



Ysbyty Maelor Wrexham Wrexham Maelor Hospital

Bwrdd Iechyd Prifysgol Betsi Cadwaladr yw enw gweithredol Bwrdd Iechyd Lleol Prifysgol Betsi Cadwaladr.
Betsi Cadwaladr University Health Board is the operational name of Betsi Cadwaladr University Local Health Board.

Gât 1

Prif Fynedfa A

**Damweiniau ac
Achosion Brys**

 **Parcio Anabl**

 **Maes Parcio**

Gate 1

Main Entrance A

Accident & Emergency

 **Disabled Parking**

 **Car Park**



Smoking is not permitted on this site

haniaateir ysmegu

**Comisiwn
Bevan
Commission**

DECISION MAKING USING PLACENTAL GROWTH FACTOR (PLGF) IN PREECLAMPSIA - PILOT STUDY IN WALES



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Ashwin Ahuja, O&G Clinical Fellow
Lynda Verghese, O&G Consultant
Wrexham Maelor Hospital

- Pilot project
- Test the clinical utility of PLGF point of care blood test in the management of preeclampsia.
- NICE validated tool (20 and 34⁺⁶ weeks)
- This project aspires to positively contribute to the following initiatives:
 1. National Maternity Reviews “Better Births”: Improving Outcome of Maternity Services in England (Feb 2016) ✓
 2. The Five-Year Forward View for Mental Health: Perinatal Mental Health (Feb 2016). ✓
 3. Maternal and neonatal health safety collaborative (Feb 2016) ✓
 4. Maternity Steering Network All Wales ✓



What is PLGF

What is PLGF – Placental Growth Factor



~~6 to 18 weeks :
Remodelling of the uterine spiral arteries~~

Responsible angiogenic growth factors

- VEGF
- Placental growth factor

PLGF is a chemical secreted by the placenta (afterbirth). It is the level of this chemical which helps risk stratify women to appropriately direct surveillance and care provided and optimize the pregnancy outcome.



Sensitivity of commonly used tests for diagnosis of PET

| ≥20w <35w GA | % Sens | % Spec | % PPV | % NPV | OR |
|--------------------|--------|--------|-------|-------|-------|
| PIGF <100 pg/mL | 96.1 | 55.9 | 44.0 | 97.5 | 30.87 |
| Dipstick ≥2+ | 66.7 | 81.0 | 52.6 | 88.4 | 8.50 |
| Uric acid* | 43.6 | 88.0 | 60.0 | 79.1 | 5.66 |
| SBP ≥170 mmHg | 26.7 | 86.3 | 60.0 | 79.1 | 2.22 |
| DBP ≥110 mmHg | 17.3 | 85.8 | 30.2 | 74.5 | 1.27 |
| ALT ≥32 IU/L | 11.3 | 88.7 | 28.6 | 7.14 | 1.00 |

PELICAN study

- A cut-off of 100 pg./mL was used to predict a diagnosis of pre-eclampsia delivering within 14 days
- Sensitivity 96.1% (73 of 76)
- Specificity 55.9% (118 of 211)
- NPV of 97.5% (118 out of 121)
- Women with PIGF ≥100 pg/mL did not have pre-eclampsia needing delivery in 14 days.

PLGF protocol

INDICATIONS

Presentation **between 20 – 34⁺⁶ weeks**

- Isolated new hypertension (>140/90 mmHg)
- Borderline proteinuria (30-50mg/mmol)
- Worsening chronic hypertension /proteinuria in preexisting renal disease/diabetes
- Adjunct to cases with preeclampsia to decide who is likely to need delivery within next two weeks.



METHODOLOGY

- Retrospective and Prospective audit
- 36 cases identified
- Reviewed 34 cases



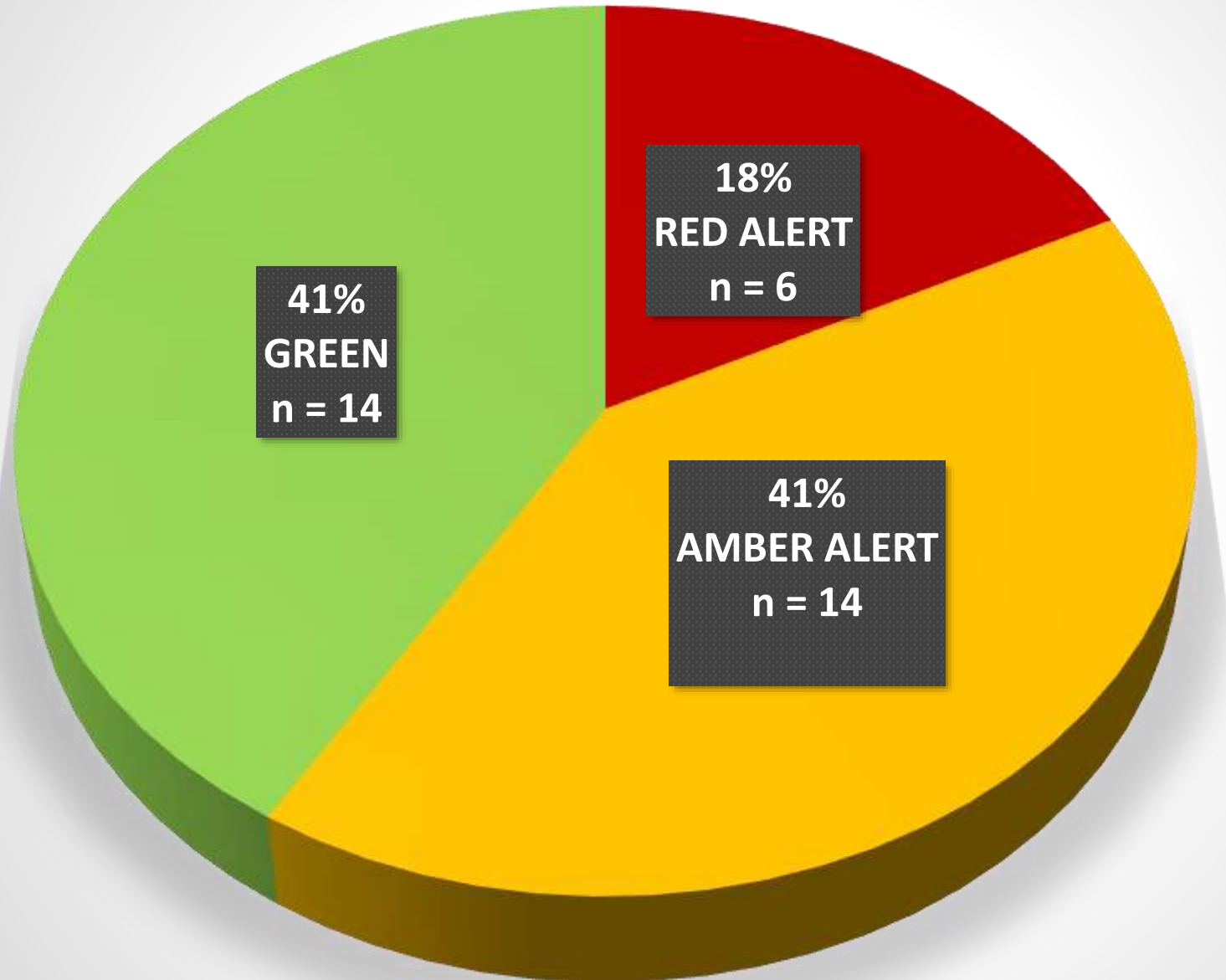
RISK STRATIFICATION

**PIGF ≥ 100
NORMAL**

**PIGF 12-99
LOW**

**PIGF < 12
VERY LOW**

OUTCOMES OF THE PLGF TRIAGE TESTING



- VERY LOW/RED ALERT
- LOW/AMBER ALERT
- NORMAL/GREEN ALERT

Sub-analysis of the VERY LOW PLGF Group

- 6 out of the 34 women had a very low PLGF (<12 pg/ml)
- The commonest indication to do the test was as an adjunct to pre-eclampsia or due to worsening chronic HTN.
- All 6 cases in the red group were admitted, developed severe preeclampsia (2 of which developed HELLP) within the same week, were stabilized, given prophylactic steroids in view of prematurity and were delivered .
- This test helped identify all the high risk patients correctly and helped optimize the maternal and fetal outcomes.

| Gestation presented | Clinical course | Gestation delivered |
|-------------------------------------|---|---|
| G3P1 at 31 wks | GROWTH 3 rd centile HELLP syndrome | IUT to Liverpool , EMCS at 31 wks. |
| Primi at 32 wks. | Growth normal Severe PET , MgSO4 | Delivered at 32 weeks with baby weight of 1.9kg |
| G2P1 at 27wks, essential HTN,BM1=50 | Growth 3 rd centile, PET superimposed on chronic HTN | IUT at 31 wks. –North west and delivered |
| G4P3 at 33 wks | HELLP syndrome, severe PET | Delivered at 33 wks Bt wt 1.8 kg |
| G2P0 at 27+6 weeks | Partial Nephrectomy, Anxiety Depression | IUT to YGC, delivered by EMCS |
| G2P0 at 29+6 weeks | Factor V Leiden Heterozygous | IUT to YGC, delivered by EMCS |

Management

- **PLGF <12 (Red)** – Highly suggestive of placental dysfunction needing early delivery
- **PLGF 12- 99 (Amber)** – Increased outpatient surveillance up to maximum of 37- 38 wks.
- **PLGF \geq 100 (Green)** – Normal and reassuring, can carry on till term if possible.

And why this is important is because.....



One of our patients who was a **third time pregnant mom**, was at **24 weeks and developed severe COVID 19**.

She had to be **cared for in the Intensive Care** after being **intubated**.

However, her **PLGF was in the green i.e. low-risk group**.

We allowed her to continue with the pregnancy – reassured by her PLGF status – and

she **recovered**, was **extubated** and **came out of the ICU**; had a **normal uncomplicated vaginal delivery at 39 weeks**.

Had it not been for the PLGF, she would have had to be delivered at 24 weeks by a Caesarean and the live with the morbidities of extreme preterm delivery.



DISCLAIMER

****INTERPRETATION OF PLGF RESULT****

****LOW OR VERY LOW PLGF RESULT IS NOT AN INDICATION
FOR DELIVERY IN ITSELF****

**PLGF ALONG WITH THE COMPLETE CLINICAL PICTURE
SHOULD BE USED TO DETERMINE THE ULTIMATE
MANAGEMENT OF THE PRE-ECLAMPSIA**



BUDGET
HM TREASURY



STOCKNSHARES / GETTY IMAGES

Financial impact on BCUHB



- Every 24 hr bed day costs taxpayers £ 518
- 2700 deliveries at Wrexham Maelor Hospital – roughly 81 cases of PET/annum
- In 6 months – we had 36 cases – analysed 34
- In this project , 10/34 cases were admitted and 24 cases managed on outpatient surveillance.

- Bed day costs saved = $24 \times 518 \times 3 = \text{£ } 37,296$ (saved in last 6 months)
- Cost of PLGF testing = $88 \times 34 = \underline{\text{£ } 2,992}$
- Total Net Savings £ 34,304
- In addition, SCBU cost/day saved = £ 1,118/day to look after iatrogenic preterm in intensive care
- Medicolegal costs saved in terms of litigation for stillbirth /eclampsia /stroke
- Impact on midwifery and HCA staffing and turnover in labour ward ensuring smooth transition of patient from the induction bay to the labour ward.

Environmental Impact



- Interestingly, Oxford Sustainable Coalition has pointed out that each patient avoiding hospitalization by the use of this PIGF triage testing – the greenhouse gas emissions saved equating to 35 T on CO2 across the Oxford University Trust.
 - ~ 3 million car miles Saved
 - ~ 37 cu. mts less water was used therefore saved.

<https://www.oxfordahsn.org/our-work/strategic-and-industry-partnerships/the-innovation-exchange/spread-and-adoption-of-supported-innovations/pre-eclampsia/case-study-widely-adopted-pre-eclampsia-test-has-additional-environmental-benefits/>



Obstacles & Challenges

| CHALLENGES | HOW WE OVERCAME IT |
|------------|--------------------|
| | |
| | |
| | |
| | |

PLGF Triage testing proved to be of **excellent diagnostic value** in the local cohort of patients with suspected pre-eclampsia.

Prevented maternal complications like eclampsia / stroke / DIC / death and **foetal complications:** stillbirths, prevented iatrogenic preterm deliveries



Provided reassurance for outpatient surveillance and **alleviated maternal anxiety** and allowed patients in this sub-group to safely go home – implications on her and her family's mental and physical health (especially in the pandemic), associated childcare costs and travel costs.



Total net Bed Cost savings from the outpatient management of the 68% of the cases identified in the study period saved the hospital about £ 34,304 in 6 months alone.

SCBU cost/day saved about £ 1,118/day to look after preterm babies in intensive care.



Saved medicolegal costs saved in terms of litigation for stillbirth/eclampsia/stroke.

Impact on **midwifery and HCA staffing** and turnover in labour ward ensuring smooth transition of patient from the induction bay to the labour ward.





Interestingly, each patient that avoids hospitalization due to the PLGF triage test reduces the greenhouse gas emissions equating to 35 T on CO₂ making the project's entire footprint to be near carbon neutral and in fact be greatly beneficial to the environment.

Our Vision for the future



- Roll out PIGF Triage Testing to the 2 sister sites at BCUHB with support from Quidel and Clinical Biochemistry
- Re-audit on all 3 sites when practically possible in the foreseeable future.
- We are working with Action on Pre-eclampsia Charity (APEC) to raise awareness on Pre-eclampsia and PLGF with particular focus in Wales.
- We were enrolled in the PARROT-2 trial and look forward to the outcome of this study to see if we should repeat PLGF testing
- Last but not the least,
'PIGF Triage Testing in the management of suspected pre-eclampsia' is our BEVAN EXEMPLAR PROJECT for pioneering its use in Wales.



STOP
CHECK THE
PRESCRIPTION
CHART BEFORE
GIVING
MEDICATION

Check Agreement for the Submission and Interpretation of
Quality Test Results - Standard 1

| Test | Frequency | Method | Reference Range |
|--------------------------------|-----------|---------------|----------------------------------|
| Glucose | Q1 | Point of Care | 70-100 mg/dL |
| Hemoglobin A1c | Q1 | Lab | 5.7-7.0 % |
| Cholesterol | Q1 | Lab | 125-200 mg/dL |
| Triglycerides | Q1 | Lab | 75-150 mg/dL |
| BUN | Q1 | Lab | 7-20 mg/dL |
| Cr | Q1 | Lab | 0.6-1.2 mg/dL |
| INR | Q1 | Lab | 0.8-1.2 |
| PT | Q1 | Lab | 11-14 sec |
| PTT | Q1 | Lab | 25-35 sec |
| WBC | Q1 | Lab | 4,000-11,000 /mm ³ |
| Hgb | Q1 | Lab | 12-16 g/dL |
| Hct | Q1 | Lab | 37-47 % |
| Plat | Q1 | Lab | 150,000-400,000 /mm ³ |
| Urea Nitrogen | Q1 | Lab | 7-20 mg/dL |
| Creatinine | Q1 | Lab | 0.6-1.2 mg/dL |
| Alkaline Phosphatase | Q1 | Lab | 44-146 U/L |
| Aspartate Aminotransferase | Q1 | Lab | 0-37 U/L |
| Alanine Aminotransferase | Q1 | Lab | 0-40 U/L |
| Lactate Dehydrogenase | Q1 | Lab | 100-250 U/L |
| Bilirubin | Q1 | Lab | 0.1-1.2 mg/dL |
| Prothrombin Time | Q1 | Lab | 11-14 sec |
| Partial Thromboplastin Time | Q1 | Lab | 25-35 sec |
| International Normalized Ratio | Q1 | Lab | 0.8-1.2 |
| White Blood Cell Count | Q1 | Lab | 4,000-11,000 /mm ³ |
| Hemoglobin | Q1 | Lab | 12-16 g/dL |
| Hematocrit | Q1 | Lab | 37-47 % |
| Platelet Count | Q1 | Lab | 150,000-400,000 /mm ³ |

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| Platelet Count | Q1 | Lab | 150,000-400,000 /mm ³ |

Please record
Results in
this book
for all PLRF
TESTS.



DIOLCH



Thank
You

Thank you for a patient hearing

Our Attitude of Gratitude

Dept of Biochemistry , WXM

Dr Yee Ping Teoh, Consultant Clinical Pathologist and Clinical Lead For Biochemistry , WXM

Marge Everall POCT Manager

Ann Marie Parry, POCT Site Lead

Geoffrey Armstrong ,Clinical Biochemist

Gareth Davies , Clinical Biochemist

Quidel

Eoin Morgan -Sales Manager

Dawn Hannah - Clinical Trainer

BCUHB Health Board for business case approval

Dino Tedaldi ,Business and Performance Lead

Maria Atkin, General Manager and Business Lead

Nichola Bray , Financial Advisor

Fiona Giraud – Director of Women’s Services

Last but not the least

All the Midwives, Registrars and Consultants at Wrexham Maelor for implementing it in clinical practice.