



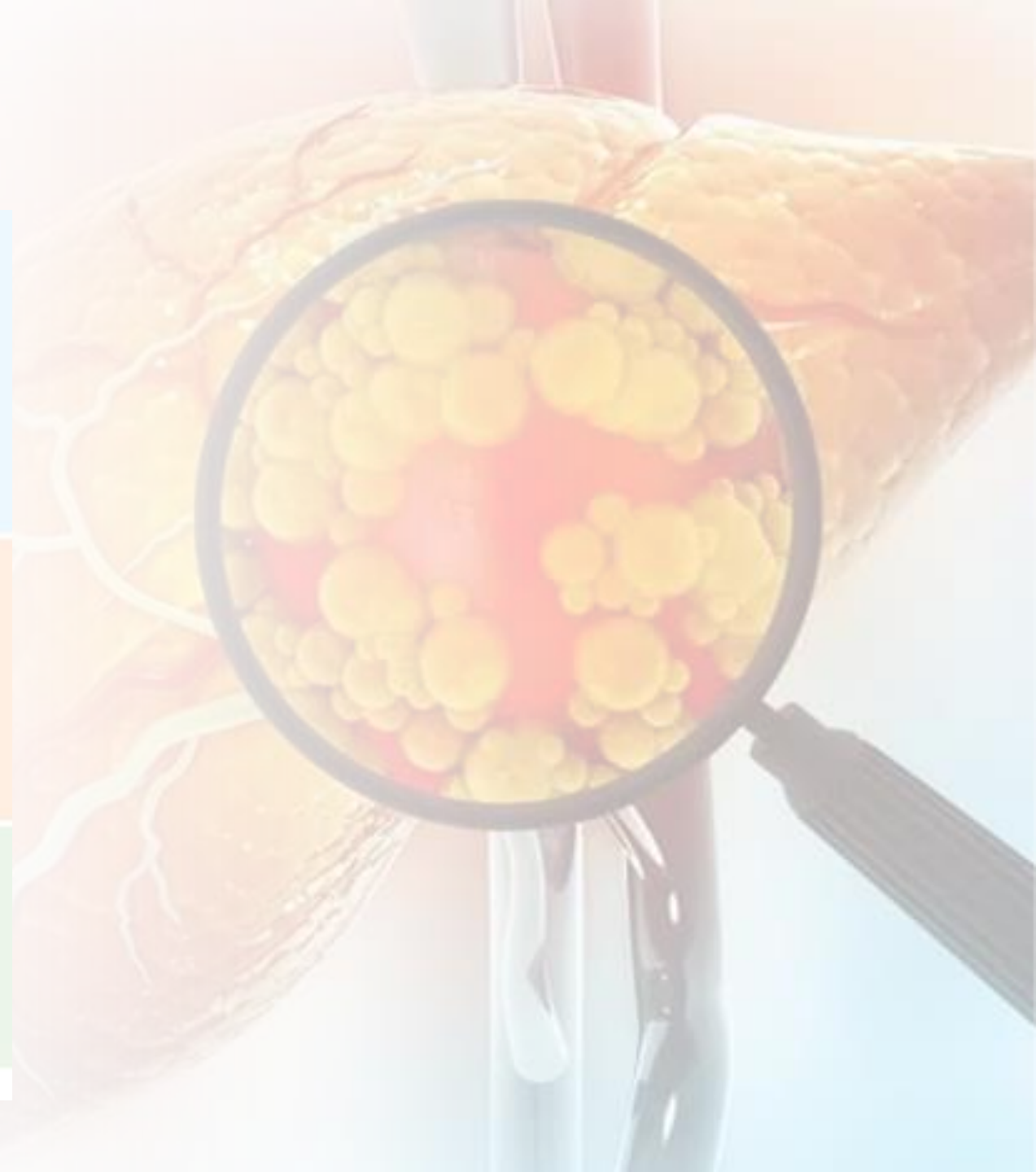
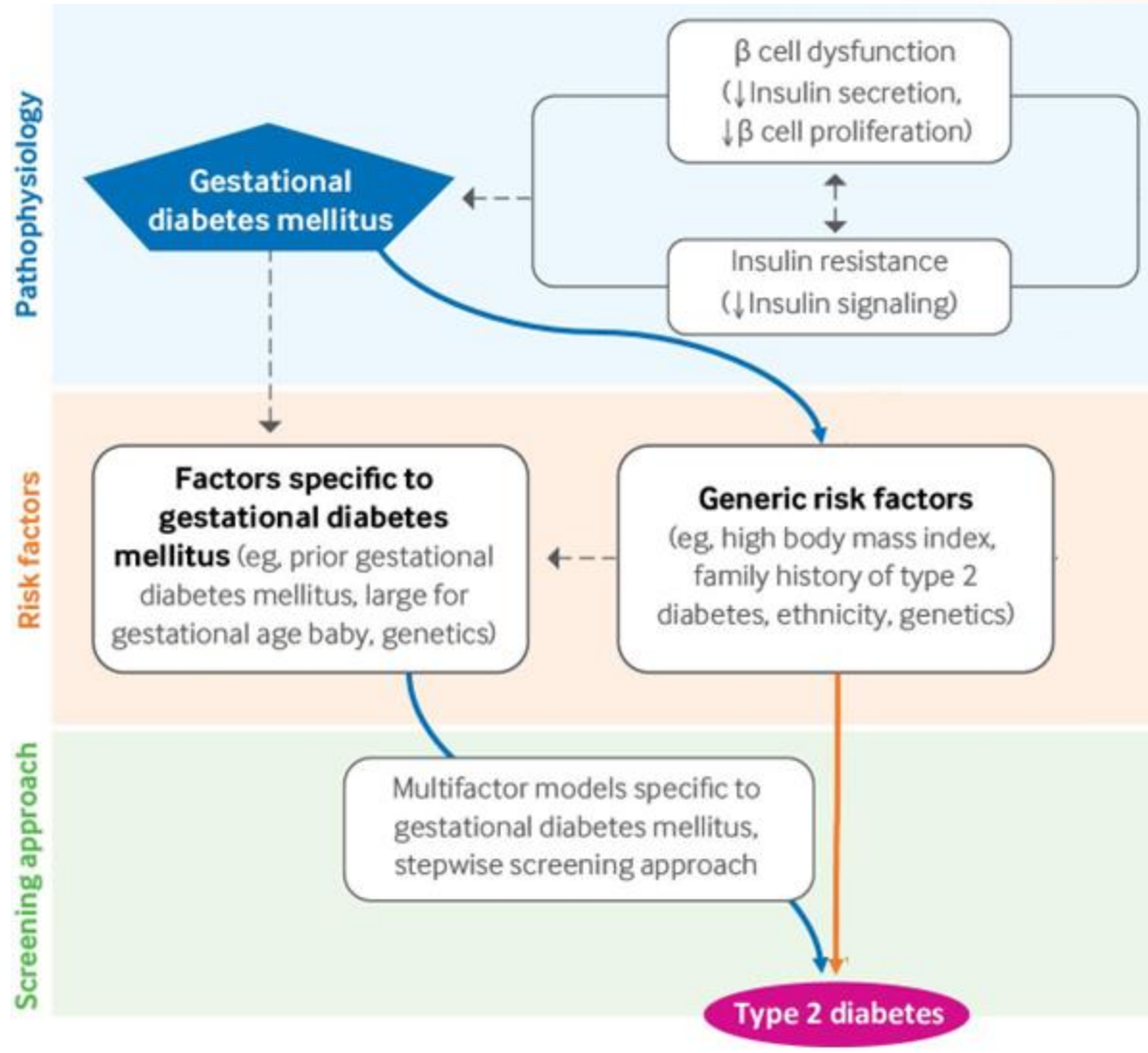
FLiP-GD2 Pilot Study:

Is Fat in the Liver a marker of post-Pregnancy
Glycaemic Deterioration in women with Gestational
Diabetes?

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Gestational diabetes mellitus





Background: Gestational diabetes and fatty liver

• **NAFLD is strongly associated with obesity, insulin resistance, and T2DM.** The general population prevalence: 46%, rising to 74% in T2DM.

• **Relationship Between NAFLD and T2DM-**

- The incidence of T2DM is higher with moderate/severe NAFLD (12.3%) and mild NAFLD (3.4%). 1
- Worsening NAFLD severity linked to the highest T2DM risk (19.2%).
- UK Commission data shows liver disease, including NAFLD, on the rise.
- One-third of overweight individuals have NAFLD; 8% are at risk of fibrosis and have potential for progression to cirrhosis and cancer.



• **NAFLD in Pregnancy**

- Limited assessment of NAFLD in pregnancy and it's link to post-partum glycometabolic status.
- 2002 study, 14/27 (52%) young, obese, non-diabetic women with previous GDM demonstrated NAFLD postnatally that was associated with insulin resistance (3).
- 2011 study on postpartum women, demonstrated NAFLD was more than twice as common in women with previous GDM compared to those without previous GDM (4).

• **Post-Partum NAFLD in Women with Previous GDM**

- NAFLD is twice as common in women with previous GDM.
- It is unclear if NAFLD during pregnancy links to glycaemic abnormalities and insulin resistance post-partum, potentially leading to T2DM.

Study Objectives

	Objectives
PRIMARY	To determine whether the presence of NAFLD during pregnancy in women with gestational diabetes mellitus is a marker of worse metabolic profile (insulin resistance)
SECONDARY	To obtain detailed metabolic profiling in those with NAFLD vs those without NAFLD
	To validate the performance of Ultrasound guided Fatty Liver Index in women with GDM against MRI as a gold standard as a potential screening tool for NAFLD post-partum
	To validate the performance of third trimester ultrasound scan against MRI as a gold standard as a potential screening tool for NAFLD antenatally.

NAFLD: U/S vs MRI

Advantages

- Cost-Effective: Ultrasound is generally more cost-effective compared to MRI.
- Real-time Imaging.
- **Widely Available: Can be done as part of antenatal screening ultrasound**

Limitations

- Operator Dependence.
- Limited Sensitivity: It may **have limited sensitivity in detecting mild to moderate hepatic steatosis.**


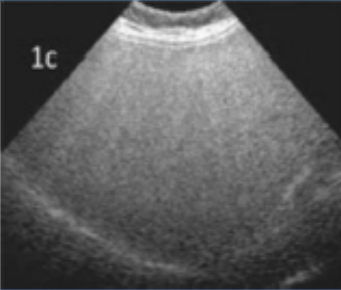
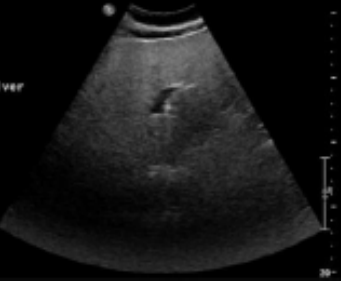
Advantages

- High Sensitivity: MRI is highly sensitive to the presence of fat, making it **effective in detecting even mild hepatic steatosis.**
- Quantitative Assessment: MRI can provide **quantitative measures** of fat content, such as the proton density fat fraction (PDFF).

Limitations

- Cost and Availability:
- Time-Consuming
- **Not performed in pregnancy.**

Ultrasound: Making the subjective objective

Fatty liver Index		
Grading	Criteria	Representative Example
Mild	-Enhanced parenchymal echogenicity - NORMAL visualization of the: <ul style="list-style-type: none"> • Intrahepatic vessel borders • Posterior segments • Diaphragm 	
Moderate	- Increased parenchymal echogenicity - DISTURBED visualization of the: <ul style="list-style-type: none"> • Intrahepatic vessels • Posterior segments • Diaphragm 	
Severe	- Increased parenchymal echogenicity resulting in - POOR/ABSENT visualization of the: <ul style="list-style-type: none"> • Intrahepatic vessels • Posterior segments • Diaphragm 	

Report template:

FLIP – GD2

Parenchymal echogenicity: Normal / Enhanced / Increased

Visualization:

Intrahepatic vessels - YES/ DISTURBED/ POOR /ABSENT

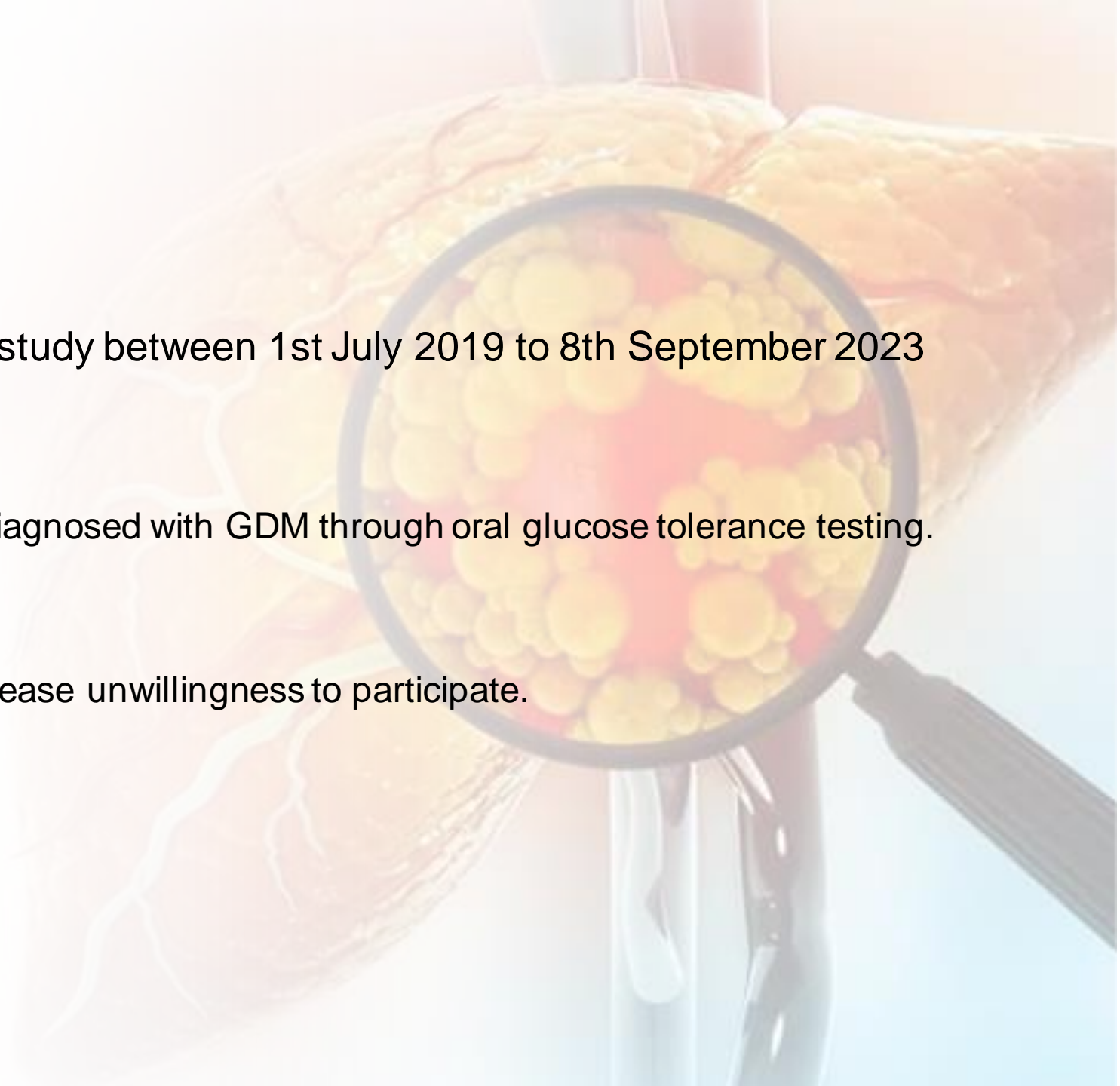
Posterior Segments- YES / DISTURBED / POOR / ABSENT

Diaphragm- YES/ DISTURBED / POOR / ABSENT

Steatosis grading: No observable fatty changes / Mild / Moderate / Severe

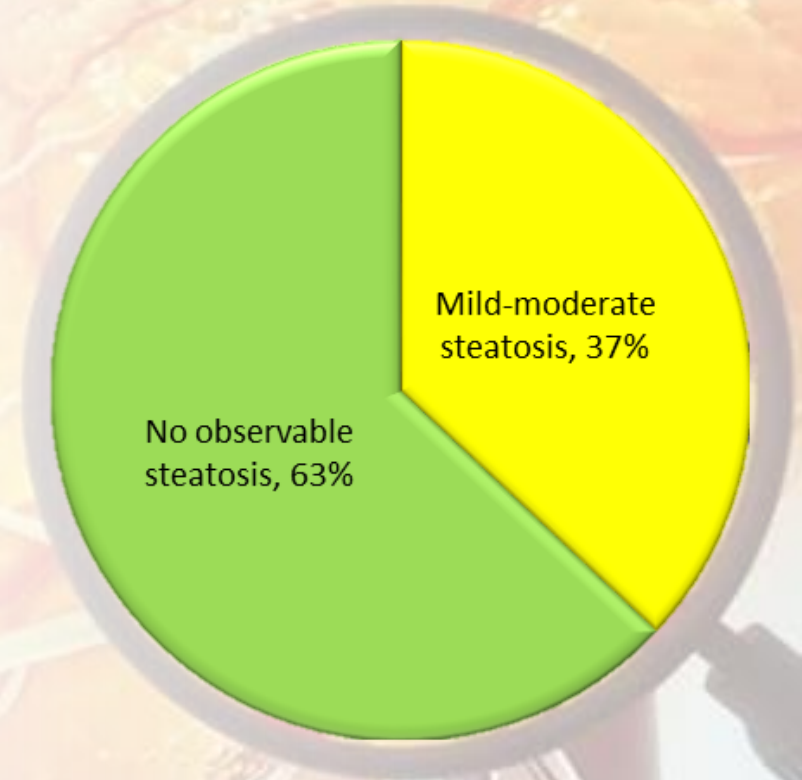


Methods:

- 27 patients were included in the pilot study between 1st July 2019 to 8th September 2023
 - **Inclusion criteria:**
 - Pregnant women aged 18 and above diagnosed with GDM through oral glucose tolerance testing.
 - **Exclusion criteria:**
 - Pre-existing diabetes history of liver disease unwillingness to participate.
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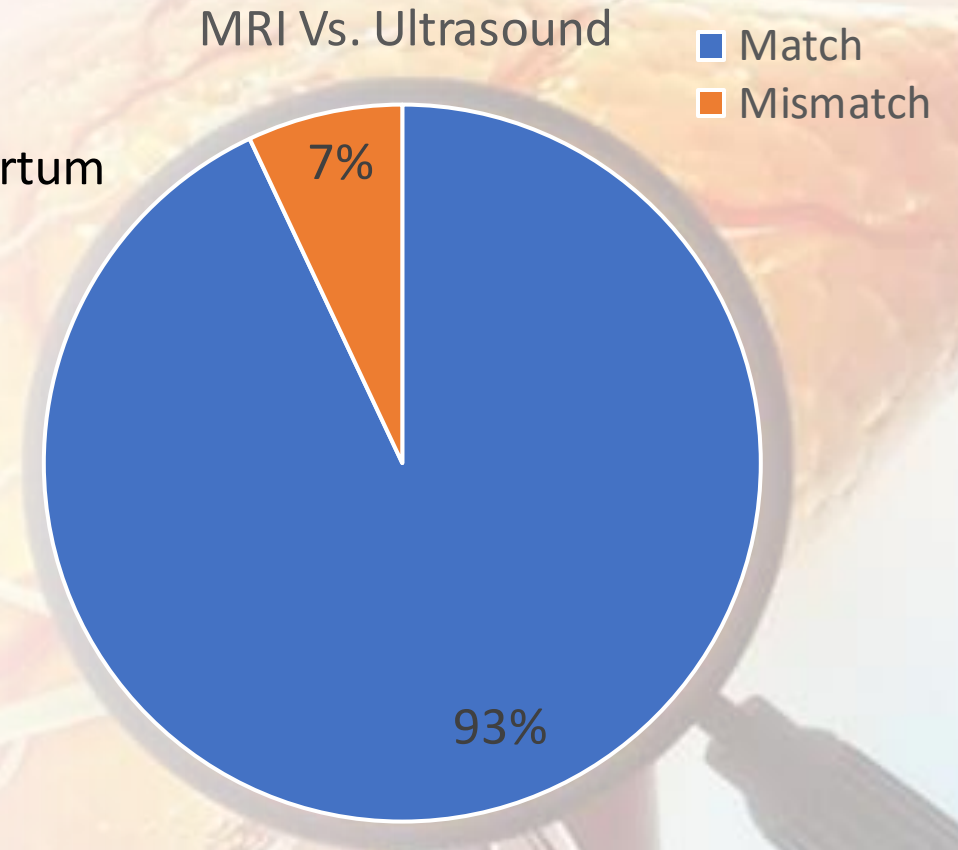
Results: 3rd Trimester Ultrasound Screening

37% of GDM females screened in the third trimester showed positive NAFLD on ultrasound.

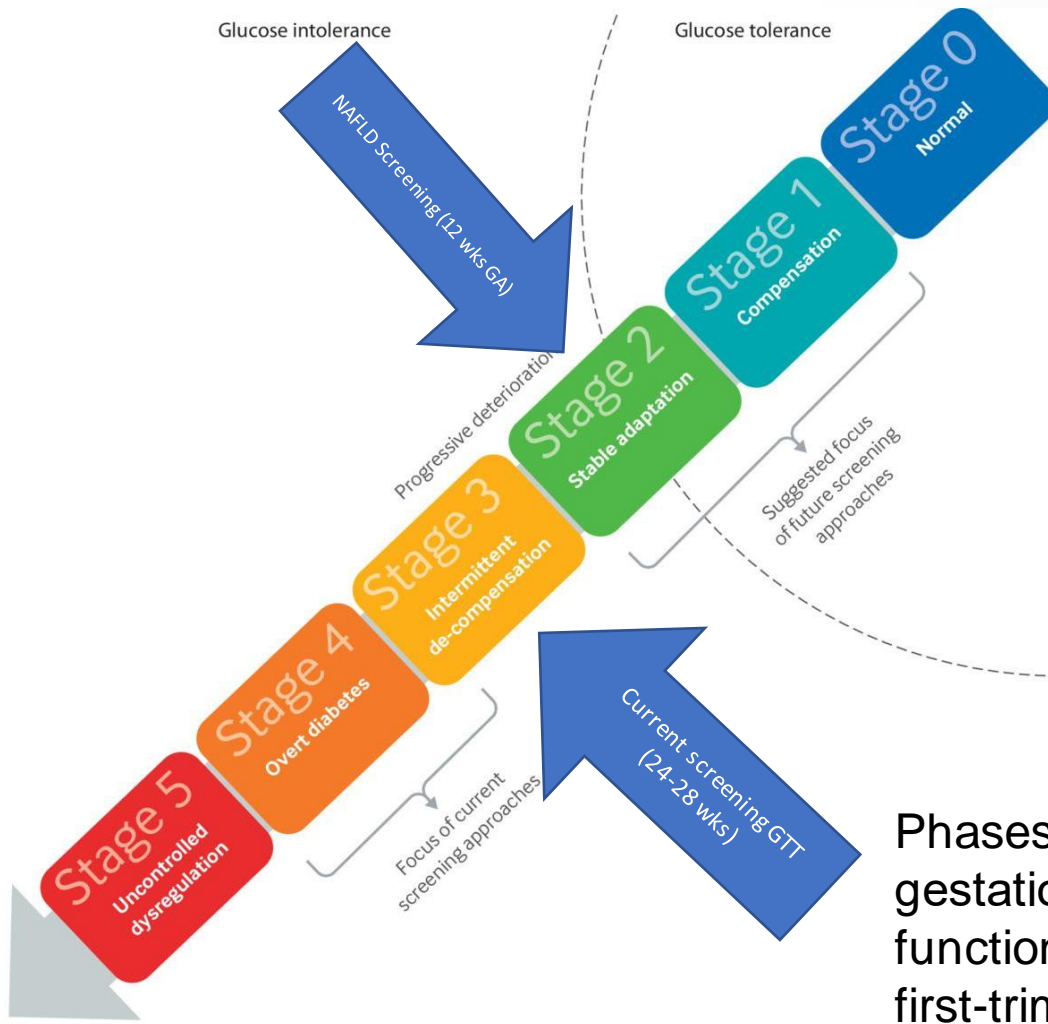


Results: Postpartum MRI

- Correlating ultrasound with MRI- 26 patients underwent postpartum MRI
- Correlation coefficient was 0.901 (Strong positive concordance between ultrasound and MRI)



Impact:



GLYCEMIC DYSREGULATION	
Glycemic status	
Normal	Normal β cell function.
1 - Compensation	Moderate increase in β cell activity, but no glycometabolic consequences.
2 - Stable adaptation	Significant increase in β cell activity with reduced glucose stimulated insulin secretion. No additional stress on glycometabolic system.
3 - Intermittent decompensation	Transient unstable period with sharp rise in glucose levels, when β cell mass drops below a critical level.
4 - Overt diabetes	Persistent decompensation with frank hyperglycemia and severe β cell de-differentiation.
5 - Uncontrolled decompensation	Severe decompensation with progression to ketosis.

Phases of progressive deterioration in glycaemic status in women with gestational diabetes mellitus in relation to insulin resistance and β cell function. Integrating an NAFLD screening ultrasound into the early 12-week first-trimester ultrasound, instead of the current 24-week gestational age Glucose Tolerance Test (GTT) screening, offers the potential to arrest the progressive deterioration leading to gestational diabetes mellitus.



Conclusion:

This study highlights the potential of antenatal ultrasound as a non-invasive, imaging-focused screening tool for NAFLD in pregnant women with GDM.

The high concordance of U/S with postpartum MRI supports its viability as an antenatal screening tool for NAFLD.

Considering NAFLD as an additional predictor for GDM provides an opportunity for early intervention during routine first-trimester scans → holding significant implications given that 70% of GDM patients develop diabetes within 10 years postpartum



References



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