





# Enlightening the Path for Next Generation of Gynaecologists

*Dedicated Issue:* Medical Disorders in Pregnancy: An algorithmic approach



## **AOGD SECRETARIAT**

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# AOGD Bulletin

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# From the President's Pen



Dear Friends, Greetings from AOGD

This deciated issue of AOGD Monthly Bulletin on 'Medical Disorders in Pregnancy" is in your hand. We all strive to achieve a Healthy and Safe Motherhood but on occasions pregnancy is complicated by disorder, which compromise maternal outcome and affects various statistics associated with child birth. In India we have made tremendous achievement in reducing maternal mortality rate to current level of 134/100000 live birth. Such a thing has been possible with active efforts of all of you. To have latest knowledge about various disorders that will help improve outcomes further we present this issue of AOGD Bulletin to you.

Happy Reading

Dr Sunesh Kumar President, AOGD

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**AOGD Bulletin** 

# From the Secretary's Desk



Dear Friends, Greetings from AOGD.

The November issue of AOGD bulletin dedicated to contraception has been received with great enthusiasm. I thank one and all for it.

Continuing with the theme of "Enlightening the Path of Next Generation of Gynaecologists", the present issue is dedicated to Medical disorders in Pregnancy. In this issue of the bulletin, we intend to provide management algorithms and practice points for the various medical disorders in pregnancy as easy reference for practitioners and students at large. As the algorithms incorporates current guidelines, it will prove to be very useful for all generations of obstetricians.

This was a busy month that saw many conferences and CME being organized.

A conference on Aesthetic Gynaecology was organized on 9<sup>th</sup> & 10<sup>th</sup> November, 2019 by Indian Academy of Cosmetic Dermatology & Gynecology Under the aegis of AOGD & FOGSI.

An Update in Gynaecologic Oncology was organized on 13<sup>th</sup> November, 2019 under the aegis of Oncology Committee of AOGD at AIIMS, New Delhi

A CME on "Tackling Obstetric Dilemmas" was organized on 16<sup>th</sup> November, 2019 at The Surya under the aegis of FOGSI Medical Disorders in Pregnancy Committee & AOGD.

Qyality improvement Workshop was organized on 16<sup>th</sup> November, 2019 at MAMC & LNJP Hospital under the aegis of Quality Improvement Sub-committee AOGD.

A CME on "High Risk Pregnancy" on 16<sup>th</sup> November 2019, was organized under aegis of Safe Motherhood Committee AOGD at ESI Basaidarapur.

Breast Cancer screening Health Camp was organized on 19<sup>th</sup> November, 2019 by GTB under the aegis of Rural Health Committee, AOGD.

NARCHI pre congress workshop on "PPH" on conducted on 22<sup>nd</sup> November under aegis of Safe Motherhood Committee AOGD at the Northern Railway Hospital.

A CME was organized under the aegis of DGF North, Breast and Cervical cancer awareness screening & prevention subcommittee AOGD & Breast committee FOGSI on 27<sup>th</sup> November, 2019.

We look forward to your continued support.

I take this opportunity to wish a Happy, Healthy and Prosperous 2020 Warm Regards

Dr Vatsla Dadhwal

Hon. Secretary

# **Monthly Clinical Meeting**

Monthly Clinical Meet will be held at Sir Ganga Ram Hospital, New Delhi on **Friday**, **27<sup>th</sup> December**, **2019 from 04:00pm to 05:00pm**.

# From the Editor's Desk



Dr J B Sharma Editor



Dr Reeta Mahey



Dr P Vanamail



Dr Vidushi Kulshreshtha





Dr Aparna Sharma — Guest Editors —



Dr Archana

We are pleased to write from the editorial desk for this issue of AOGD Bulletin on the very special topic of 'Medical Disorders in Pregnancy'. The previous issue on 'contraception' was very much appreciated by our fellow AOGD members. With increasing age at conception, increasing obesity and better optimization of medical disorders, more women with medical disorders are now presenting with complex scenarios during pregnancy.

We have invited a group of eminent obstetricians to author articles that are both cutting edge and pertinent to changing obstetric practice. Our issue begins with an important article "Anemia in pregnancy" which is an important indirect cause of maternal mortality, an excellent starting point written by Dr JB Sharma and Dr Neha Varun. The survey on hypertensive disorders is a current, concise single reference for management of all hypertension during gestation. Comprehensive information on management of hypertensive disorders in pregnancy as well as the newer protocols for screening for preeclampsia have been discussed by Dr Aparna Sharma and Dr Archana Kumari. This is followed by an algorithm on HIV in pregnancy, a problem still in epidemic proportions in India written by Dr Rinchen Zangmo and Dr Rohini Sehgal. The next articles are related to medical autoimmune disorders like antiphospholipid antibody syndrome and SLE in pregnancy authored here by Dr Jyoti Meena, Dr Kusumlata and Dr Akansha Tiwari. We then shift focus to more well-known medical disorders, including a renewed assessment of pregnant woman with heart disease written by Dr Jyotsna Suri and Dr Zeba Khanam. The current reports on both gestational and pregestational diabetes highlighting recommendations on diagnosis and management has been prepared by Dr Garima Kachhawa and Dr Richa Vatsa. The management of pregnant women with chronic kidney disease has been elucidated beautifully by Dr Juhi Bharti and Dr Deepali Garg.

There is an article on liver disease in pregnancy authored by Dr Sharda Patra who has worked extensively in the field and an update on immunization in pregnancy written by Dr Vidushi Kulshrestha and Dr Soniya Dhiman. An algorithmic approach to Thyroid disease in pregnancy includes the most recent recommendations on antepartum and postpartum management by Dr Anubhuti and Dr Vatsla Dadhwal.

The opportunity to edit this issue of AOGD has been challenging, rewarding, and a learning experience. We hope you will find these articles as interesting and valuable as we have.

We wish our esteemed readers a happy reading and shall welcome their comments and contributions to further improve the bulletin.

#### **Editorial Team**

# Anemia in Pregnancy: Diagnosis and management

#### J B Sharma<sup>1</sup>, Neha Varun<sup>2</sup>

<sup>1</sup>Professor, <sup>2</sup>Assisstant Professor, Department of Obstetrics and Gynaecology, All India Institute of Medical Sciences, New Delhi

Anemia is defined as the reduction in the total circulating red cell mass below normal limit. It is the most commonly seen medical disorder during pregnancy. Complete blood count testing is the recommended method for diagnosing anemia during pregnancy.

In the settings where complete blood count testing is not available, onsite hemoglobin testing with a haemoglobinometer is recommended over hemoglobin color scale method use, for diagnosing anemia in pregnancy.<sup>[1]</sup>

# Management



#### **IRON DEFICIENCY ANEMIA**

#### HISTORY

#### EXAMINATION

**PRESENT HISTORY**: H/O bleeding PV, passage of worms in stool, jaundice, easy bruisability / petechiae, hematuria/ bleeding PR/epistaxis, chronic fever, IFA intake, breathlessness, pedal edema, palpitation, pica /pagophagia

**PAST HISTORY**: Chronic medical/ surgical illness, bleeding disorder, recurrent UTI, fever, blood transfusions

**FAMILY HISTORY**: hemoglobinopathy, Repeated blood transfusion

**PERSONAL HISTORY**: Dietary history, malabsorption, socioeconomic status

**SYMPTOMS**: Easy fatiguability, weakness, anorexia, palpitation, giddiness, swelling of legs, pica, pagophagia, restless leg syndrome

SIGNS: Pallor, glossitis, cheilosis, koilonychia, plummer-Vinson syndrome, tachycardia, pedal edema, ejection systolic murmur in mitral area

### INVESTIGATION

- Severity of anemia
- Typing of anemia
- Assessment of iron stores
- · Assessment of heme iron
- Assessment of iron
   absorption

# Indications of bone marrow biopsy

- Failure of therapy
- Kala azar
- Hypoplastic or aplastic anemia

#### DIETARY MEASURES

- Consumption of iron rich foods
- Cooking food in iron utensils

#### DEWORMING

mg stat to all after 1st trimester WHO 2017: Anti-helminthic to all where prevalence is >20%

GOI 2018: Tab albendazole 400

TREATMENT

- 1) ORAL IRON\*
- 2) PARENTERAL IRON #
- 3) BLOOD TRANSFUSION<sup>\$</sup>

#### **RESPONSE TO THERAPY**

- 1) Improvement in symptoms
- 2) Improved appetite
- **3)** Increase in reticulocyte count after 7-10 days
- 4) Rise in Hb after 2 weeks (0.8-1 g/dl/week)
- 5) Hb reaches normal levels by 6-8 weeks

If no significant improvement seen in 3 weeks, diagnostic re-evaluation needed

#### **\*Oral Iron supplementation** (GOI 2108)

- 1) Prophylactic- elemental Fe 60mg OD+ folic acid 500 mcg OD till term-min 180 days
- 2) Therapeutic: Mild-mod anemia-2 IFA tab/day
- 3) Postpartum: elemental Fe 60mg OD+ folic acid 500 mcg OD for 6 months

#### **# PARENTRAL Fe THERAPY:**

- 1) INDICATIONS: Intolerance to oral iron, no response to oral iron, advanced pregnancy, severe anemia, patients on EPO
- 2) CONTRAINDICATIONS: Iron overload conditions, hypersensitivity to IV Iron. 1st trimester of pregnancy, active infection, hepatic dysfunction (OT/PT > twice)

#### **CALCULATION OF IRON DOSE:**

REQUIRED IRON DOSE (mg): 2.4 (target Hb-actual Hb) pre-pregnancy weight (kg) + 1000 mg

#### **COMMONLY USED PREPARATIONS:**

- Before giving parenteral Fe ( I/V or I/M), test dose is to be given with emergency drugs and equipment's to be ready for any anaphylactic reaction.
- 1) Iron sucrose complex: Each ml contains 20 mg of elemental iron. 200mg after dilution in 100 ml of 0.9% NaCl as a slow infusion over 15-20 minutes can be given on alternate days and maximum 3 doses /week.
- 2) Ferric carboxy-maltose: Maximum single dose 1000 mg (20 ml) diluted in 250 ml of sterile 0.9% NaCl over 15-20 min not more than once a week and single injection not exceeding 20mg of iron/ Kg body weight.

#### **\$ BLOOD TRANSFUSION (WHO OBSTETRIC MANUAL 2017)**

#### 1) Pregnancy < 36 weeks POG:

- Hb < 5 g/dL
- Hb 5-7 g/dL with presence of 1 or more of following conditions: Established or incipient cardiac failure or clinical evidence of hypoxia, pneumonia or any other serious bacterial infection, malaria, pre-existing heart disease, not causally related to the anemia
- 2) Pregnancy  $\geq$  36 weeks:
  - Hb < 6 g/dL
  - Hb 6-8 g/dL with presence of one or more of following conditions: Established or incipient cardiac failure or clinical evidence of hypoxia, pneumonia or any other serious bacterial infection, malaria, pre-existing heart disease, not causally related to the anemia
- 3) Other indications of blood transfusion in pregnancy:
  - Acute hemorrhage with Hb < 6 g/dl
  - · Anemia not due to hematinic deficiency
  - Intrapartum: if Hb is < 7g/dl in labor





# **Key Points**

- 1) India is categorized as high prevalence area for anemia in pregnancy
- 2) Nutritional deficiency anemia during pregnancy continues to be a major health problem in India.
- 3) During pregnancy iron and folate prophylaxis is recommended
- 4) Complete work up and diagnosis of anemia during pregnancy should be made before initiating any therapy
- 5) First line therapy for iron deficiency anemia is oral iron

6) Parenteral iron therapy is safe and should be considered for moderate to severe anemia

#### References

- 1) WHO recommendation on the method for diagnosing anemia in pregnancy. 08 March 2018
- 2) CDC Issues Guidelines for Prevention, Detection and Treatment of Iron Deficiency. Am Fam Physician. 1998 Oct 15; 58(6): 1475-1477
- 3) Am Fam Physician. 2017 Sep 15;96(6):384-389

# **Screening & Management of Preeclampsia and Eclampsia**

#### Aparna Sharma<sup>1</sup>, Archana Kumari<sup>2</sup>

<sup>1</sup>Additional Professor, <sup>2</sup>Assisstant Professor, Department of Obstetrics and Gynaecology, AIIMS, New Delhi



#### <sup>1</sup>Definition:

- Pregnant women  $\geq 20$  weeks
- SBP  $\geq$  140 mmHg and / or DBP  $\geq$  90 mmHg (on two separate occasions taken 4 hours apart)
- SBP $\geq$ 160mH or DBP $\geq$ 110
- Proteinuria 2+ or more (may be absent)

In absence of proteinuria, new onset hypertension with new onset of any of the followings:

- Headache (not relieved by regular analgesics or accounted by alternative diagnosis or visual symptoms)
- Thrombocytopenia <11ac
- Raised liver function tests (twice the normal)
- Renal insufficiency (S.cr.>1.1mg/dl or doubling in absence of other renal ds
- Pulmonary edema

#### <sup>2</sup>Symptoms of Severe Preeclampsia SBP≥160mH or DBP ≥ 110

- Proteinuria 2+ or more (may be absent) OR
- Headache (not relieved by regular analgesics)
- · Blurred vision
- Upper abdominal pain (Epigastric or right upper quadrant pain)
- Oliguria (Urine output <400ml in 24 hours)
- Pulmonary edema

#### <sup>3</sup>Clinical Assessment:

- Check PR, BP, RR, Temperature
- Secure IV line
- Catheterise for monitoring urine output
- Monitor sings & symptoms of impending eclampsia / pulmonary oedema
- Calculate gestational age
- · Check for fetal well being

#### <sup>4</sup>Investigations:

- Complete haemogram with platelet count with peripheral smear for haemolysis Blood urea nitrogen & creatinine
- Liver function tests
- Urine albumin by dipstick & 24 hours urine protein
- Fundus examination
- Ultrasound for fetal growth and well being if available

#### <sup>5</sup>Control BP:

(Keep DBP<110mmHg)

- If  $BP \geq 160/110 \ mmHg$
- Hydralazine 5 mg IV slowly over 5 minutes, repeat ever 20-40 mts (maximum 20 mg) OR
- IV labetalol 10-20 mg then 20-80 mg Every 10-30 mts (maximum 300 mg) OR
- tab nifedipine 10-20 mg, repeat in 20 mts if needed, then 10-20 mg every 2-6 hrly(maximum 180 mg/day)

#### Once BP is stabilised start oral antihypertensives:

- Tab. Labetelol 100 mg TDS
- (maximum 2400 mg) (First Line)
- OR
- Tab. Methyldopa 250 mg TDS (maximum 2grams)

#### <sup>6</sup>Contraindication to expectant management:

- · Impending eclampsia
- Eclampsia
- HELLP Syndrome
- Pulmonary edema
- DIC
- Renal dysfunction
- Abruption
- Gestational age  $\geq$  37 weeks
- Fetal death
- Non-reassuring fetal status
- · Previable gestation
- Persistent REDF in umbilical artery doppler

#### $^{7}MgSO_{4}$ for seizure prophylaxis :

#### Regimen # 1

Loading dose:

- 4 gram of 20% MgSO4 IV over 5 minute AND
- 10 gram of 50% MgSO4 deep IM with 1 ml of 20% lignocaine (5gram each buttock) Followed by
- 5 gram of 50% MgSO4 IM every 4 hour for 24 hour

#### Regimen # 2

 4 gram of 20% MgSO4 IV over 5 minutes followed by 1 gram/hour IV solution of 20% MgSO<sub>4</sub> for 24 hours

#### Monitoring

- Respiratory rate (≥ 16breath/minute) Urine output (>30ml/hour)
- Knee jerk (present)
- Contraindications of MgSO4-
- Mysathenia gravis, Hypocalcemia, moderate to severe renal failure, cardiac ischemia, heart block, myocarditis
- · Give diazepam or phenytoin

#### <sup>8</sup>Maternal assessment:

- Monitor every 8 hours Vital signs
- Input/Output
- Signs & Symptoms of impending eclampsia Presence of contractions and/or rupture of membranes, Bleeding
- Laboratory testing biweekly (Complete blood count, platelets counts, ALT, AST, Creatinine)

#### Fetal assessment:

- Daily kick count & NST
- Twice weekly Biophysical profile
- Serial growth studies
- \* Antenatal steroids should be given if the gestational age is between 24 weeks 34 weeks for lung maturity

#### <sup>9</sup>Severe maternal & fetal complications:

- Uncontrolled BP
- Pulmonary edema
- Eclampsia
- Signs & symptoms of impending eclampsia
- Abruption
- · Progressive labor
- · Non-reassuring fetal heart
- Fetal death
- Severe IUGR

#### **Timing of delivery**

Delivery is recommended when gestational hypertension or preeclampsia with severe features is diagnosed at or beyond 34 0/7 weeks of gestation, after maternal stabilization or with labor or prelabor rupture of membranes.



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#### **CONTINGENT SCREENING IN LIMITED RESOURCE SETTING**

In limited resource settings, routine screening for preterm preeclampsia should be done by maternal factors and MAP in all pregnancies and preserving PLGF and UTPI for the subgroup of population based on risk assessment by MAP and maternal risks as shown in algorithm below



### **Suggested Reading**

- ACOG guidelines for Preeclampsia and eclampsia 2019
- FIGO guidelines for Screening of Preeclampsia 2019

# **HIV Infection in Pregnancy**

#### Rinchen Zangmo<sup>1</sup>, Rohini Sehgal<sup>2</sup>

<sup>1</sup>Assistant Professor, <sup>2</sup>Scientist, Department of Obstetrics and Gynaecology AIIMS, New Delhi



#### **Intra Partum Management in Women with HIV Infection**

Status	Intra-partum	Post -partum
Presenting in active labour,	Initiate TDF (300 mg) + 3TC	Continue TDF (300 mg) + 3TC (300mg) +
No prior ART	(300 mg) + EFV (600 mg)	EFV (600 g)

NACO recommends normal vaginal delivery unless the woman has obstetric indications (like foetal distress, obstructed labour) for a Caesarean section.

# In places where facilities for viral load monitoring are available, intrapartum management should be done as per viral load (British HIV Association Guidelines 2018):

- Plasma viral load <50 HIV RNA copies/mL at 36 weeks, with no obstetric contraindications → planned vaginal delivery.
- Vaginal birth after CS (VBAC) offered to women with a viral load <50 HIV RNA copies/ml.
- Plasma viral load 50–399 HIV RNA copies/mL at 36 weeks. → pre-labour CS (PLCS) should be considered, taking into account the actual viral load, the trajectory of the viral load, length of time on treatment, adherence issues, obstetric factors and the woman's views.
- Viral load ≥400 HIV RNA copies/mL at 36 weeks. → PLCS recommended at 38-39 weeks.
- Viral loads more than 1,000 copies/mL at or near delivery independent of antepartum antiretroviral therapy, or those with unknown levels → PLCS.

#### Planned vaginal delivery:

Minimize vaginal examinations during labour

Avoid prolonged labour; consider oxytocin to shorten labour

Avoid artificial rupture of membranes

Use non-invasive fetal monitoring

Early cord clamping

#### Precautions to be taken during Caesarean section:

ARV prophylaxis 4 hours prior (elective)

Avoid rupturing membranes until the head is delivered through the surgical incision

Early cord clamping

Use of round blunt tip needles

Use forceps to receive and hold the needle

Peripartum antibiotics

#### Care of Newborn:

Avoid invasive nasogastric suctioning.

Wash away blood from newborn.

Start ARV prophylaxis within 1 hour.

Early infant testing-

DNA PCR at 6wks and 6 months age (Dried Blood Sample)  $\rightarrow$  95% sensitive & specific.

If positive- confirm with Whole Blood Sample PCR  $\rightarrow$  start ART if positive.

Final diagnosis  $\rightarrow$  at 18 months.

Start Cotrimoxazole prophylaxis for all neonates from 6 weeks to 18 months.

All babies detected positive <2 years of age are given Paediatric ART irrespective of CD4 count.

#### **ARV Prophylaxis for Infant:**

Daily infant NVP prophylaxis can be started even if more than 72 hours have passed since birth and should continue.

The duration of NVP given to infant is a minimum of 6 weeks, regardless of whether the infant is exclusively breast-fed or exclusive replacement fed.

Infants of women with prior exposure to NVP should get syrup Zidovudine (AZT) in place of syrup Nevirapine.

## **Keypoints**

- Provider-initiated Opt-out testing recommended as a part of battery of routine ANC tests.
- Start ART with confirmed positive test report to reduce mother to child transmission.
- Monitor Plasma HIV RNA levels at initial prenatal visit, 2–4 weeks after initiating (or changing) cART drug regimens; monthly until RNA levels are undetectable; and then at least every 3 months during pregnancy
- Check CD4 count- at initial visit and every 3-6 months.
- In women conceiving on cART there should be a minimum of one CD4 cell count at baseline and one at delivery.
- In women who commence cART in pregnancy, a CD4 cell count should be performed as per routine initiation of cART with the addition of a CD4 count at delivery even if starting at CD4 >350 cells/mm3.
- In women who commence cART in pregnancy, an

HIV viral load should be performed 2–4 weeks after commencing cART, at least once every trimester, at 36 weeks and at delivery.

- Vaginal delivery can be done with plasma viral load <50 HIV RNA copies/mL at 36 weeks, with no obstetric contraindications
- Pre-labour CS (PLCS) should be considered Plasma viral load 50–399 HIV RNA copies/mL at 36 weeks
- PLCS recommended at 38-39 weeks with viral load 2400 HIV RNA copies/mL at 36 weeks and in those with unknown levels

## **Further Reading**

- 1. WHO update 2017: What is new in treatment monitoring: Viral load and CD4 testing
- 2. BHIVA guidelines on the management of HIV in pregnancy and postpartum 2018
- 3. Clinical Obstetrics and Gynaecology 2018: HIV infection: Antepartum Treatment and Management
- 4. NACO 2018: National technical guidelines on ART.

# **Thyroid Disorders in Pregnancy**

Anubhuti Rana<sup>1</sup>, Vatsla Dadhwal<sup>2</sup>

<sup>1</sup>Assistant Professor, <sup>2</sup>Professor, All India Institute of Medical Sciences, New Delhi.



#### **Thyroid function tests:**

- Marked and early increase (7 weeks) in TBG and hCG
- Increase TBG  $\rightarrow$  Increase serum TT4 levels (peak at 16 weeks)
- Increase hCG→ increase FT4 levels→ this transient increase causes negative feedback on TSH (level decrease)
- Except for transient increased FT4 levels between 7-12 weeks, levels essentially unchanged
- Serum TSH→ after the first trimester, TSH levels return to baseline values and progressively increase in the third trimester related to placental growth and production of placental deiodinase

#### Changes in thyroid physiology during pregnancy





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#### How to start treatment and monitor in pregnancy?

- T4 replacement therapy, beginning with levothyroxine in dosages of 1–2 micrograms/kg daily
- Target TSH: -0.1-2.5 mU/L
- Measure TSH four weekly during the first half of pregnancy
- Can be monitored less often, at least once each trimester in latter half of pregnancy, as long as the dose is unchanged



#### Screening for hypothyroidism in neonate

- Screening should be done for every newborn by blood spot analysis typically 2–5 days after birth (ideally at 48 to 72 h of age).<sup>1</sup>
- Primary TSH assay is recommended for newborn screening.
- Preterm and LBW/VLBW infants should undergo routine screening for at 48–72 h postnatal age.
- Sick neonates should be screened at least by 7 days of age





#### Measure TSH 4-8 weekly

Continue levothyroxine up to 12 months post-partum

After 12 months, start tapering using serum TSH value for titration

Annually follow the patient with TSH value to prevent permanent disease



**AOGD Bulletin** 







Anti Thyroid Drugs (ATD) used in hyperthyroidism:

-Start at smallest lowest dose possible 5 mg MMI=100 mg of PTU (Maintain a ratio of 1:20 when shifting from MMI to PTU) -Doses: MMI=5-30 mg daily (OD dosing) CM=10-40 mg daily PTU=100-600 mg dailyPTU divided into 2-3 dosing

-Teratogenic effects:

MMI related: (3-4%) Aplasia cutis Choanal /esophageal atresia PTU related: (2-3%) Face and neck cysts Urinary tract abnormalities

Targets: Serum FT4/TT4 should be at/just above the reference range. Monitoring: FT4 (preferably) and TSH monitored 4 weekly

#### Anti thyroid drugs in post partum

Some amount secreted in breast milk

Use lowest dose of MMI/CM or PTU

Routine assessment of thyroid function in breastfeeding infant not recommended

Screening of neonate in mother with hyperthyroidism

- Routine evaluation of fetal thyroid function, including fetal thyroid ultrasonographic assessment, umbilical cord blood sampling, or both is not recommended.<sup>2</sup>
- Umbilical cord blood sampling should be done only when the diagnosis of fetal thyroid disease cannot be reasonably excluded based on clinical and ultrasonographic data.<sup>7</sup>

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# **Antiphospholipid Antibody Syndrome Pregnancy**

Akanksha Tiwari<sup>1</sup>, Jyoti Meena<sup>2</sup>

<sup>1</sup>Senior Resident, <sup>2</sup>Associate Professor, All India Institute of Medical Sciences

An autoimmune thrombophilic condition marked by the presence of antibodies in blood - attack phospholipid-binding proteins, rather than phospholipid itself Incidence: 5 new cases/100,000 persons per year Prevalence: 40-50cases/100,000 persons per year Prevalence of aPL in healthy indiviuals - aCL-10% - LAC-<1% In patients with SLE:20-30% have aPL



- Unexplained 2<sup>nd</sup> or 3<sup>rd</sup> trimester fetal death
- Preterm birth

- epilepsy, chorea, migraineRenal involvement- glomerulonephritis and interstitial nephritis
- Puci immune vasculitis

#### Management of Pregnancy with APS

#### Aim of Management

- 1. To maximise the chance of successful fetal outcome.
- 2. To prevent thrombosis and other clinical manifestations of APS in the mother.
- 3. To ensure good counselling and planning for future pregnancies.

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- Pre-eclampsia screen
- Surveillance for thrombosis
- Screen for Ro/La antibody

- Unfractionated (UFH)

- Low-molecular-weight heparin (LMWH)

- low-dose aspirin (75 mg once daily)

**Treatment of choice** 

#### When to start

- LDA-pre-conceptionally or as soon as UPT is positive

• Close fetal surveillance from >32 weeks with BPP

- LMWH- started in early first trimester after
- documenting intrauterine fetus rising hCG

Agent	Prophylactic	Therapeutic
UFH (unfractionated Heparin)	nated Heparin) 1st Trimester: 5000-7500 units s/c q12hr, 2nd Trimester:7500-10000 units s/c q12hr, 3rd Trimester:10000 units s/c q12hr	
LMWH (Low molecular weight heparin)	Enoxaparin 40mg s/c OD Dalteparin 5000 units s/c OD	Enoxaparin 1mg/kg s/c q12hr Dalteparin 200 units/kg s/c OD or 100 units/kg s/c q12hr Tinzaparin 175 units s/c OD (may target anti-Xa level:0.5-1.0 iu/ml)



#### Labour and delivery:

• Switched over LMWH to UFH (shorter half life) at 37 weeks POG

Reg	imen	When to stop	When to start after delivery
Prop LM	ohylactic WH	10-12 hrs	6-8 hrs
The LM	rapeutic WH	24 hrs 24-36 hrs	24 hrs

Contraception					
	Cu-IUD	LNG-IUD	Injectable DMPA	POP	COC
CDC 2016	1	2 2	2	4	
WHO 2015	1	2	2	2	4

\*LDA should be continued till at least upto 36weeks POG and should be stopped 24 hours to scheduled IOL or CS

## **Suggested Reading**

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- 3. Green-top guideline No.37a. Reducing the Risk of Venous Thromboembolism during Pregnancy and the Puerperium. RCOG 2015;205:1-40.
- 4. ACOG practice Bulletin. Antiphospholipid Syndrome. 2012;20(6):1514-21.

## **Forthcoming Events**

- Next Monthly Clinical Meeting on 27<sup>th</sup> December, 2019 (4:00-5:00 pm) at Sir Ganga Ram Hospital.
- NOTE : Monthly Meeting date of 31<sup>st</sup> January, 2020 has been changed to 17<sup>th</sup> January, 2020 due to AICOG Conference.



# Royal College of Obstetricians & Gynaecologists AICC Northern Zone India

Website: www.aiccrcognzindia.com

Chairperson: Dr Nirmala Agarwal: (n.menoky@gmail.com /9811888732)

Vice Chairperson Dr Anita Kaul Hon. Secretary Dr Arbinder Dang

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For Accommodation, Hotel Bookings, Travel Enquiry Contact Miss Carolina Fernandez Cox & Kings +919711992043/ Carolina.fernandes@cox&kings.com

#### Certificate of attendance for this course will be provided by the RCOG UK

#### **Registration Guidelines** (Online registration available on website)

- Registration form to be downloaded from website www.aiccrcognzindia.com
- Bank Transfer or Demand Draft must be made in favour of "Royal College of Obstetricians and Gynaecologists NZ India" payable at New Delhi. (Cheques not accepted).
- There will be no refunds on cancelation.
- Registration request along with Demand Draft to be posted to the Secretariat mailing address as given below:-

# Mailing Address:

## RCOG North Zone Secretariat

OT Complex 3<sup>rd</sup> Floor Sant Parmanand Hospital, 18 Shamnath Marg, Civil Lines, Delhi 110054 **Mr Asif Muniri** (Administrative Assistant) +919560069925 / 9716801190, Tel No - 91-11-23981260, 23994401-10 Ext 314 Email: rcognz2017@gmail.com/ n.menoky@gmail.com/ arbidang@gmail.com

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# **Events Held**

• Conference on Aesthetic Gynaecology, 2019 on 9<sup>th</sup> & 10<sup>th</sup> November, 2019 organized by Dr Ragini Aggarwal under the aegis of AOGD & FOGSI.



 Update in Gynaecologic Oncology on 13<sup>th</sup> November, 2019 at AIIMS under the aegis of Oncology Committee AOGD.







• A CME on "High Risk Pregnancy" on 16<sup>th</sup> November 2019 under aegis of Safe Motherhood Committee AOGD at ESI Basaidarapur, organizes by Dr Taru Gupta.





**AOGD Bulletin** 

CME on "Tackling Obstetric Dilemmas" on 16<sup>th</sup> November, 2019 at The Surya under the aegis of FOGSI Medical Disorders in Pregnancy Committee & AOGD







• Breast Cancer screening Health Camp on 19<sup>th</sup> November, 2019 by GTB under the aegis of Rural Health Committee, AOGD.







• NARCHI pre congress workshop on "PPH" on 22<sup>nd</sup> November under aegis of Safe Motherhood Committee AOGD at the Northern Railway Hospital.



CME on 27<sup>th</sup> November, 2019 under the aegis of DGF North, Breast and Cervical Cancer Awareness Screening & Prevention Subcommittee AOGD & Breast Committee FOGSI.







• Monthly Clinical Meeting on 29th November, 2019 at MAMC & LN Hospital, New Delhi.







• Dr Shashi Lata Kabra Maheshwari, Dr Mrinalini Mani , Dr Sushma Sinha and Dr Richa Sharma received Wonder FOGSI Award, 2019 at Mumbai by Dr Nandita Palshetkar.



**AOGD Bulletin** 

# **Management of SLE in Pregnancy**

#### Kusum Lata<sup>1</sup>, Jyoti Meena<sup>2</sup>

<sup>1</sup>Assistant Professor <sup>2</sup>Associate Professor, All India Institutes of Medical Sciences, New Delhi

#### Epidemiology

- Incidence: 1:250
- 1/3 remain the same, 1/3 worsen, 1/3 improve
- Newly diagnosed cases in pregnancy: tend to be more severe

#### **Best outcomes:**

- 1. Quiescent for  $\geq 6$  months
- 2. No lupus nephritis
- 3. No APLA syndrome
- 4. No superimposed pre-eclampsia

#### **Risk Stratification before pregnancy**

Increased risk of flares, hypertension, fetal morbidity, FGR, preterm delivery associated with:

- 1. Flares in last 6 months
- 2. Nephritis: increases risk
- 3. High titer of C3/C4/dsDNA
- 4. APLA positive: need for LMWH, aspirin
- 5. Anti Ro/La positive: risk for neonatal lupus, congenital heart block
- 6. End organ damage: increased maternal and fetal risk

#### Planned pregnancy is important in SLE

- Women may have menstrual irregularities, conceive while on teratogenic drugs, or during high disease activity.
- If SLE is active at the time of conception, frequency of flare higher (61-67 %) during pregnancy
- · Patients with Lupus Nephritis may have renal failure during pregnancy
- · Advise conception based on co-morbidities: HTN, DM, APLA

#### **Preconception Work up**

• Anticardiolipin antibodies and lupus anticoagulant

- Anti-Ro and anti-La antibodies, Anti dsDNA and C3 (to assess disease activity)
- S. Free T3, T4 and TSH, Complete blood count (CBC)
- Liver Function tests, Uric acid
- Renal function (creatinine, urinalysis with urine sediment)

#### Maternal Monitoring

- 1. Weekly BP monitoring
- 2. Monthly CBC, KFT
- 3. Proteinuria screen at every visit
- 4. Early GDM screen, especially if on steroid therapy
- 5. Supplement Ca, Vit D, Iron
- 6. Monthly bacteruria screen

Fetal Complications

#### Neonatal Lupus Syndrome

- Rash (photosensitive), haematologic (thrombocytopenia and neutropenia) and hepatic abnormalities
- **Congenital Heart Block (**CHB affects about 2%, develops between18-24 weeks of gestation
- Increased PR interval (first degree heart block) forewarns the subsequent development of CHB
- · Pacemakers may be required in the majority as lesion is permanent
- · Efficacy of maternal fluorinated steroids unproven in CHB

#### Management in Pregnancy

· Requires Multidisciplinary approach

#### Treatment

- HCQ: associated with good outcomes if given before, during pregnancy
- Others: Prednisolone, Azathriopine, tacrolimus
- Refractory: IV Methylprednisolone 1000 mg for 3 days/ IVIG/ plasmapheresis
- Low dose aspirin continued throughout gestation.

#### **Fetal monitoring**

- Ist trimester: NT-NB and dual screen
- 18-20 weeks: CMF, Doppler with uterine artery
- 26-28 weeks: monthly growth parameter, liquor, doppler (umbilical artery)
- Fetal ECHO: if Anti Ro/La +, dysrhythmia, if previous child affected, from 16 weeks onward weekly. Counsel on 16% recurrence if previous child affected.

#### Contraception

- Effective contraceptive measures to be discussed with the patient by weighing the individual risk factors, disease activity and thrombotic risk (presence of antiphospholipid antibodies.
- Copper IUD can be offered to all.
- OCPs & POP in patients with inactive or stable SLE and negative APLA.
- Emergency Contraception with LNG is not contraindicated in SLE or APS.
- Tubal sterilization to be performed when disease is quiscent

#### **Take Home Message**

- Pregnancy in SLE is a high risk condition and active disease at the time of conception is is associated with worse maternal and fetal outcomes.
- Pregnancy should be planned when disease is quiescent for 6 months.
- Fetal surveillance based on biometric and Doppler findings during the third trimester helps to better tailor the time of delivery and reduce perinatal morbidity and mortality

## **Suggested Reading**

Andreoli L, Bertsias GK, Agmon-Levin N, *et al. Rheum Dis* 2017;76:476–485. Williams Obstetrics 25<sup>th</sup> edition Williams

# **Congratulations !!**

Dr Neerja Varshney, Dr Supriya Chaubey and Dr Rohit Raina for correctly answering the Crossword and Pictorial Quiz of November issue

## **Answer: November Issue**

#### Crossword

- Down
- 1. Three
- 2. Progestasert
- 3. Chhaya

## Across

1. Ten

- Frameless
   Breakthrough
- Essure
   Ulipristal
- 6. Antara
- 7. Ormeloxifene
- **Pictorial Quiz Figure 1:** 1) Implant
- 2) 0.02%
- Figure 2: 1) Essure
  - 2) 3 mths
- Figure 3: 1) Frameless 10D
  - 2) Young nullipare

# **Rheumatic Heart Disease in Pregnancy: An overview**

#### Zeba Khanam<sup>1</sup>, Jyotsna Suri<sup>2</sup>

<sup>1</sup>Senior Resident, <sup>2</sup>Professor, Department of Obstetrics & Gynaecology, VMMC & Safdarjung Hospital

**Box 1:** Pre-conceptional counseling in a patient of rheumatic heart disease

- Careful preconception planning, as well as care, monitoring and support during pregnancy, can improve outcomes for both mother and baby.
- Modified WHO (mWHO) risk stratification of cardiac diseases should be done during pre-conceptional period<sup>1</sup>
- A careful note of woman's pre-existing medications is made, namely anti hypertensives, anti-coagulants, diuretics, inotropics and beta blockers and necessary modifications made with the opinion of the cardiologist.
- Conditions where pregnancy is contraindicated or where termination is warranted if pregnancy is diagnosed
  - 1. Pulmonary arterial hypertension of any cause
  - Severe systemic ventricular dysfunction (LVEF <30%, NYHA III–IV)
  - 3. Previous peripartum cardiomyopathy with any residual impairment of left ventricular function
  - 4. Severe mitral stenosis, severe symptomatic aortic stenosis
  - 5. Marfan syndrome with aorta dilated >45 mm
  - 6. Aortic dilatation >50 mm in aortic disease associated with bicuspid aortic valve
  - 7. Native severe coarctation of aorta

Box 2: Diagnosis of Rheumatic heart disease in pregnancy

- 1. Careful history and examination.
  - History of dyspnea, chest pain, syncopal attacks and productive cough.
  - Any new murmur, changes in murmur, pathological murmur (any diastolic murmur or greater than grade II systolic murmur)
  - Signs of heart failure
  - Echocardiography in unexplained or new cardiovascular signs or symptoms.
- 2. Blood pressure measurement in left recumbent position using standardized method.
- 3. Holter monitoring should in patients with known paroxysmal or persistent documented arrhythmia (ventricular tachycardia, atrial fibrillation or flutter)
- 4. Trans esophageal echocardiography-
  - Rarely needed in pregnancy.
  - Risk of vomiting, aspiration, sudden increase in intraabdominal pressure should be taken into account. Fetal monitoring should be performed if sedation is used.
- 5. Exercise testing-
  - To objectively access functional capacity, chronotropic and BP response and exercise-induced arrhythmias
  - Should be performed with known heart disease, preferably prior to pregnancy to assist in risk assessment.
  - No evidence that it increases risk of spontaneous abortion<sup>2</sup>.
  - Semi recumbent cycle ergometry appears to be the most comfortable modality

- Treadmill walking or upright cycle ergometry may also be used.
- Dobutamine stress should be avoided.
- 7. A chest radiograph, with shielding of the foetus, may be considered if other methods are not successful in clarifying the cause of dyspnoea.
- 8. Cardiac catheterization may be considered with very strict indications, timing, and shielding of the foetus.
- 9. CT and electrophysiological studies, with shielding of the foetus, may be considered in selected patients for vital indications.
- 10. Magnetic resonance imaging (MRI)
  - Useful in diagnosing complex heart disease or pathology of the aorta<sup>3</sup>.
  - It should be only performed if other diagnostic measures, including transthoracic and trans oesophageal echocardiography fail to diagnose the cause.
  - Gadolinium use to be avoided<sup>4</sup>.

Box 3: Antenatal concerns in a woman with Rheumatic heart disease

- 1. Pre-pregnancy risk assessment and counselling is indicated in women with known or suspected cardiovascular Rheumatic heart disease.
- 2. Risk assessment should be performed in all women with cardiac diseases of childbearing age and after conception.
- 3. High risk patients should be treated in specialized centres by a multidisciplinary team<sup>[5,6]</sup>.
- 4. For the prevention of infective endocarditis in pregnancy the same measures as in nonpregnant patients should be used.
- 5. Prevention and management of anaemia and any infections
- 6. Operative interventions- best time is after four months of gestation in second trimester
  - Valvular surgery may be considered when conservative and medical therapy has failed, in situations that threaten the mother's life and that are not amenable to percutaneous treatment<sup>7,8</sup>.
  - When gestational age is at least 28 weeks, delivery before necessary cardiac surgery should be considered<sup>9</sup>.

#### Specific concerns in mitral stenosis

- 1. In patients with symptoms or pulmonary hypertension, restricted activities and beta1-selective blockers (Metoprolol or Bisoprolol) are recommended <sup>10,11</sup>.
- 2. Diuretics are recommended when congestive symptoms persist despite beta blockers<sup>11</sup>.
- 3. Patients with severe MS should undergo intervention before pregnancy. If at all it is planned percutaneous mitral commissurotomy is preferably performed after 20 week gestation and only in NYHA class III/IV and/or estimated systolic PAP>50 mmHg at echocardiography despite optimal medical treatment, in the absence of contraindications and if patient characteristics are suitable<sup>10,11</sup>.
- 4. Therapeutic anticoagulation is recommended in the case of atrial fibrillation, left atrial thrombosis, or prior embolism.

#### Specific concerns in aortic stenosis

- 1. Echocardiography is mandatory for diagnosis <sup>10,12</sup>.
- Patients with severe AS should undergo intervention prepregnancy if they are symptomatic or have LV dysfunction ( LVEF <50%).</li>
- 3. Asymptomatic patients with severe AS should undergo intervention pre-pregnancy when they develop symptoms during exercise testing.
- 4. Asymptomatic patients with severe AS should be considered for intervention pre-pregnancy when a fall in blood pressure below baseline during exercise testing occurs.
- 5. Medical treatment and restricted activities are indicated for patients developing signs or symptoms of heart failure during pregnancy.
- 6. Diuretics can be administered for congestive symptoms.
- 7. A beta-blocker or a non-dihydropyridine calcium channel antagonist should be considered for rate control in AF. If both are contraindicated, digoxin may be considered<sup>13</sup>.
- During pregnancy in severely symptomatic patients not responding to medical therapy, percutaneous valvuloplasty can be undertaken. If not possible and patients have life-threatening symptoms, valve replacement should be considered after early delivery by caesarean section<sup>14</sup>.

#### Specific concerns in regurgitant lesions

- 1. Women with severe regurgitation and symptoms or compromised LV function are at high risk of heart failure<sup>15</sup>.
- 2. In asymptomatic women with preserved LV function the most frequent complications are arrhythmias.
- 3. Patients with severe aortic or mitral regurgitation and symptoms or impaired ventricular function or ventricular dilatation should be treated surgically pre-pregnancy.
- 4. Medical therapy is recommended in pregnant women with regurgitant lesions when symptoms occur.

#### Follow up during pregnancy

- I. Mitral stenosis
  - Clinical and echocardiographic follow-up is indicated monthly or bimonthly depending on haemodynamic tolerance.
  - In mild MS, evaluation is recommended every trimester and prior to delivery.
- II. Aortic stenosis
  - Regular follow-up during pregnancy is required by an experienced team.
  - In severe AS, monthly or bimonthly cardiac evaluations including echocardiography are advised to determine symptom status, progression of AS, or other complications.
- III. Regurgitant lesions
  - Follow-up is required every trimester in mild/moderate regurgitation, and more often in severe regurgitation.
  - Follow-up plans need to be individualized according to clinical status and symptoms.

#### Box 4: Intrapartum and postpartum care

- 1. Vaginal delivery is recommended as first choice in most patients.
- 2. While there is no absolute contraindication to misoprostol or dinoprostone, there is a theoretical risk of coronary vasospasm and a low risk of arrhythmias. Dinoprostone also has more profound effects on BP than misoprostol and

is therefore contraindicated in active cardiovascular disease. Mechanical methods such as a Foley catheter would be preferable to pharmacological agents, particularly in the patient with cyanosis<sup>16</sup>.

- 3. Continuous fetal heart rate monitoring should be done.
- 4. Maternal BP and heart rate should be monitored in all patients.
- 5. Pulse oximetry and continuous ECG monitoring are advised to detect early signs of decompensation.
- 6. In some high-risk patients (PH), right atrial pressure monitoring may be considered.
- Epidural analgesia reduces labour pain and can be used to provide anesthesia for caesarean section if necessary. However, it can cause systemic hypotension (10%) and must be carefully titrated in patients with obstructive valve lesions or diminished ventricular function.
- 8. All intravenous fluids need to be infused carefully.
- 9. At all times lateral decubitus position is recommended <sup>17</sup>.
- 10. A careful watch of foetal head descent is required.
- 11. In severe symptomatic AS, particularly during the second half of the pregnancy, caesarean delivery should be preferred with endotracheal intubation and general anaesthesia. In non-severe AS, vaginal delivery is favoured.
- 12. The active phase of the second stage should be delayed for 2 h to allow maximal descent of the foetal head
- 13. Assisted delivery with forceps or a ventouse during the second stage of labour may be used to further reduce maternal efforts.
- 14. Active management of the third stage of labour is recommended. Slow i.v. infusion of oxytocin (2 U of oxytocin given over 10 min immediately after birth, followed by 12 mU/min for 4 h) reduces the risk of post-partum haemorrhage. Misoprostol (200–1000 mg) can be used to treat postpartum haemorrhage; however, ergometrine and prostaglandin F analogues should be avoided.
- 15. Prophylactic antibiotic therapy during delivery is not recommended <sup>18</sup>.
- 16. The post-partum period is associated with significant haemodynamic changes and fluid shifts, particularly in the first 24–48 h after delivery, which may precipitate HF.
- 17. Haemodynamic monitoring should, therefore, be continued for at least 24–48 h in those at risk<sup>19</sup>.
- 18. Meticulous leg care, elastic support stockings, and early ambulation are important to reduce risk of thromboembolism.
- 19. With preceding beta-blockade, infant monitoring for 48 h is recommended.
- 20. Lactation is associated with a low-risk of bacteremia secondary to mastitis and should be encouraged in patients with heart disease whenever possible.

Box 5: Indications for caesarean section in heart disease<sup>10,20,21</sup>

Caesarean delivery should be considered for obstetric indications and/or

- 1. Dilatation of the ascending aorta >45 mm
- 2. Severe aortic stenosis
- 3. Patient presenting in labour while on oral anticoagulants
- 4. Severe form of pulmonary artery hypertension
- 5. Eisenmenger syndrome
- 6. Acute intractable heart failure

Box 6: Contraception and follow up

1. Injectable progesterone only contraceptives (DMPA and NET en) are safe in valvular heart disease

- 2. Low dose oral contraceptives containing 20 mg of ethinyl estradiol are safe in women with a low thrombogenic potential, but not in women with complicated valvular disease (pulmonary hypertension, risk of atrial fibrillation, history of subacute bacterial endocarditis).
- 3. Levonorgestrel-based long-acting reversible contraception implants or intrauterine devices are the safest and most effective contraceptives.

Barrier methods are unreliable. A good approach is the combination of barrier methods and long-acting reversible contraception.

- 4. A copper intrauterine device is acceptable.
  - a. Antibiotic prophylaxis is not recommended at the time of insertion or removal.
  - b. If excessive bleeding occurs at the time of menses, the device should be removed.
- 5. Tubal ligation is usually accomplished safely, even in relatively high risk women.

a. Risks are increased in cases of pulmonary artery hypertension.

6. Vasectomy for the male partner is another efficacious option.



Figure 1: Recommendations for women with mechanical prosthetic valves

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Months	Name of the Institute
27 <sup>th</sup> December, 2019	Sir Ganga Ram Hospital
17 <sup>th</sup> January, 2020	Dr RML Hospital
28 <sup>th</sup> February, 2020	UCMS & GTB Hospital
27 <sup>th</sup> March, 2020	LHMC
24 <sup>th</sup> April, 2020	Apollo Hospital

## **Calendar of Monthly Clinical Meetings 2019-20**

# **Gestational Diabetes in Pregnancy: Screening and management**

#### Richa Vatsa, Garima Kachhawa

<sup>1</sup>Assistant professor, <sup>2</sup>Additional Professor, All India Institute of Medical Sciences, New Delhi.



β

Sugar not controlled on metformin

#### Insulin

Human premix insulin (30/70) or regular (Actrapid) with intermediate acting (Insulitard) DIPSI guideline recommend only insulin (Human premix 30/70) for GDM

 $FBS \ge 95 \text{ mg/dl}$ : Pre Dinner 4U insulin

2 Hr PP BS  $\geq$  120: Pre Breakfast insulin

Blood sugar (mg/dl)	Insulin (U)
120-140	4
140-160	6
160-180	8

Still FBS  $\ge$  95 mg/dl and/ or 2 Hr PP BS  $\ge$  120: Increase dose of insulin by 2u Pre dinner and /or Pre breakfast respectively

Repeat FBS and PPBS every 3rd day till dose of insulin adjusted.

#### **Fetal monitoring**

Fetal growth scan should be performed at 28-30 wk POG & repeated at 34-36 wk POG At least 3 wk gap between two ultrasounds for fetal biometry & AFI estimation. Explain about daily fetal movement activity assessment

#### Labour and delivey

GDM with well controlled BS: IOL at or after 39 wk POG (DIPSI)/  $40^{+6}$  wk (NICE 2013) GDM with poor BS control, those with risk factors like hypertensive disorder of pregnancy, previous still birth & other complications timing of delivery: Individualised

Fetal macrosomia (EFW >4 kg) consideration for a primary CS at 39 weeks to avoid shoulder dystocia

#### Special precaution during labour

Omit morning dose of insulin/metformin on day of induction/labour, monitor 2 hourly monitoring of blood sugar. IV infusion with NS & add regular insulin according to BS levels as per table below.

Blood Sugar Level	Insulin added in 500 ml NS	Rate of NS Infusion
90-100	0U	100 ml/hr (16 drops/min)
120-140 mg/dL	4U	100 ml/hr (16 drops/min)
140-180 mg/dL	6U	100 ml/hr (16 drops/min)
>180 mg/dL	8U	100 ml/hr (16 drops/min)

#### Post delivery follow up of pregnant women with GDM

OGTT with 75 gm glucose (F and 2 hr PP) 6 wk postpartum: cut off values FBS:  $\geq$  126 mg/dl 2 hr PP: Normal: < 140 mg/dl, IGT: 140-199 mg/dl, DM:  $\geq$  200 mg/dl

## **Suggested Reading**

NICE guideline on diabetes in pregnancy 2015 IADPSG Consensus Panel. Diabetes care 2010 DIPSI guideline, Govt Of India 2018

**AOGD Bulletin** 

# **Pregnancy with Chronic Kidney Disease (CKD)**

#### Juhi Bharti, Deepali Garg

Assistant Professor, Department of Obstetrics & Gynaecology, All India Institute of Medical Sciences, New Delhi.



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Table 1: Recommendations of immunosuppressive medications in pregnancy				
Drug	Effects on Pregnancy	Recommendation in pregnancy		
Prednisolone	Maternal HTN & GDM Risk of thymic hyperplasia & adrenal suppression in neonate	May be continued Avoid prolonged high doses		
Azathioprine	No teratogenicity	May be continued		
Tacrolimus	Increased risk of GDM Transient neonatal renal dysfunction and hyperkalemia	May be continued Monitor & adjust levels Early OGTT		
Cyclosporine A	No teratogenicity, maternal HTN	May be continued, monitor levels		
Mycophenolate mofetil	Teratogenic & embryopathic Multiple congenital defects	Stop 12 weeks prior to conception		
Cyclophosphamide	Teratogenic in first trimester	Stop 12 weeks prior to conception		

<sup>1</sup>GFR: Glomerular Filtration rate, <sup>2</sup>HTN: Hypertension, <sup>3</sup>PE: Preeclampsia, <sup>4</sup>PCR: Protein Creatinine ratio, <sup>5</sup>ACEI: angiotensin -converting enzyme inhibitors, <sup>6</sup>ARBs: angiotensin receptor blocker, <sup>7</sup>MTP: Medical termination of pregnancy



**AOGD Bulletin** 

#### Maternal surveillance

Frequent ANC visits\* q 2 weekly initially >32 weeks-weekly

Investigations\* Renal function: monthly Proteinuria- monthly BP- weekly Bacteriuria-each trimester GDM screening: early

#### Fetal surveillance

Routine screening Risk of false positive due to increased B hCG (renal excretion) cffDNA or invasive testing may be required Fetal growth from 28 weeks q 2-3 wks Weekly/biweekly BPS after 28 weeks



#### **Optimize Condition**

**Preeclampsia prevention** Tab Ecosprin 150 mg HS from 12 weeks till 36 weeks

**Nutrition (Dietician counseling)** Diet suitable for pregnancy & renal disease Limit non nutritive sources of Phosphate

Anemia Iron supplementation Add erythropoietin to non dialysis CKD patients if Hb<10

Hypertension Target BP-110/70 to 135/85 Safe antihypertensive drugs: Labetatol, CCB-Nifedipine, α methyl dopa. Add diuretics if associated edema and reduced GFR

Thromoprophylaxis with LMWH if nephrotic

range proteinuria

#### Metabolic Bone Disease

(Women on treatment in preconception period) Monitor Calcium, Phosphorus, 25 OH Vit D, PTH once every trimester Use Phosphate binders: Calcium carbonate

#### Indications of delivery

Severe Preeclampsia Fetal Growth Restriction Non reassuring fetal heart testing Worsening renal disease

#### Indication of hemodialysis

- eGFR <20ml/min/1.73 m<sup>2</sup>
- · Signs/symptoms of uraemia
- · Persistent volume overload
- · Refractory acidosis
- · Hyperkalemia
- · Hyperphosphatemia

#### **Postpartum period**

- · ACE inhibitors may be resumed
- Captopril, Enalapril not excreted in breast milk
- Continue Erythropoietin and IV Iron as necessary
- Monitor electrolytes, Creatinine & Urine PCR
- Maintain BP-140/90
- · Avoid dehydration and pulmonary edema
- Measure urine output and look for sign of fluid overload
- Avoid NSAID
- Safe & effective contraception

# **Algorithmic Approach to Liver Disease in Pregnancy**

#### Sharda Patra

Professor, Dept of Obstetrics & Gynecology, Lady Hardinge Medical College, New Delhi

## Background

Liver disease in pregnancy manifests mostly with jaundice as first sign or with abnormal liver function tests. Jaundice in pregnancy is one of the major indirect cause of maternal mortality and is responsible for 5-30% of all maternal deaths. Jaundice in pregnancy when presents in a milder form with mild elevation in liver enzyme is associated with a favorable maternal fetal outcome. However when it presents in a more severe form associated with an alarmingly abnormal elevations in liver enzymes, the resulting liver failure and death of the mother and her fetus, then becomes a cause of concern.

### Management

Algorithmic management depending on the cause of liver disease and biochemical markers is shown in Fig 4, 5, 6

## Conclusion

Liver disease in pregnancy poses a diagnostic and management dilemma. Thus the management of jaundice in pregnancy do present with many dilemmas from diagnosis to its management but an in depth understanding of the physiologic changes during pregnancy, ability to identify and treat liver disorders, and a proper vigilance in recognizing clinical and laboratory abnormalities in a timely manner along with a coordinated team approach management involving the primary care physician, obstetrician, hepatologist, and a transplant surgeon can to some extent promote a favourable maternal and fetal outcomes.



Figure 1: Causes of liver disease in Pregnancy

\*Most commonly manifests in postpartum period

Abbreviations: IHCP- Intrahepatic Cholestasis of pregnancy, AVH- Acute Viral Hepatitis AFLP- Acute fatty liver of pregnancy HELLP- Hemolysis, Elevated Liver enzymes and Low Platelet count syndrome, PH- Portal Hypertension



Figure 2: Algorithmic Initial Work up of a pregnant women presenting with jaundice in Pregnancy





Figure 3: Algorithm of work up of a pregnant women presenting with Severe jaundice based on Lab investigations

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Figure 4: Algorithm of Management of a pregnant women presenting with mild jaundice





Aim of the management: To deal with liver failure by giving supportive treatment, Correct coagulation, prevent progression of hepatic encephalopathy, correction of metabolic disturbances and to deal with obstetric complications



Figure 6a: Algorithm of management of a pregnant women presenting with Severe jaundice with suspected liver failure-Medical management

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- Vaginal packing should not be done as routine as this lead to more lacerations.
- · Delivery should always be conducted by an experienced obstetrician.
- PPH from uterine atony not responding to medical management.

Figure 6b: Algorithm of management of a pregnant women presenting with Severe jaundice with suspected liver failure-Obstetrical management

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# **Immunization in Pregnancy**

#### Soniya Dhiman<sup>1</sup>, Vidushi Kulshrestha<sup>2</sup>

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- Vaccines safe in pregnancy: Toxoids - Diphtheria, tetanus Bacterial vaccines - Acellular pertussis Inactivated viral vaccines -Inactivated influenza Inactivated Polio Vaccine (IPV) Hepatitis A Rabies Fractional Typhoid (parenteral) Hepatitis B Pneumococcal Meningococcal Yellow Fever, though live attenuated, can be given if indicated
- Vaccines contraindicated during pregnancy: **Live Attenuated Vaccines** Bacterial -Viral -Measles BCG Mumps Oral Typhoid Rubella Human papilloma virus (HPV) Vaccinia Varicella Herpes Zoster Rotavirus Live attenuated influenza Oral Polio • After administration, pregnancy should be avoided for at least 4 weeks. • Termination is not recommended on inadvertent administration.
  - Termination is not recommended on inadvertent administration.
  - HPV: If HPV vaccine series was interrupted for pregnancy, the series should be resumed postpartum with the next dose.
  - HPV vaccine can be given to breastfeeding women 26 years and younger who have not been previously vaccinated.
  - Though there is viral shedding with live vaccines, close contacts of pregnant women may be immunised.

#### Vaccines Recommended during every pregnancy: TT, Td or Tdap

#### TT (Tetanus toxoid): As per National Immunization Programme 2019

- 0.5 mL IM, 2 doses 4 weeks apart, in upper arm, first dose early in pregnancy.
- Only TT Booster- if received 2 TT doses in a pregnancy within the last 3 years.
- Second dose TT or TT booster can be given before 36 weeks or after 36 weeks of pregnancy or in labour, if she has not received previously.

#### Td (Tetanus, diphtheria toxoid): As per WHO 1998

- 0.5 mL IM, 2 doses 4 weeks apart. Td Booster- if received 2 Td doses in a pregnancy within the last 3 years.
- National Technical Advisory Group on Immunization (NTAGI), Ministry of Health & Family Welfare has recommended the replacement of TT vaccine with Td vaccine in India's immunization programme for all age groups, including pregnant women.

#### Tdap (tetanus toxoid, reduced diphtheria toxoid and acellular pertussis): As per ACOG Committee 2017

- Single IM dose, preferably between 27-36 weeks of gestation or at any time during pregnancy if needed such as in pertussis outbreak or wound management.
- Once received, no need to repeat the dose at 27-36 weeks.
- Tdap vaccination at 27–36 weeks gestation maximises passive antibody transfer to the newborn and protects infant from pertussis till age of routine infant vaccination (8 weeks).
- if not given during pregnancy, give postpartum if woman has never received a prior dose of Tdap as an adolescent, adult or during a previous pregnancy.
- In patients with unknown or incomplete tetanus vaccination--dT, 3 doses, at 0,1, and 6 -12 months after the 2nd. One dose of dT replaced with Tdap preferably given between 27-36 weeks of gestation. (ACOG Committee 2017)

#### Influenza:

#### Inactivated Influenza:

- One dose IM, should be given with each influenza season. (Vaccine should be administered at least one month prior to the commencement of the season.
- Can be given in any trimester, irrespective of duration of pregnancy.
- Provides passive immunity to the fetus.
- In suspected/documented infection- immunization should be given along with treatment with oseltamivir.

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## Special vaccines for certain population:

S. No	Immunological agent	Indications for immunization during pregnancy	Dose Schedule	Comments
1	Pneumococcus a) 23-valent pneumococcal polysaccharide vaccine (PPSV23)	<ul> <li>a) Heart disease, lung disease, sickle cell disease, diabetes, other chronic illnesses.</li> </ul>	In adults, one dose only; repeat dose in 6 years in high risk women	a) PSPV13, safety in 1st trimester has not been evaluated.
	b) 13-valent pneumococcal vaccine (PCV13)	b) Human immunodeficiency virus (HIV) infection and asplenia.		b) PCV13 vaccine should be deferred in pregnant women.
2	<ul> <li>Meningococcus</li> <li>a) Quadrivalent conjugate meningococcal vaccine (MenACWY)</li> <li>b) Meningococcalsero group B vaccine</li> </ul>	<ul> <li>Individuals with HIV infection</li> <li>Complement component deficiency (including eculizumab use)</li> <li>Functional or anatomic asplenia (including sickle cell disease)</li> <li>Exposure during disease outbreak</li> <li>Travel to endemic</li> <li>Microbiologist routinely exposed to Neisseria meningitidis.</li> </ul>	One dose, tetravalent vaccine	Meningococcal serogroup B vaccine should be deferred in pregnant women, unless the woman is at increased risk of serogroup B meningococcal disease
3	Rabies	Postexposure prophylaxis	Each case considered individually	
4	Hepatitis A	Chronic liver disease, clotting-factor disorders, traveling, drug abusers	Two doses IM 6 months apart	
5	Hepatitis B	<ul> <li>Women who have</li> <li>Hepatitis B surfaceantigen– positivehousehold contacts or sex partners</li> <li>More than one sex partner during the previous 6 months</li> <li>Evaluated or treated for a sexually transmitted infection</li> <li>Current or recent injection-drug users</li> <li>Chronic liver disease</li> <li>HIV infection</li> </ul>	Three doses series IM at 0,1 and 6 months	Used with hepB IgG for some exposure
6	Poliomyelitis	For women at risk	two doses of IPV SC at at4-8 week intervals and a 3rd dose 6-12 months after the 2nd.	Travelling in endemic areas
7	Yellow Fever	For women at risk	Single dose SC	Travelling in endemic areas
8	Typhoid	Women having close, continuous exposure or travel to endemic areas	Two doses IM 4 weeks apart	Inadequate data.

### **Specific Immunogobulins:**

S. No	Immunological agent	Indications for immunization during pregnancy	Dose Schedule	Comments
1	Hepatitis B	Post exposure prophylaxis	Depends on exposure	Given with Hep B vaccine to exposed neonates.
2	Rabies	Post exposure prophylaxis	Half dose at injury site, half dose at deltoid.	given with vaccine.
3	Tetanus	Post exposure prophylaxis	One dose IM	given with tetanus toxoid.
4	Varicella	To exposed pregnant women	one dose IM within 96 hours of exposure	

#### Vaccines for the future:

- 1. Respiratory syncytial virus (RVS) Vaccine: boost the neonatal immunity by trans placental transfer, specially for the first three months of life.
- 2. Group B streptococcal vaccine: maternal immunisation will help in prevention of GBS disease in mothers and newborns.

#### Take Home Message

- Obstetrician-gynaecologists and other obstetric care providers should routinely assess their pregnant patient vaccination status.
- All vaccines are safe in lactating women.
- MMR, Varicella, Tdap, Inactivated influenza vaccines should be given postpartum if not received at all.

## **Suggested Reading**

ACOG Committee Opinion. Williams 25th edition CDC 2016/2017 NIS 2019- (mohfw.gov.in)

Journal Scan - I

Shikha Sharma All India Institute of Medical Sciences, New Delhi

# Placental growth factor testing to assess women with suspected pre-eclampsia: a multicentre, pragmatic, stepped-wedge cluster-randomised controlled trial.

Kate E Duhig, Jenny Myers, Paul T Seed, Jenie Sparkes, Jessica Lowe, Rachael M Hunter, Andrew H Shennan, Lucy C Chappell, on behalf of the PARROT Trial Group

**Introduction:** Hypertension affects 10% of pregnant women, and pre-eclampsia complicates around 3% of singleton pregnancies. Diagnosis is based on clinical features such as hypertension and raised urinary protein excretion, both of which are subject to observer error, heterogeneity in test accuracy, and an insufficient ability of clinicians to predict important adverse pregnancy outcomes. The presentation of pre-eclampsia is often clinically ambiguous, and risk stratification of women with suspected pre-eclampsia is complex. This ambiguity leads to repeated hospital attendances for antenatal monitoring, increased use of health resources, and considerable anxiety for women, while missing at-risk cases

Angiogenic factors being associated with the pathophysiology of pre-eclampsia, have shown good performance in predicting the need for delivery in women with suspected pre-eclampsia. In a study of the accuracy of tests in diagnosing pre-eclampsia, low circulating maternal placental growth factor (PIGF) concentrations had a high sensitivity (96%; 95% CI 89–99) and negative predictive value (98%; 93–99.5) in diagnosing pre- eclampsia that required delivery within 14 days in women who presented with suspected pre-eclampsia.

Aims and Objectives: This study aimed to determine whether knowledge of circulating PIGF concentration, integrated with a clinical management algorithm, decreased the time for clinicians to make a diagnosis in women with suspected pre- eclampsia. Also, to determine whether this approach reduced subsequent maternal or perinatal adverse outcomes.

**Background:** Previous prospective cohort studies have shown that angiogenic factors have a high diagnostic accuracy in women with suspected pre-eclampsia, but there is still uncertainty regarding the effectiveness of these tests in a real-world setting. It was aimed to determine whether knowledge of the circulating concentration of placental growth factor (PIGF), an angiogenic factor, integrated with a clinical management algorithm, decreased the time for clinicians to make a diagnosis in women with suspected pre-eclampsia, and whether this approach reduced subsequent maternal or perinatal adverse outcomes.

Methods: This was a multicentre, pragmatic, stepped-wedge cluster-randomised controlled trial in 11 maternity units in the UK, which were each responsible for 3000-9000 deliveries per year. Women aged 18 years and older who presented with suspected pre-eclampsia between 20 weeks and 0 days of gestation and 36 weeks and 6 days of gestation, with a live, singleton fetus were invited to participate by the clinical research team. Suspected pre- eclampsia was defined as new-onset or worsening of existing hypertension, dipstick proteinuria, epigastric or right upper-quadrant pain, headache with visual disturbances, fetal growth restriction, or abnormal maternal blood tests that were suggestive of disease (such as thrombocytopenia or hepatic or renal dysfunction). Women were approached individually, they consented for study inclusion, and they were asked to give blood samples. The maternity units, representing the clusters, were randomly allocated to blocks. Blocks represented an intervention initiation time, which occurred at equally spaced 6-week intervals throughout the trial. At the start of the trial, all units had usual care (in which PIGF measurements were also taken but were concealed from clinicians and women). At the initiation time of each successive block, a site began to use the intervention (in which the circulating PIGF measurement was revealed and a clinical management algorithm was used). Enrolment of women continued for the duration of the blocks either to concealed PIGF testing, or after implementation, to revealed PIGF testing. The primary outcome was the time from presentation with suspected pre-eclampsia to documented pre-eclampsia in women enrolled in the trial who received a diagnosis of preeclampsia by their treating clinicians. This trial was registered with ISRCTN, number 16842031.

Findings: Between June 13, 2016, and Oct 27, 2017, 1035 women with suspected pre-eclampsia were enrolled

and assessed. Of the 1023 eligible women, 576 (56%) women were assigned to the intervention (revealed testing) group, and 447 (44%) women were assigned to receive usual care with additional concealed testing (concealed testing group). Three (1%) women in the revealed testing group were lost to follow-up, so 573 (99%) women in this group were included in the analyses. One (<1%) woman in the concealed testing group withdrew consent to follow-up data collection, so 446 (>99%) women in this group were included in the analyses. The median time to pre-eclampsia diagnosis was 4·1 days with concealed testing versus 1·9 days with revealed testing (time ratio 0·36, 95% CI 0·15–0·87; p=0·027). Maternal severe adverse outcomes were reported in 24 (5%) of 447 women in the concealed testing group versus 22 (4%) of 573 women in the revealed testing group (adjusted odds ratio 0·32, 95% CI 0·11–0·96; p=0·043), but there was no evidence of a difference in perinatal adverse outcomes (15% vs 14%, 1·45, 0·73–2·90) or gestation at delivery (36·6 weeks vs 36·8 weeks; mean difference -0.52, 95% CI -0.63 to 0.73).

**Interpretation:** It was found that the availability of PIGF test results substantially reduced the time to clinical confirmation of pre-eclampsia. Where PIGF was implemented, there was a lower incidence of maternal adverse outcomes, consistent with adoption of targeted, enhanced surveillance, as recommended in the clinical management algorithm for clinicians. Adoption of PIGF testing in women with suspected pre-eclampsia is supported by the results of this study.

**Summary:** A major challenge in modern obstetrics is early identification of pregnancies at high-risk of early onset PE and undertaking the necessary measures to improve placentation and reduce the prevalence of the disease.

Extensive research in the last 20 years, mainly as a consequence of the shift in screening for aneuploidies from the second- to the first-trimester of pregnancy, has identified a series of early biophysical and biochemical markers of impaired placentation. Using a novel Bayes-based method that combines prior information from maternal characteristics and medical history, uterine artery pulsatility index (PI), mean arterial pressure (MAP), and maternal serum pregnancy-associated plasma protein-A (PAPP-A) and placental growth factor (PIGF) at 11–13 weeks' gestation can identify a high proportion of pregnancies at high-risk for early onset PE.

A large number of biochemical markers have been investigated for the prediction of PE. Maternal serum PAPP-A and PIGF are two biochemical markers that have been investigated extensively and have shown promising results in the early prediction of PE. The addition of maternal serum PAPP-A and PIGF to maternal factors improves the detection rates from 36% to 60% and from 33% to 43%, at false-positive rate of 5%, and from 51% to 74% and from 43% to 56%, at false-positive rate of 10%, for PE requiring delivery before 34 and 37 weeks' gestation, respectively.

This trial has demonstrated that, the clinical use of PIGF measurement could present a change for antenatal care that improves speed of diagnosis and improves pregnancy outcomes. The findings of this study provide novel evidence supporting the adoption PIGF testing as a diagnostic adjunct for suspected pre-eclampsia. Evaluation of the intervention with women stratified by PIGF category could further elucidate the mechanisms by which PIGF testing and our management algorithm affect maternal outcomes.

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**Journal Scan - II** 

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## What Is New in Maternal Heart Disease?

**Best Articles From the Past Year** 

Metz TD. Obstet Gynecol 2019;133:181-2.

Dr Metz reviewed four recent papers on heart disease with pregnancy.

## **Pregnancy Outcomes in Women with Heart Disease: The CARPREG II Study** Silversides CK et al. J Am Coll Cardiol 2018;71: 2419–30.

Several models exist to predict the likelihood of serious morbidity or mortality in women with heart disease who opt to pursue pregnancy. Investigators from two large Canadian tertiary care hospitals prospectively followed 1,938 pregnancies among women with maternal heart disease for cardiac outcomes from 1994 to 2014. Aims were to examine temporal trends and identify predictors of complications antepartum or postpartum. Cardiac complications (maternal cardiac death, cardiac arrest, sustained arrhythmia requiring treatment, left- or right-sided heart failure, stroke or transient ischemic attack, cardiac thromboembolism, myocardial infarction, vascular dissection) were present in 16% of the cohort. Complication rates did not change over time. General cardiac factors, lesion specific variables, and late pregnancy care variables were ultimately included in the CARPREG-II risk prediction model. With a C-statistic of 0.77 (95% CI 0.74–0.81), the CARPREG-II model exceeded the predictive capability of existing models and may allow for improved risk assessments for women with cardiac disease. However, unique individual circumstances must still be considered.

PREDICTOR	POINTS
Prior cardiac events or arrhythmias	3
Baseline NYHA III-IV or cyanosis	3
Mechanical valve	3
Ventricular dysfunction	2
High risk left-sided valve disease/ left ventricular outflow tract obstruction	2
Pulmonary hypertension	2
Coronary artery disease	2
High risk aortopathy	2
No prior cardiac intervention	1
Late pregnancy assessment	1

The CARPREG (Cardiac Disease in Pregnancy Study) II risk score is based on 10 predictors, shown in the box above. Each predictor is assigned a weighted point score. The sum of points represents the risk score. Risk scores are categorized into the 5 groups: 0 to 1 points; 2 points; 3 points; 4 points and >4 points The predicted risks for primary cardiac events stratified according to point score were 0 to 1 points (5%), 2 points (10%), 3 points (15%), 4 points (22%), and >4 points (41%).

Bottom Line: The CARPREG-II prediction model integrates both lesion-specific and general factors to improve prediction of cardiac morbidity among women with heart disease in pregnancy and postpartum.

# Pregnancy Outcomes in Women with Rheumatic Mitral Valve Disease: Results from the Registry of Pregnancy and Cardiac Disease

Van Hagen IM et al. Circulation 2018;137:806–16.

Investigators used data from the international, prospective Registry of Pregnancy and Cardiac Disease to examine pregnancy outcomes for women with rheumatic mitral valve disease and no history of valve replacement. Of 390 included women, 75.4% were from developing countries. The majority (70%) had mitral stenosis with or without regurgitation. Before pregnancy, 26.9% had a percutaneous or surgical valve repair. During pregnancy, heart failure was common, affecting 23.1% of women with moderate or severe mitral regurgitation. Similarly, hospital admission, typically for heart failure, was common among women with mitral stenosis (23.1% overall and 49.1% with severe stenosis). A New York Heart Association Classification of II or greater was associated with adverse cardiac events (odds ratio 3.77, 95% CI 1.93–7.38). In women with severe stenosis (valve area less than 1.0 cm2), those with an intervention before pregnancy had fewer adverse cardiac events than those without a repair (14%versus 66%, P5.014).

Bottom Line: Almost 50% of pregnant women with severe mitral stenosis and 23% of those with significant mitral regurgitation developed heart failure. Preconception counselling with pre-pregnancy valve repair or replacement should be considered in women with mitral valve disease.

## Subsequent Pregnancy Outcomes in Patients with Peripartum Cardiomyopathy Codsi E, Rose CH, Blauwet LA. Obstet Gynecol 2018; 131:322–7.

Data regarding how to counsel women with a history of peripartum cardiomyopathy about the risk of relapse in subsequent pregnancies are limited. Investigators used a retrospective cohort design to evaluate neonatal and maternal outcomes of all pregnant patients with prior peripartum cardiomyopathy at a single center from January 2000 to March 2017. There were 43 subsequent pregnancies among 25 participants; all but one woman had recovery of cardiac function to a normal ejection fraction (50% or greater) before any subsequent pregnancy. The majority (76.7%) of subsequent pregnancies resulted in live births (four late preterm). Relapse occurred in nine women over the 43 pregnancies, for a relapse rate of 20.9%. There were no deaths, and all women who relapsed again recovered normal ventricular function, with a median recovery time of 1 month. Although the cohort is limited by small numbers and may be biased to women with favourable outcomes after the incident pregnancy, the authors provide valuable information for counseling women with a history of peripartum cardiomyopathy.

Bottom Line: Women with a history of peripartum cardiomyopathy and recovered cardiac function (normal ejection fraction) had a 21% chance of relapse in subsequent pregnancies.

# Improving Healthcare Response to Cardiovascular Disease in Pregnancy and Postpartum

### Hameed AB, Morton CH, Moore A. Retrieved October 23, 2018.

Cardiovascular disease is the leading nonobstetric cause of maternal mortality. As such, the California Maternal Quality Care Collaborative partnered with the California Department of Public Health to produce a toolkit for clinicians to improve the health care response to cardiovascular disease in pregnancy and postpartum. This toolkit is free and available to health care providers on the California Maternal Quality Care Collaborative's website: https://www.cmqcc. org/resources-toolkits/toolkits/improving-healthcare-response-cardiovascular-disease-pregnancy-and. The toolkit consists of algorithms to guide risk stratification and evaluation of women with known cardiac disease or signs and symptoms of cardiac disease. Specifically, signs and symptoms that should prompt evaluation by a specialist are delineated. In addition, the toolkit provides health care providers with recommended resources for caring for women with complex cardiovascular disease in pregnancy, which includes discussions of appropriate contraceptive counseling and information about the risk profiles of various

cardiac medications. Finally, the document focuses on known racial disparities in cardiovascular disease prevalence and outcomes.



#### CARDIOVASCULAR DISEASE ASSESSMENT IN PREGNANT and POSTPARTUM WOMEN

Bottom Line: A toolkit to improve health care provider response to cardiovascular disease in pregnancy and postpartum is free for download and use by clinicians. The purpose of the toolkit is to prevent maternal morbidity and mortality related to cardiovascular disease in pregnancy.

**Editor's comments:** Heart disease is one of the most important cause of maternal morbidity and mortality especially in the developing countries. Lack of access or late access to health care is the major cause of poor outcome in such patients. The complication rate increases with severity of lesion and NYHA functional status. Hence early booking in pregnancy, early diagnosis by thorough clinical examination at initial visit and prompt management will help to avert untoward consequences. Preconception counselling with valve repair or replacement prior to pregnancy should be considered in women with mitral valve disease. Balloon mitral valvotomy during pregnancy is a safe and technically feasible procedure which will help to reduce the cardiac complications with better maternal fetal outcome.

Peripartum cardiomyopathy is a rare form of pregnancy associated idiopathic dilated cardiomyopathy developing in last trimester to first five months postpartum. There is around 21% chance of recurrence of disease in subsequent pregnancy. The 2011 European Society of Cardiology guidelines state that subsequent pregnancies should be discouraged in patients with nonrecovered ventricular function, providing the nondirective recommendation of the "need for counselling because of the risk of recurrence with a new pregnancy" in patients who recover LV function. A recent scientific statement from the American Heart Association makes very similar recommendations. Hence, proper counselling of women is must before planning for next pregnancy. CARPREG II risk score is a good predictor of morbidity in women with heart disease in pregnancy.

# Clinical Proceedings of AOGD Clinical Meeting held at MAMC & LN Hospital, New Delhi on 29<sup>th</sup> November, 2019

#### **Aggressive Angiomyxoma**

#### Y M Mala, Sarah F Siddiqui, Shikha Sharma

#### Background

Aggressive Angiomyxoma is a rare (only 350 cases reported till now worldwide), locally invasive mesenchymal tumor that most commonly arises in the vulvovaginal region, perineum and pelvis of women. We present here 2 rare cases of Aggressive Angiomyxoma with varying presentations one arising from pelvis and other from the vulva. While one underwent complete surgical excision of the neoplasm with no evidence of recurrence in the subsequent follow-ups the other patient succumbed to pulmonary and renal metastasis.

#### **Case Presentations**

A 45 year old lady referred from a private hospital with provisional diagnosis of cervical fibroid based on her MRI findings. On examination, a large mass was felt in the vagina which was firm, smooth, nontender and immobile. She later developed retention of urine. CT scan revealed a large (13 x 12x 10 cms) mass displacing bowel, uterus, ureters, causing, hydroureteronephrosis and bilateral renal metastasis. Her chest x-ray too revealed probable metastasis. Biopsy of the mass was performed which confirmed diagnosis of Angiomyxoma. Immunohistochemistry was performed and found to be positive for vimentin, estrogen and progesterone receptors. She opted for neoadjuvant GnRH therapy. Unfortunately, the patient expired soon after initiating the therapy.

The second patient presented with mass growing on right side of vulva having increased massively in size during her recent pregnancy, now measuring  $10 \times 8 \text{ cms}$  with a pedicle of  $10 \times 2 \text{ cms}$ . Histopathology and immunohistochemistry confirmed Aggressive Angiomyxoma for which wide local excision was done. There was no evidence of recurrence noted till date.

#### Discussion

Aggressive angiomyxoma has a very high risk for local relapse even though metastasis is very rare. Hence we need to differentiate it from other mesenchymal tumors occurring in this region. Clinically it presents

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mostly as asymptomatic mass varying in size from 1 to 60 cms with cut section revealing homogenous, gelatinous and glistening appearance. Microscopically it reveals spindle and stellate cells in a myxoid stroma with vessels of varying caliber scattered throughout. While pathogenesis is not well understood, it is hormonally responsive and recent studies have identified chromosomal abnormality involving chromosome 12 as one of the causes.

Complete resection with negative margin is the mainstay of treatment. Chemotherapy and radiotherapy are not effective due to scarce mitotic figures. As it is estrogen and progesterone positive GnRH may be given pre and post-operatively. Vascular embolization also plays a role in the management of the cases. Since incidence of recurrence is high, follow-up is advised as long as 15 years.

## Intrabdominal Fibromatosis – An intriguing presentation in gynecological setting

Deepti Goswami, Divya Singh, Varuna Mallya, Nidhi Verma, Gauri Gandhi

#### Case-

18-year old unmarried girl presented with low grade fever and abdominal distension, on workup she was diagnosed as a case of Stage III C mixed germ cell tumor. She underwent staging laparotomy and six cycles of adjuvant chemotherapy (Bleomycin, Etoposide, Paclitaxel). She was discharged and followed up subsequently, on follow up a n abdominopelvic mass of 24 weeks size was noted which kept on growing at a very rapid pace. An exploratory laparotomy was done and the mass was removed which was histopathologically proven as intrabdominal fibromatosis. Post laparotomy there was recurrence and the abdominopelvic mass continued to grow at a very fast speed in view of which weekly chemotherapy with injection methotrexate and vinblastin. She did not tolerate the chemotherapy well and developed severe neutropenia, lung consolidation and she succumbed to these complications after second cycle of chemotherapy.

#### Discussion

Fibromatosis is a slow growing proliferation of fibroblast like cells with no atypia. In a rapidly growing abdominopelvic mass fibromatosis should be considered as a differential and a multidisciplinary team should be involved in management.

## Complete Androgen Insensitivity Syndrome

Patient Miss X, 18 years old with chief complaints of swelling in bilateral inguinal area and primary amenorrhea.

The patient has 4 sisters. On review of history, the two sisters of age 15 years and 13 years respectively, were called to our OPD with similar complaints of bilateral inguinal swellings. On examination,

- Breast development- Tanner stage 3/4, soft no mass palpable
- Pubic hair- very scant, Tanner stage 1
- Axillary hairs- absent

#### **Per Abdomen Examination**

- A soft cystic to firm fluctuant swelling of around 4x4 cm palpable in left inguinal area, non tender.
- Another small mobile lump of size 2x2 cm felt in right inguinal area, firm consistency with no tenderness.

#### **Local Examination**

- External genitalia- normal female, with developed labia majora.
- Blind vagina of 2-3 cm.

#### **Per Rectal Examination**

- Uterus and cervix not felt
- No mass felt in PR examination

#### **Hormonal Profile**

- S. FSH- 3.4 mIU/ml (1.5-12.4 mIU/ml)
- S. LH- 23.5 mIU/ml (1.7-8.6 mIU/ml)
- S. Testosterone- 4.6 ng/ml (male: 2.8-8 ng/ml, female: 0.06-0.82 ng/ml)
- S. Estradiol- 51.04 pg/ml (male:10-40 pg/ml, female: 50-250 pg/ml)
- S. AMH- 54.1 ng/ml (male: 20-40 ng/ml, female: 2-5 ng/ml)

#### Ultrasound

• Uterus and cervix not visualised likely absent. There is e/o solid mass lesion of 3.6x6.5 cm in left inguinal

region with multiple oval hypoechoic nodules showing internal vascularity within, largest nodule around 15.1x6.8 mm, few cystic areas also noted.

• Right inguinal region shows a similar lesion of size 2.2x1.2 cm with multiple hypoechoic nodules with internal vascularity.

Impression: ? undescended testis

FNAC: Likely streak gonads/gonadal dysgenesis possibility of atrophic testes.

#### MRI

- Non visualization of uterus, cervix and upper two third of vagina, distal vagina appear normal.
- No definite ovarian tissue seen at normal expected location
- Oblong ovoid soft tissue intensities areas seen in b/l inguinal region with prominent in left inguinal region with few follicles like structures at periphery D/D gonads (?herniated ovaries).
- Bladder is normal and bilateral kidneys are normal.
- Review masses suggestive of undescended testes

#### **Genetic Testing**

- Karyotype- 46, XY
- AR Gene Analysis
  - Mutation in AR gene, EXON 2 deletion.
     (x:66,863,091-66,863,258) Homozygous zygosity
  - Same mutation in the two sisters with bilateral inguinal masses (46, XY).
- Patient & her parents counselled
- Patient was operated and laparoscopic gonadectomy was done.
- Histopathology of bilateral inguinal masses was suggestive of leydig cell hyperplasia and sertoli cells.
- Complete androgen insensitivity syndrome was first described by John Morris in 1953 and Coined "Testicular Feminization"
- Its is a Disorder of hormone resistance with XY Karyotype and Female phenotype

#### **Disorder of androgen action**

- Complete Androgen Insensitivity (CAIS)
- Partial Androgen Insensitivity (PAIS)
- Mild Androgen Insensitivity (MAIS)
- The mutation can lead a spectrum of complete phenotype female to infertile male
- Four widely investigated mutations

- o Single point mutations
- o Deletions/ insertions
- o Partial or complete gene deletions involving a large part of the gene sequence
- o Mutations involving introns altering RNA splicing

#### Our Patient- Exon 2 deletion- Homozygous

**Differential Diagnosis:** Mayer-Rokitansky-Kuster-Hauser syndrome

Other mullerian duct anomalies—eg, transverse vaginal septae

Management is Individualised and Multidisciplinary.

#### **Treatment:**

- Gonadectomy is delayed until puberty is complete (~16-18 years) and Overall risk of testicular germ cell tumour (TGCT)- ~5-10%. (Seminoma most common) which before puberty is ~0.8-2%
- · Hormone replacement with estrogens

# DR. ASHOK KHURANA

C-584, DEFENCE COLONY \* NEW DELHI – 110024 Consultant in Reproductive Ultrasound

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# The Maze of Knowledge

Abhijeet Kumar, Harpreet Kour Isher All India Institutes of Medical Sciences, New Delhi

- 1. The ultrasonic image below is suggestive of
- a. Amniotic band b. Dichorionic twins c. Monochorionic twins d.TTTS



- 2. Planned vaginal delivery is appropriate for HIV+ mothers when HIV RNA copies /ml are less than
  - a. 50 b. 500
  - c. 1000 d. 200
- 3. Regarding SLE in pregnancy, all the following are true except
  - a. SLE flares up in pregnancy
  - b. Risk of neonatal lupus with one affected child is 10%.
  - c. Fall of complement level more than 25% indicates active lupus
  - d. Titres of ANA do not change with disease activity
- 4. Regarding pre-existing diabetes in pregnancy, all are true except
  - a. Risk of congenital malformations is 25% if HbA1c>10%
  - b. Diabetic nephropathy does not worsen in preganancy
  - c. Diabetic retinopathy can progress in pregnancy
  - d. Fetus tolerates maternal hyoglycemia better than maternal ketoacidosis
- 5. Charecteristic features of acute fatty liver of preganacy are all except
  - a. Profund hypoglycaemia
  - b. Thrombocytopenia
  - c. Coagulopathy
  - d. Marked hyperuricemia
- 6. Which of the following is least likey to be used as a treatment in ITP in pregnancy, remote from term
  - a. Anti D
  - b. IV Ig
  - c. Corticosteroids
  - d. Platelet transfusion

- 7. Intracranial hemorrhage of the fetus can be caused by all of the following in the mother except
  - a. Immune thrombocytopenic purpura
  - b. TORCH infections
  - c. APLA
  - d. Factor VIII deficiency
- 8. Regarding ionising radiation in pregnancy, all of the following are true except
  - a. Fetal exposure to background radiation is greater than 3 mGy.
  - b. Use of lead shielding is able to reduce fetal exposure by 80%.
  - c. Computerised tomography pulmonary angiogram (CTPA) is associated with an increased risk of maternal breast cancer.
  - d. There is a proven link between fetal exposure to diagnostic doses of ionising radiation and childhood cancer.
- 9. The risk of congenital heart disease in the baby of a mother with congenital heart disease is
  - a. 10% b. 1%
  - c. 5% d. 2%
- 10. The above ultrasound image of the fetus showing the distance in callipers measuring 0.91 cm is suggestive of



- a. Normal cistern magna
- b. Abnormal cistern magna
- c. Normal nuchal fold thickness
- d. Abnormal nuchal fold thickness
- 11. All of the following anomalies can be diagnosed on the scan at 11-13+6 weeks, except
  - a. Agenesis of corpus callosum
  - b. Holoprosencephaly
  - c. Hypoplastic left heart
  - d. Omphlocele

- 12.Quadruple test in triplet pregnancy for aneuploidy screening
  - a. Performs better than triple test
  - b. Performs better than dual test
  - c. Not recommended
  - d. Not validated
- 13. Which of the following second trimester soft tissue markers shown in the images increases the risk of Down's the most



14.A pregnant patient develops heart failure without an identifiable underlying cause. I she does not recover baseline cardiac function by 6 months postpartum, which of the following best approximates her 5-year mortality rate?

a.	5%	b.	20%
c.	40%	d.	80%

- 15. Which of the following pregnancy complications is reduced when antihypertensive therapy is started during pregnancy?
  - a. Preterm delivery
  - b. Maternal morbidity
  - c. Development of severe hypertension
  - d. Neonatal intensive care admissions

Watsapp your answers to **9211656757.** Names of first three correct entries will be mentioned in the next issue

- 16. Women receiving therapeutic doses of low molecularweight heparin should not receive neuraxial blockade (e.g., epidural anesthesia) or how long after the last dose was administered?
  - a. 12 hours
     b. 24 hours

     c. 36 hours
     d. 48 hours
- 17. Which of the following viral infections has been associated with a marked increase in the risk or intrahepatic cholestasis of pregnancy?
  - a. Hepatitis C b. Hepatitis B
  - c. Cytomegalovirus
  - d. Human immunodeficiency virus
- 18.Pregnancy physiology results in which of the following changes to factor VIII and von Willebrand factor (vWF) levels?
  - a. Increased factor VIII and vWF levels
  - b. Decreased factor VIII and vWF levels
  - c. Increased factor VIII levels; decreased vWF factor levels
  - d. Decreased factor VIII levels; increased vWF factor levels
- 19. During which of the following epochs in pregnancy is the peak incidence of maternal hypoglycemia noted?
  - a. 10–14 weeks b. 20–24 weeks
  - c. 28–32 weeks d. 34–38 weeks
- 20. Which systemic lupus erythematosus-specific antibody correlates with nephritis and vasculitis activity when seen in high titers?d
  - a. Anti-Ro b. Anti-La
  - c. Antinuclear d. Anti- double-stranded -DNA

Refer page 38 for previous answer key.

# Association of Obstetricians & Gynaecologists of Delhi

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