

PREIMPLANTATION BIOPSY – DECEASED DONOR TRANSPLANTATION



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INTRODUCTION



- **Marginal cadaveric kidney donors** can be **defined** as:
All donors older than 60 years, donors older than 50 years with any of the following criteria:
 - (1) Hypertension,
 - (2) Cerebro -vascular cause of brain death or
 - (3) Pre-retrieval serum creatinine (SCr) level > 1.5 mg/dl, with a degree of glomerulosclerosis $> 15\%$ and prolonged cold ischemia.
- **Pre implantation deceased donor kidney biopsy** helps in utilizing organs from marginal donors and predict long term outcome.

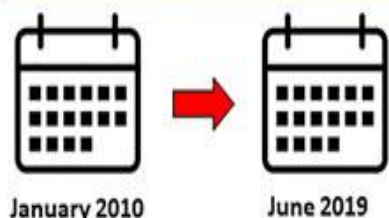
SCORING SYSTEMS



- Remuzzi protocol.
- Marryland Aggregate Pathology Index[MAPI].
- BANFF scoring system.
- Leuven scoring system.
- **Core** biopsy vs **wedge** biopsy – no much difference in scoring outcomes.
- Expert renal pathologist showed better interpretation of biopsies compared to routine pathologist.
- Stains used **Hematoxylin and Eosin** stain.
- Also **periodic acid–Schiff and Masson's trichrome** - useful in evaluating parameters such as hyaline arteriosclerosis, tubular atrophy, and interstitial fibrosis.

Evaluation of expanded criteria donors using kidney profile donor index and preimplantation renal biopsy

Material and methods



Kidney transplant (KT) recipients who received an allograft from a expanded criteria donor (ECD)

KDPI was calculated and results were stratified in quartiles

- Q1=86% (n=66)
- Q2=95% (n=72)
- Q3=99% (n=83)
- Q4=100% (n=45)

OPTN
ORGAN PROCUREMENT AND
TRANSPLANTATION NETWORK

And a preimplantation biopsy was performed



Glomeruli sclerosed *
Hyaline arteriopathy *
Fibrous thickening of the vascular intima *
Tubular atrophy *
Interstitial fibrosis *

Score 0-3 points *

Accepted for KT until a global SCORE of 7 points

Results

N= 266 KT recipients; median follow-up: 46 months



KT from KDPI Q1 presented better survival (log rank; $p=0.003$) and Q4 had worse renal function.



Biopsy score >3 was related to shorter survival (log rank; $p=0.018$) and poorer renal function.



Q1 donors with biopsy score ≤ 3 had the best survival (log rank; $p=0.014$).

Hyaline arteriopathy, glomerulosclerosis, and KDPI Q4 were predictors for graft survival



HR: 2.136 ($p=0.047$)



HR: 2.614 ($p=0.016$)



Q4 HR: 4.767 ($p=0.029$)

Conclusions: KDPI and a greater histological injury in biopsy, especially glomerular and vascular lesions, were related to a higher rate of KT graft from ECD.

Remuzzi et al scoring system

SCORING SYSTEM

Glomerular global sclerosis

- 0 = none
- 1+ = <20%
- 2+ = 20 to 50%
- 3+ = >50%

Tubular atrophy

- 0 = absent
- 1+ = <20% of tubuli affected
- 2+ = 20 to 50%
- 3+ = >50%

Interstitial fibrosis

- 0 = absent
- 1+ = <20% replacement by fibrous tissue
- 2+ = 20 to 50%
- 3+ = >50%

Arterial and arteriolar narrowing

- 0 = absent
- 1+ = increased wall thickness less than diameter of the lumen
- 2+ = wall thickness equal or slightly greater than diameter of the lumen
- 3+ = wall thickness far exceeds the diameter of the lumen

MARGINAL KIDNEY DONOR

- Age >60 yr
- History of diabetes or hypertension
- Clinical proteinuria (≤ 3 g/24 h)

Macroscopic evaluation
of the kidneys

Major vascular, ureteral, or
parenchymal abnormalities

Discard

No abnormalities

Pre-transplant biopsy
of both kidneys

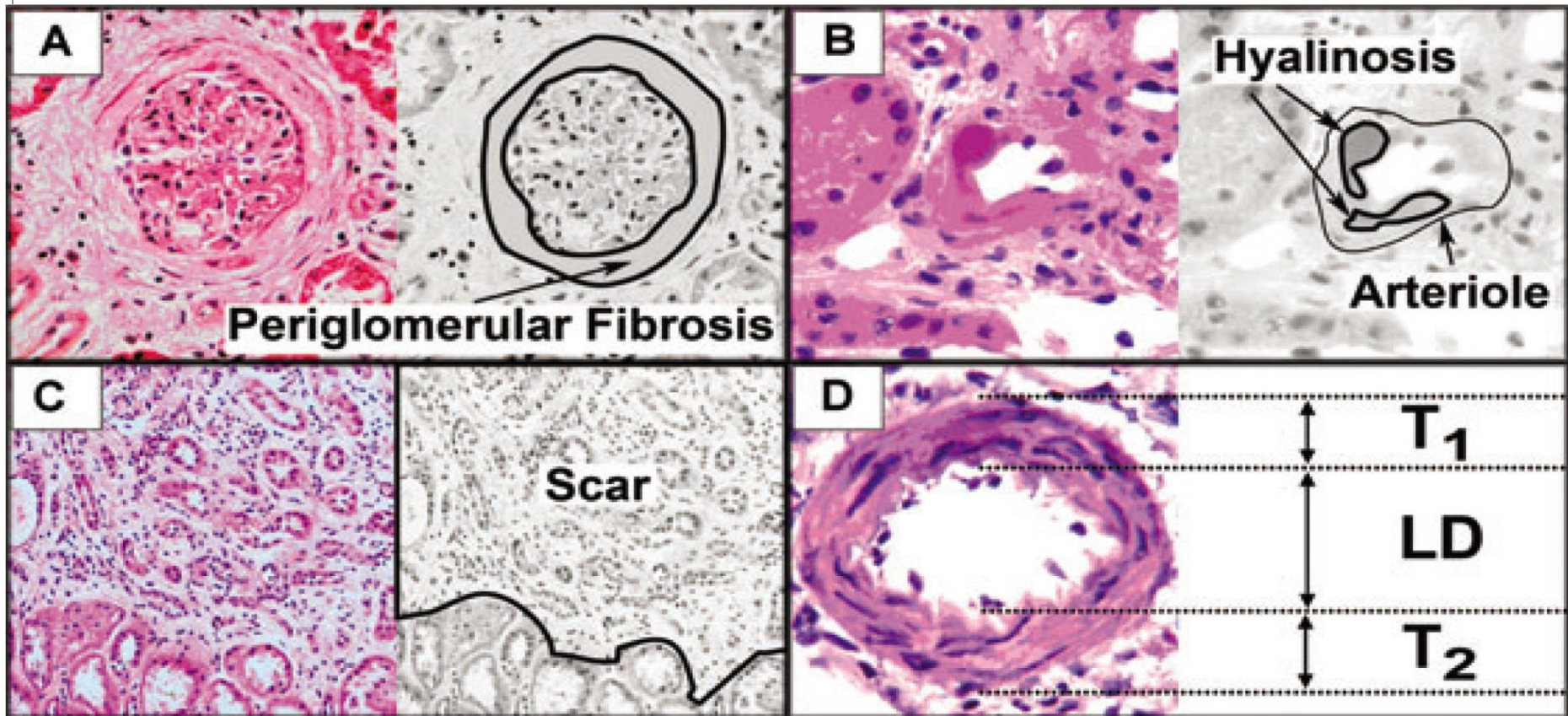
Eligibility for double or single
transplant according to
a standardized score system

Final Grading

- 0 to 3 mild → *single*
- 4 to 6 moderate → *double*
- 7 to 12 severe → *discard*

The Maryland Aggregate Pathology Index: A Deceased Donor Kidney Biopsy Scoring System for Predicting Graft Failure

Pathologic features used in MAPI as seen on frozen section preparations. (A) **Periglomerular fibrosis**, (B) **arteriolar hyalinosis**, (C) **scar** including features of interstitial fibrosis, tubular atrophy and glomerulosclerosis, and (D) measurements for **arterial wall-to-lumen ratio** (WLR) calculation, including the thickness of two opposing walls (T_1 and T_2) and the luminal diameter (LD). $WLR = (T_1 + T_2) / LD$



OUR EXPERIENCE

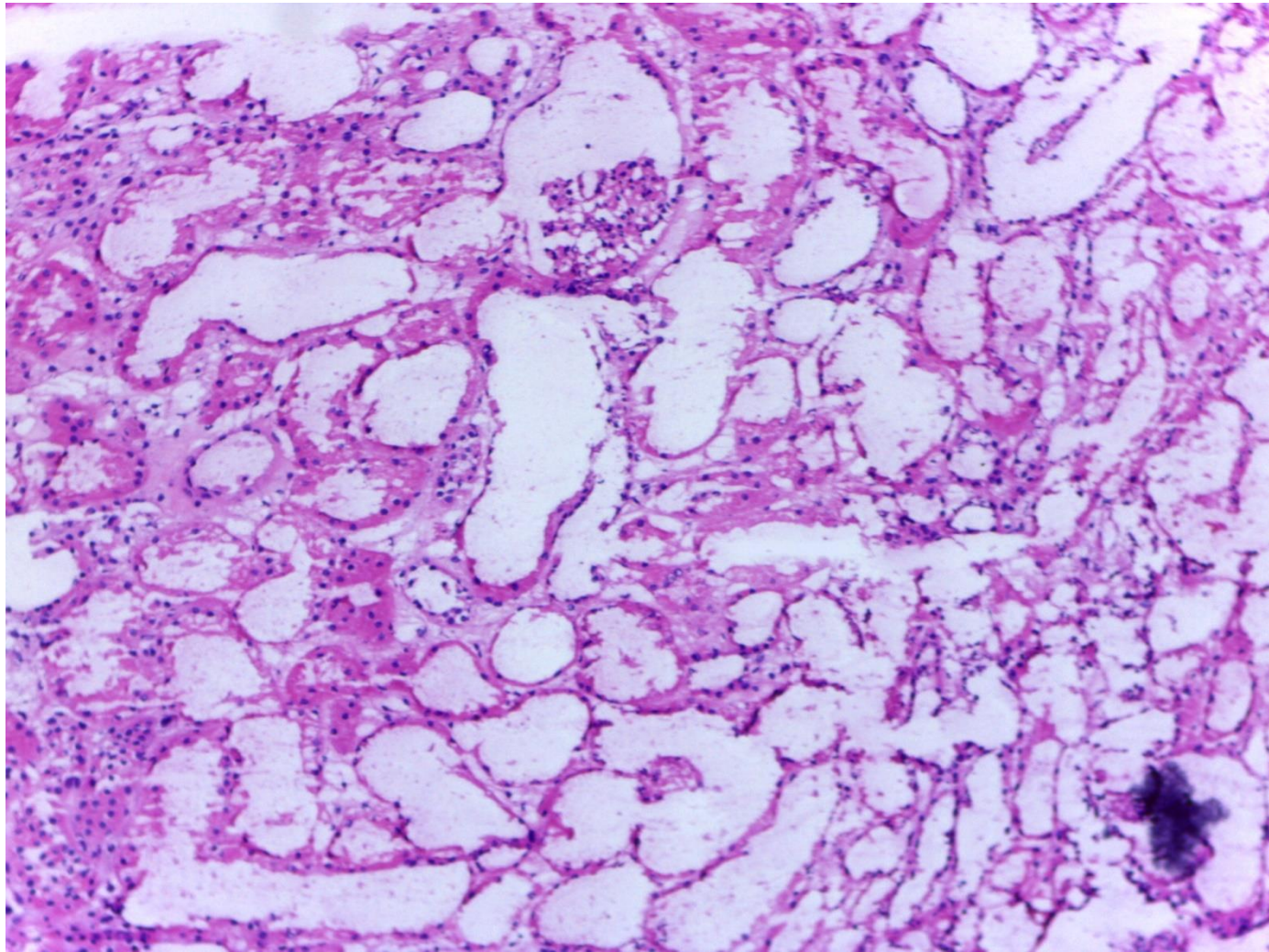


- 20 preimplantation biopsies done out of 66 cadaver transplants done.
- 4 donor kidneys rejected : 1 had extensive intra glomerular thrombi ,1 case of glomerular cystic disease and 2 cases of increased IFTA.
- Preimplantation biopsy helped procuring the organ which would have otherwise been rejected and is an important tool for expanding donor pool.

CASE 1



- 18 year old deceased donor with no comorbidities and death due to road traffic accident.
- His creatinine - 1.4 mg/dl and all other biochemical parameters were within normal limits.
- His urine routine showed protein 1+ and no RBCs. He had no history of hypertension or renal disease in family. He was declared brain dead after 48 hours of admission.
- On initial outlook he seemed to be a fair donor who is young with no comorbidities and well managed by ICU team.
- Since he had a baseline high creatinine for his age [Egfr -66 ml per min per 1.73 m²] we decided to do preimplantation biopsy.
- His biopsy showed **glomerulocystic changes** in all visualised glomeruli and therefore the kidney was discarded.
- Biopsy helped us from not transplanting a suboptimal kidney.

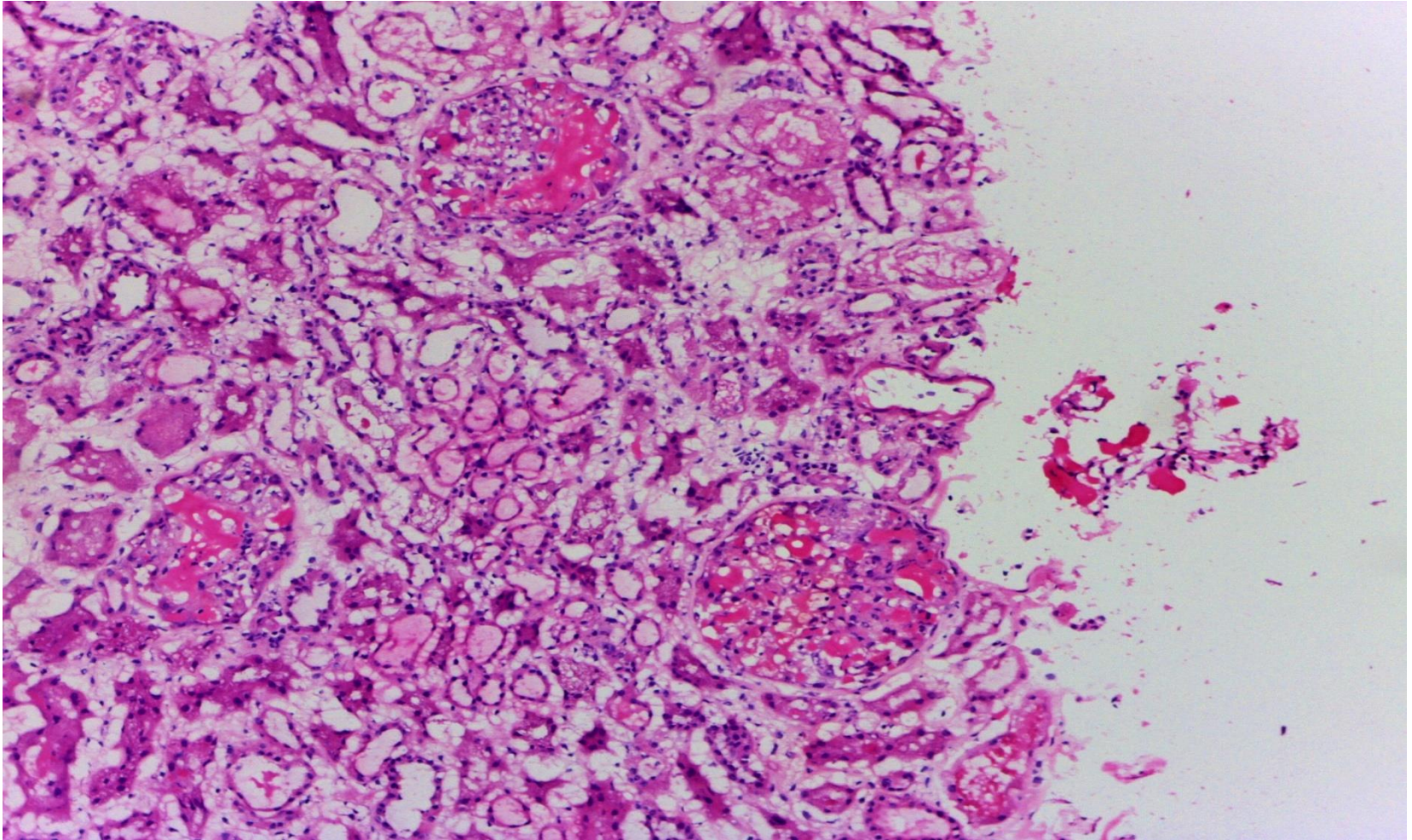


Case 1 -Glomerular cystic changes on preimplantation biopsy

Case 2



- 46/year old female deceased donor with history of hypertension and death due to road traffic accident.
- Her terminal creatinine was 1.2 mg /dl. Her urine report showed protein 2+ and few RBCs.
- Her preimplantation biopsy showed extensive intra glomerular thrombi.



Preimplantation biopsy showing extensive intra glomerular thrombi

Case 2 continued



- **Intra glomerular thrombus** is commonly seen in preimplantation biopsies.
- If present in majority of glomeruli, kidney is discarded because it causes **delayed graft function** and raises the possibility of glomerular disease of unknown etiology in the donor.
- **Recently many papers** were published on implanted kidneys with intraglomerular thrombi and follow up protocol biopsies showing resolution of thrombi early in post-transplant period and minimal residual changes in some, at the end of one year with good graft function.

Case 3



- A male deceased donor aged 49 years, was a known case of diabetes mellitus for 10 years with terminal creatinine 1.1 mg/dl and urine showing 1+protein.
- His preimplantation biopsy showed interstitial fibrosis and tubular atrophy **involving less than 10%** with Remuzzi histological score of 2.
- In view of the above findings, the donor kidney was taken up for transplant and had good graft outcome with post-transplant creatinine of 1.2 mg/dl.

Case 3 continued



- Comorbidities like **diabetes and hypertension** with urinary abnormalities usually put the transplant team in dilemma whether to take or discard the marginal kidney.
- Long standing history of diabetes mellitus in deceased donor with proteinuria is out rightly rejected on clinical criteria.
- Case 3 patient had **long standing diabetes mellitus but it did not translate to histological injury** and hence donor kidney was selected.