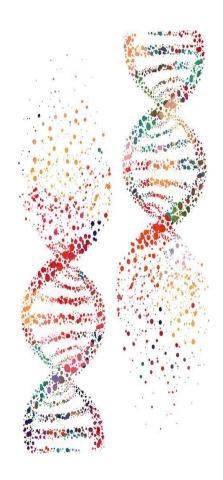
# Machine learning approaches to predict preterm birth by non-invasive means

Samantha L Wilson, PhD

Postdoc Fellow | Princess Margaret Cancer Centre | Hoffman Lab

R Ladies Toronto, June 18th 2019

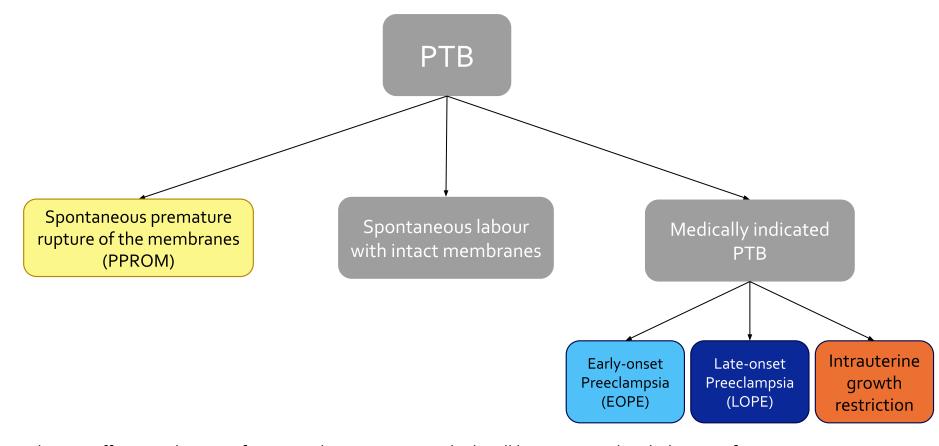


#### R Packages

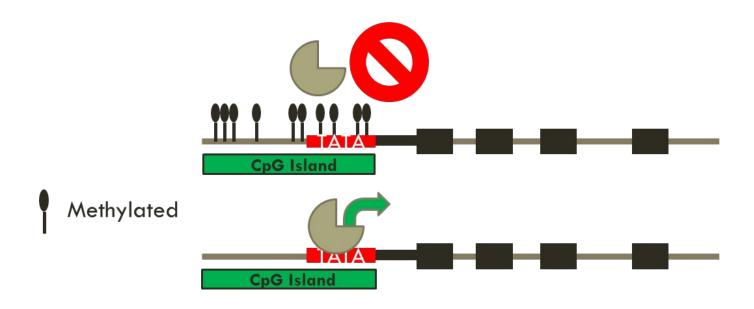
plyr rmarkdown dplyr **RSamtools** knitr stringr limma dendextend ggplot2 impute GridExtra pvclust boot GenomicFeatures reshape2 sigclust2 glm **WGCNA** lm ggjoy MLR lumi carrot

methylumi

#### What is preterm birth and why study it?

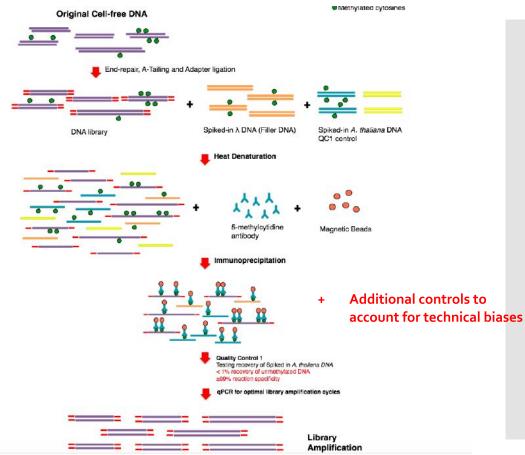


Hypothesis: Different subtypes of PTB are distinct entities which will be associated with distinct cfp-DNAm signatures which we can use to build a predictive model of PTB.



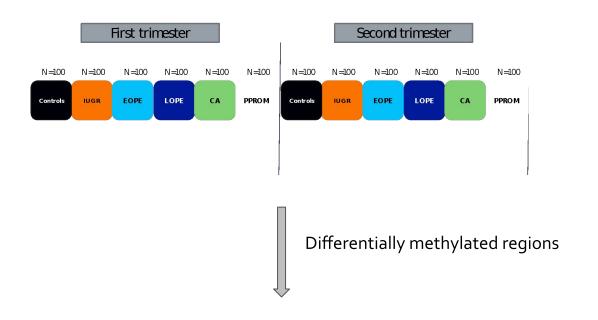
### **DNA** methylation

## Measuring DNAm: cfMeDIP



Shen et al. Nature. 2019.

Aim 2: Identify differential methylation in preterm birth subtypes



**Gene Ontology Analysis** 

Aim 3: Build a predictive model of preterm birth

| Group    | Sample<br>n=67 per<br>group | Clinical features |                                  | DNAm data per 300bp window (Adjusted allele-specific methylation) |      |  |      |  |
|----------|-----------------------------|-------------------|----------------------------------|---|------|--|------|--|
| ١.       |                             | Fetal sex         | Maternal Chronic<br>hypertension | Greater Allele<br>chr1.90300.90600 \ chr1.90601.9                 |      | Lesser Allele<br>10900 chr1.90300.90600   chr1.90601.90900 |      |  |
| Controls | 1                           | М                 | ₹.                               | 0.77  | 0.81 | 0.22   | 0.17 | B. Training data   |
|          | 67                          | F                 | +                                | 0.64  | 0.84 | 0.15   | 0.33 | Numerous predictive<br>models built on                               |
| LOPE     | 68                          | М                 | +                                | 0.98  | 0.77 | NA   | 0.27 | training data  |
|          | 134                         | F                 | +                                | 0.59  | 0.92 | 0.04   | 0.14 |  |
| EOPE     | 135                         | F                 | -                                | 0.64  | 0.81 | 0.17   | 0.34 |  |
|          | 201                         | F                 | +                                | 0.66  | 0.84 | 0.33   | NA   | Multiple-fold cross validation                                       |
| IUGR     | 202                         | М                 | -                                | 0.77  | 0.77 | 0.34   | 0.22 | Tuning data Numerous predictive models tested for                    |
|          | 268                         | <br>М             | =                                | 0.92  | 0.64 | NA   | 0.15 | utility  |
| CA       | 269                         | М                 | 7                                | 0.77  | 0.98 | 0.27   | NA   |  |
|          | 335                         | F                 | -                                | 0.64  | 0.59 | 0.14   | 0.04 | C. Testing data n=33 per group, same                                 |
| SPL      | 336                         | М                 | 8                                | 0.81  | 0.64 | NA   | 0.17 | data type and feature. Withheld from building                        |
|          | 402                         | M                 | +                                | 0.84  | 0.66 | 0.14   | 0.33 | predictive model. Best<br>model is tested on<br>validation model for |

utility