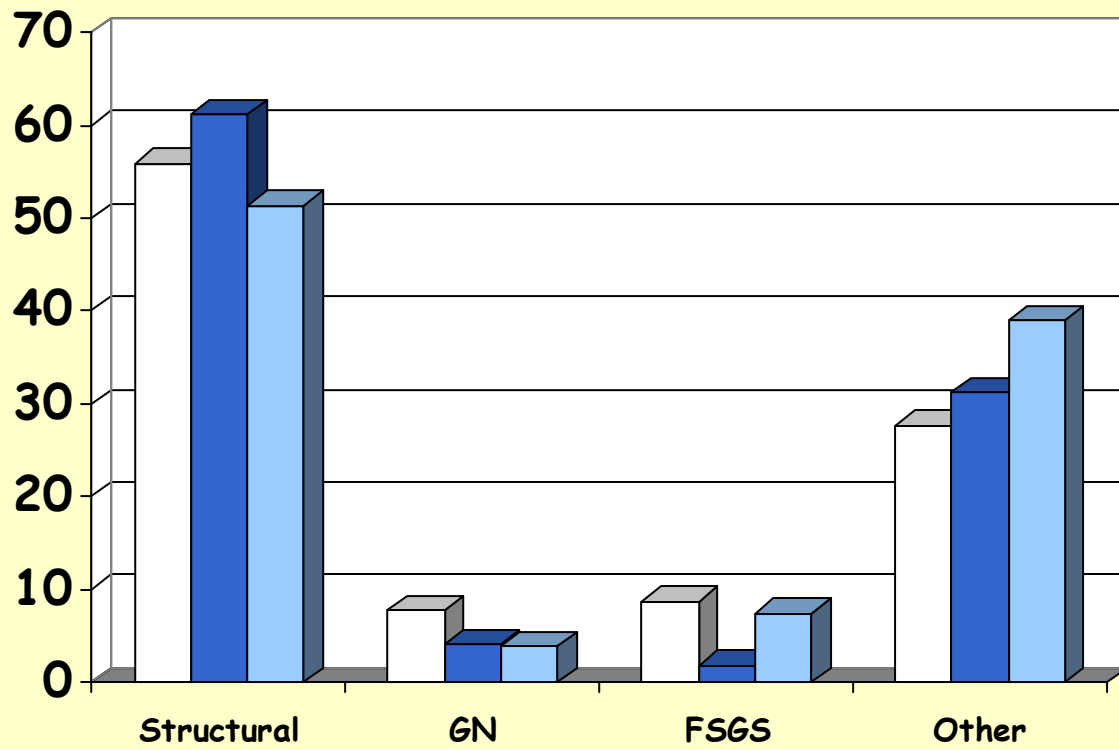
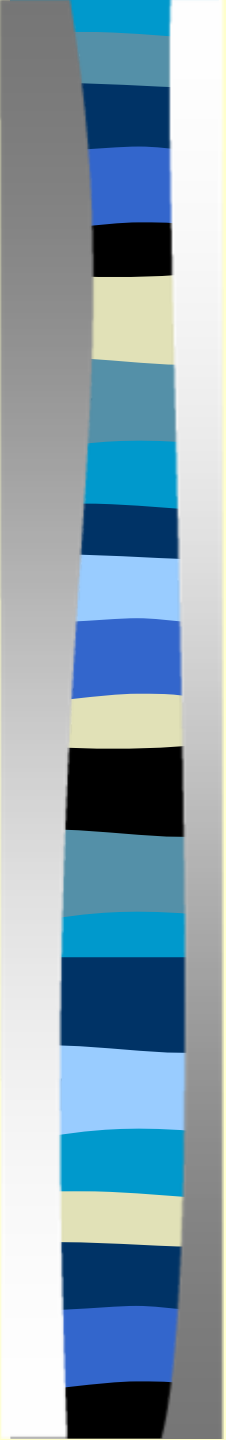
Etiology	Number	Percent	Male/female
Urological problems/tubulointerstitial diseases	143	50.7	85/58
Vesicoureteral reflux	52	18.5	26/26
Obstructive uropathy	30	10.7	26/4 ^a
Neurogenic bladder	43	15.2	22/21
Chronic pyelonephritis	7	2.2	4/3
Nephrolithiasis + chronic pyelonephritis	8	2.6	6/2
Tubulo-interstitial nephritis	3	1.0	1/2
Primary glomerulonephritis	32	11.4	19/13
FSGS	21	7.4	11/10
Others	11	4.0	8/3
Cystic renal diseases	35	12.6	18/17
Autosomal recessive PKD	8	3.0	6/2
Juvenile nephronophthisis	20	7.0	10/10
Multicystic-dysplastic kidney	5	1.9	2/3
Other	2	0.7	0/2
Secondary glomerulopathies	13	4.6	4/9
Amyloidosis	6	2.1	1/5
SLE	1	0.4	0/1
HUS	5	1.7	3/2
Other	1	0.4	0/1
Hereditary/metabolic diseases	14	4.9	10/4
Alport disease	6	2.1	4/2
Cystinosis	4	1.4	4/0
Primary hyperoxaluria type 1	1	0.4	0/1
Other	3	1.0	2/1
Hypoplasia/dysplasia/Aplasia	20	7.0	9/10
Vasculitis (HSP)	1	0.4	1/0
Renal neoplasms	1	0.4	1/0
Various syndromic cases	9	3.3	5/4
Unknown	22	7.8	11/11

^aThe only significant difference for gender was detected for obstructive uropathy ($P < 0.05$)

Causes of CKD



■ Bek et al. Turkish SPN CKDWG. *Pediatr Nephrol* 2009; 24:797-806.





Summary

- Leading causes:
 - CAKUT,
 - Other congenital and familial diseases.
- Factors affecting the pattern
 - Race, ethnicity, environment, wealth, age, time
- Preventable, curable in certain cases
- Measures to slow down progression