PREGNANCY – AKI [PRAKI]

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INTRODUCTION

- Pregnancy-related acute kidney injury (Pr-AKI) is a heterogeneous disease entity due to varying underlying etiologies.
- P-AKI in developing countries accounts for 5%–20% of total AKI population.
- Pr-AKI commonly occur in postpartum rather than the post- abortal period, reflecting a decline in septic abortions.
- 24 hr creatinine clearance is closest estimate of GFR in pregnancy.
- Diagnosis of AKI in pregnant women (any one of the three)
- (1) Sudden increase in serum creatinine >1 mg/dl,
- (2) Oliguria/anuria
- (3) Need for dialysis.

Anatomical and structural changes ↑ Size of the kidneys Pelvicalyceal dilatation R>L Hydronephrosis R>L ↑glomerular basement membrane (GBM) pore size and electrical charges Afferent and efferent arterioles dilatation Urinary stasis

Endocrine function

↑RAAS activity

↑NO production ↑erythropoietin

↑Renin activation

↑Vitamin D (active form, Calcitriol)

↓AT1R and sensitivity to ANG II
 ↑vasodilatory peptides (Ang 1-7) of RAAS

Renal haemodynamics

↑RBF by 50 - 85%

Glomerular function
↑GFR by 50%
↓plasma creatinine
↑proteinuria ("physiologic"

(less than 260 mg/24H)

↓osmostat threshold for ADH release

↓plasma osmolality by 10mOsm/kg

↓serum levels of urea and uric acid ↑bicarbonaturia (metabolic acidosis)

proteinuria)

Tubular function √sensitivity to RAAS

↑water retention

Hyponatremia (mild)

**Calciuria

**Glycosuria

↓RVR

↑RPF ↓FF

PHYSIOLOGICAL CHANGES IN PREGNANACY

CAUSES OF P-AKI

PRERENAL

Transient decrease in renal perfusion → VGFR

- > Intravascular volume depletion = hypovolemia
 - Hyperemesis gravidarum
 - · OHSS
 - Diarrhea
 - Blood loss

Miscarriage

Antepartum haemorrhage

Postpartum haemorrhage

- · Diuretics (renal loss from overuse)
- Decreased arterial pressure/Decrease effective circulating volume

Sepsis postabortal, puerperal, urosepsis

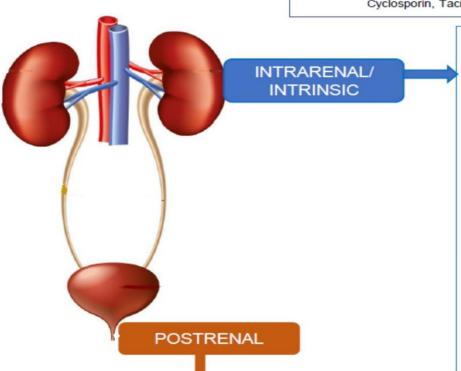
Heart failure

Impaired renal autoregulation/intrarenal vasoconstriction

NSAIDs

ACE inhibitors, ARB

Cyclosporin, Tacrolimus



Glomerulonephritis

Acute glomerulonephritis

Interstitial

- Sepsis/Infection postabortal, puerperal, urosepsis, pyelonephritis
- ▶ Ischemia
 - · miscarriage/abortion

extrarenal loss

- antepartum haemorrhage-placenta praevia, abruptio placentae, uterine rupture
- · postpartum haemorrhage- atonia, perineal tears and lacerations

Vascular

- vasculitis
- hypertension Preeclampsia/HELLP
- thrombotic microangiopathy (TMA) thrombotic thrombocytopenic purpura (TTP) atypical haemolytic uremic syndrome (aHUS) disseminated intravascular coagulation (DIC)

Medication - nephrotoxicity, acute interstitial nephritis antibiotic aminoglycosides, iodinated contrast

Acute fatty liver of pregnancy (AFLP)

Lupus nephritis

Antiphospholipid syndrome (APS)

Pulmonary embolism

Amniotic fluid embolism

Bilateral obstruction and hydronephrosis due to uterine compression of ureters/bladder Unilateral ureter obstruction due to nephrolithiasis Bladder outlet obstruction (tumor, lithiasis)

latrogenic injury of the bladder/ureter/urethra during caesarean section or vaginal delivery Spontaneous injury of the urethra/bladder during vaginal delivery

1st trimester 2nd trimester 3rd trimester **D** Postpartum

Septic Abortion

Clinical Features

➤ Prerenal azotemia or ATN

Treatment

➤Volume resuscitation, antibiotics

Hyperemesis Gravidarum Clinical Features

➤Prerenal azotemia or ATN

➤ Evaluate for molar pregnancy

Treatment

>Volume resuscitation

Preeclampsia/ HELLP

Clinical Features

➤ Hypertension and proteinuria* after 20 weeks of gestation

>Headache, visual disturbance, seizures, abdominal pain

>Hemolytic anemia, transaminitis, thrombocytopenia, high LDH

Treatment

Delivery, i.v. magnesium for seizure prevention

TTP/aHUS

Clinical Features

>TTP more common in 2nd/3rd trimester, aHUS more common in postpartum period

Neurological involvement is more common in TTP than aHUS

>Hemolytic anemia, thrombocytopenia, elevated LDH and bilirubin

Laboratory testing

>TTP: ADAMTS-13 activity <10%

>aHUS: genetic testing for complement cascade gene mutations

Treatment

>TTP: plasma exchange

>aHUS: plasma exchange + eculizumab

Acute Fatty Liver of Pregnancy

Clinical Features

Nausea, vomiting, abdominal pain, jaundice, ascites

>Transaminitis, low platelets, hypoglycemia, lactic acidosis

Laboratory Testing

> Maternal and fetal testing for LCHAD gene mutation

Treatment

>Delivery, plasmapheresis and/or liver transplant in severe cases

Lupus nephritis and/or Antiphospholipid antibody syndrome

Clinical Features

- >Dysmorphic red blood cells on urine sediment, extra-renal lupus manifestations
- >Low complements, positive anti-dsDNA, anti-cardiolipin antibodies, and/or anti-β2 glycoprotein antibodies
- >Kidney biopsy is only recommended if pathology will change management

Treatment

- Lupus nephritis: steroids + hydroxychloroquine + azathioprine/tacrolimus
- > Antiphospholipid antibody syndrome: aspirin +/- low molecular weight heparin

PREECLAMPSIA

- Definition : Characterized by
- √ New-onset hypertension (blood pressure >140/90 mmHg)
- ✓ Proteinuria (>300 mg/dl) after 20 weeks of gestation.
- Eclampsia is defined as preeclampsia with the presence of seizure.

Table 5: Severe feature of preeclampsia (one or more of these findings)

- Systolic blood pressure \geq 160 mmHg or diastolic blood pressure \geq 110 mmHg on two occasions at \geq 4 h apart while the patient is on bed rest
- Platelet count <100,000/mm³
- Elevated liver enzymes (twice normal concentrations)
- Renal insufficiency (serum creatinine concentration >1.1 mg/dl or doubling of serum creatinine concentration) or oliguria (<500 ml in 24 h)
- Pulmonary edema or cyanosis
- New-onset cerebral or visual disturbances
- Severe persistent right upper quadrant or epigastric pain

Clinical feature	HELLP*	AFLP**	aHUS***	TTP
Time of onset	3T	3T	Postpartum	2T/3T
Hypertension	80%-100%	25%-50%	+	0/+
AKI	Mild/moderate	Moderate	Severe	Mild/moderate
Renal prognosis	Recovery	Recovery	76% ESRD	Fair
CNS findings	+	Absent	Absent	Dominant
Hemolytic anemia	+	0/+	+	++
Thrombocytopenia	+	0/+	++	++
Coagulopathy	0/+	+	0	0
Liver transminases increase	++	++	0	0
LDH (IU/L)	+	0/+	++	++
Ammonia	Normal	High	Normal	Normal
ADAMTS-13 activity <10%	0	0	+	++
Alternative complement pathway	0/+	0/+	++	0
Management	Support measures/ delivery	Support measures/ delivery	Plasma infusion/ exchange	Plasma infusion/ exchange
Effect of delivery on diseases	Recovery	Recovery	None	None
*Urine sediment is bland in preeclamp AFLP, ***Isolated LDH increase with present, +: Sometimes present, ++: Al- liver of pregnancy, aHUS: Atypical he CNS: Central nervous system, LDH: I	normal hepatic transamina ways present, HELLP: Her molytic-uremic syndrome,	ase is characteristics of HUS nolysis, elevated liver enzy TTP: Thrombotic thrombo	S/TTP (P-TMA). 0: Absenmes, and low platelet cour cytopenic purpura, AKI: A	nce, 0/+: Occasionally nt, AFLP: Acute fatty Acute kidney injury,

pregnancy

RENAL CORTICAL NECROSIS

- Rare condition resulting from severe reduction of renal perfusion caused by vascular spasm, microvascular injury, or intravascular coagulation.
- Abrupt onset of oliguria, gross hematuria, flank pain and hypotension.
- Causes: Placental abruption, placenta previa, septic abortion. prolonged intrauterine fetal death and amniotic fluid embolism.
- Hypercoagulable state with increased level of coagulation factors with repressed fibrinolytic state.
- Prognostic factors: extent of necrosis, duration of oliguria, and severity of associated conditions.
- **Ultrasound**: Hypoechoic Rim and Renal Cortical Rim Sign: The hypoechoic rim seen on ultrasound, potentially with an echogenic (brighter) layer outside the rim.
- CT Scan: Lack of Cortical Enhancement and Reverse Rim Sign: A specific CT finding where the cortex appears hypo attenuating (darker) and non-enhancing, while the medulla enhances.
- Dialysis as indicated and treatment of the underlying diseases.

HD Prescription during Pregnancy

Frequency	5–6 time/week
Duration of dialysis	>36 h/week (>6 h/day)
Dry weight	Increase by 0.5 kg/week during the second and third trimesters
UF rate	6–8 mL/kg/h
Dialysate	K = 3 mmol/L, Ca = 1.5 mmol/L, HCO3 = 28–32 mmol/L
Anticoagulation	Low-dose unfractionated heparin

HD: hemodialysis; UF: ultrafiltration.