



## Webinar

# Correcting Partial Verification Bias in Diagnostic Accuracy Studies Using R

**Wan Nor Arifin**

PhD Candidate, School of Computer Sciences,  
Universiti Sains Malaysia.

**Organized by**

**Epidemiological and Statistical Modelling Team, USM**

# Background

- Diagnostic test
  - Discriminate diseased vs non-diseased<sup>O'Sullivan et al. (2018)</sup>
  - Extremely important role in medical care<sup>Kosinski & Barnhart (2003)</sup>
  - Objective assessment<sup>Gotzche (2007)</sup>
- Requires evaluation<sup>Linnet et al. (2012)</sup> → Diagnostic accuracy study.

# Background

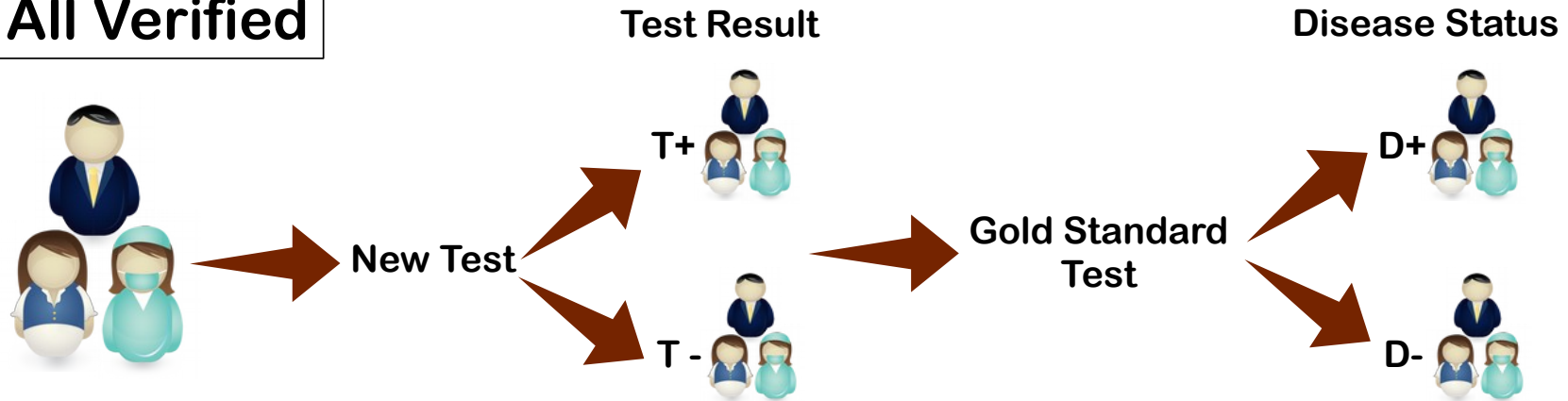
- Diagnostic accuracy study: New test vs Gold Standard<sup>Hall et al. (2019), O'Sullivan et al. (2018)</sup>
  - Covid-19: RT-Ag vs RT-PCR
  - HIV: HIV Rapid Test vs ELISA
  - Breast CA: Mammogram vs Histology

# Background

- Accuracy measures for binary test:
  - Sensitivity (True Positive Rate)
  - Specificity (True Negative Rate)
  - Positive Predictive Value (PPV)
  - Negative Predictive Value (NPV)

# Background

**All Verified**



Test Result	Disease Status		
	D+	D-	
T+	TP	FP	PPV = $\frac{TP}{TP+FP}$
T-	FN	TN	NPV = $\frac{TN}{FN+TN}$
	Sn = $\frac{TP}{TP+FN}$	Sp = $\frac{TN}{TN+FP}$	
	Sensitivity	Specificity	

Positive Predictive Value  
Negative Predictive Value

# Background

- Estimates, esp. Sensitivity and Specificity are often biased.
- Sampling bias in diagnostic accuracy study → Verification bias (VB) O'Sullivan et al. (2018)

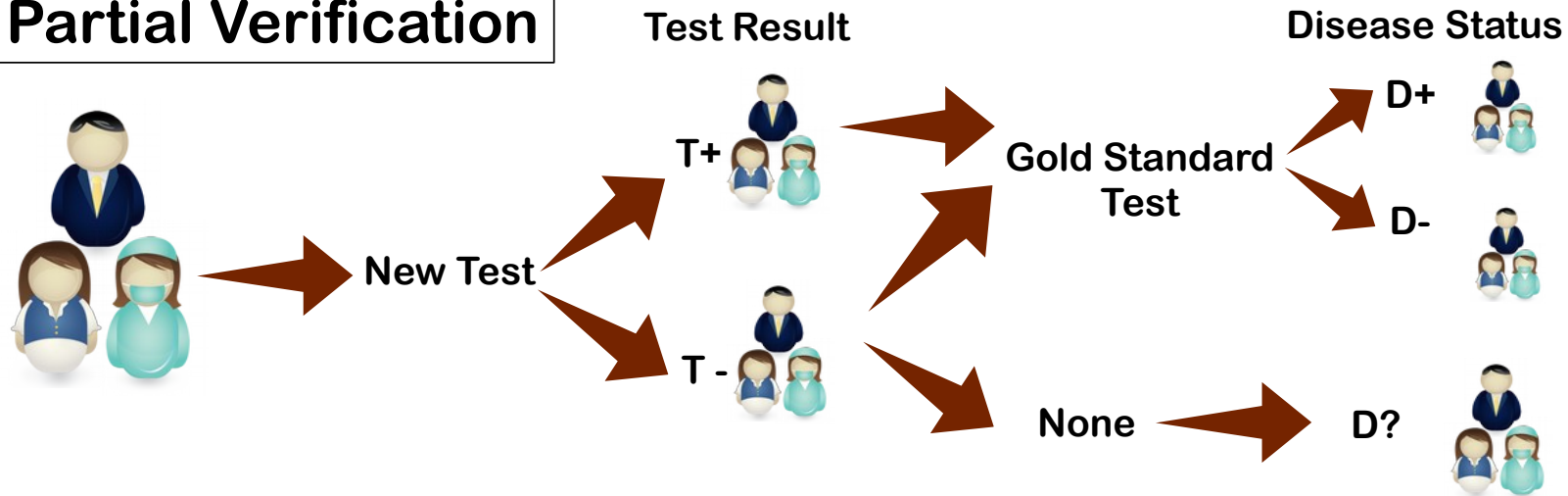
# Background

- Patients are selectively chosen for verification by gold standard.
- Test positive more likely selected + other clinical criteria O'Sullivan et al. (2018)
- Reasons Naaktgeboren et al (2016):
  - Study design: Efficiency, technical, ethical.
  - Clinical practice: Clinical likelihood.
  - Infeasibility: Invasive procedures, postmortem diagnosis.
- Partial and Differential Vb de Groot et al. (2011a)



# Background

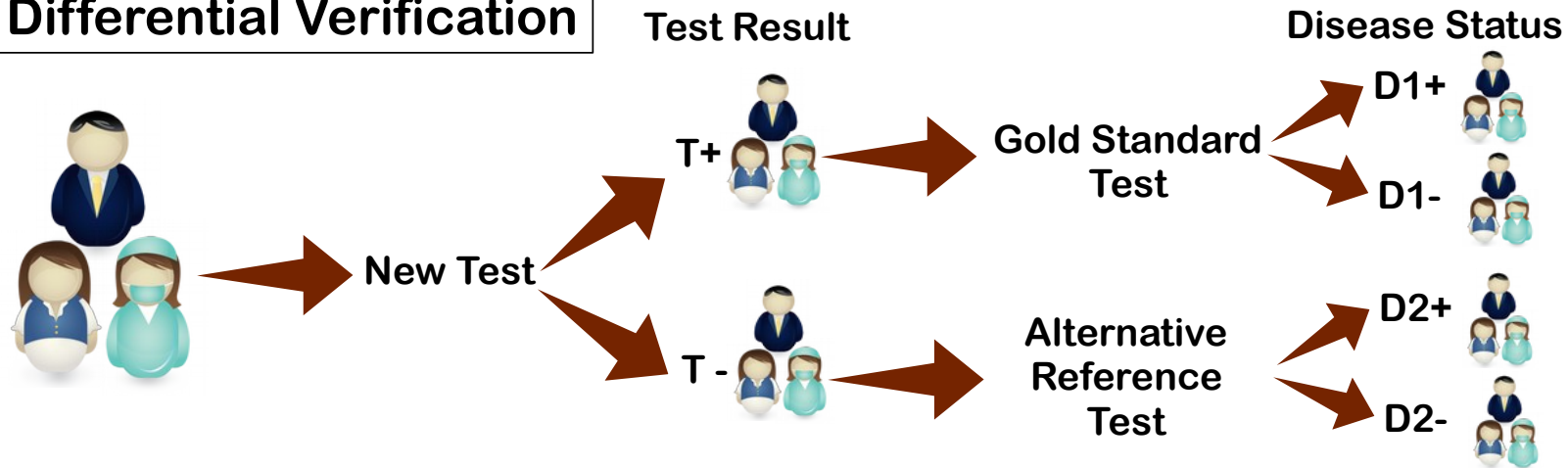
## Partial Verification



Test Result	Disease Status			
	D+	D-	D?	
T+	TP	FP	?	Positive Predictive Value?
T-	FN	TN	?	Negative Predictive Value?
	Sensitivity?	Specificity?		

# Background

## Differential Verification



Test Result	Disease Status (Gold)		Disease Status (Alternative)		Positive Predictive Value? Negative Predictive Value?
	D1+	D1-	D2+	D2-	
T+	TP	FP	TP?	FP?	
T-	FN	TN	FN?	TN?	

Sensitivity?

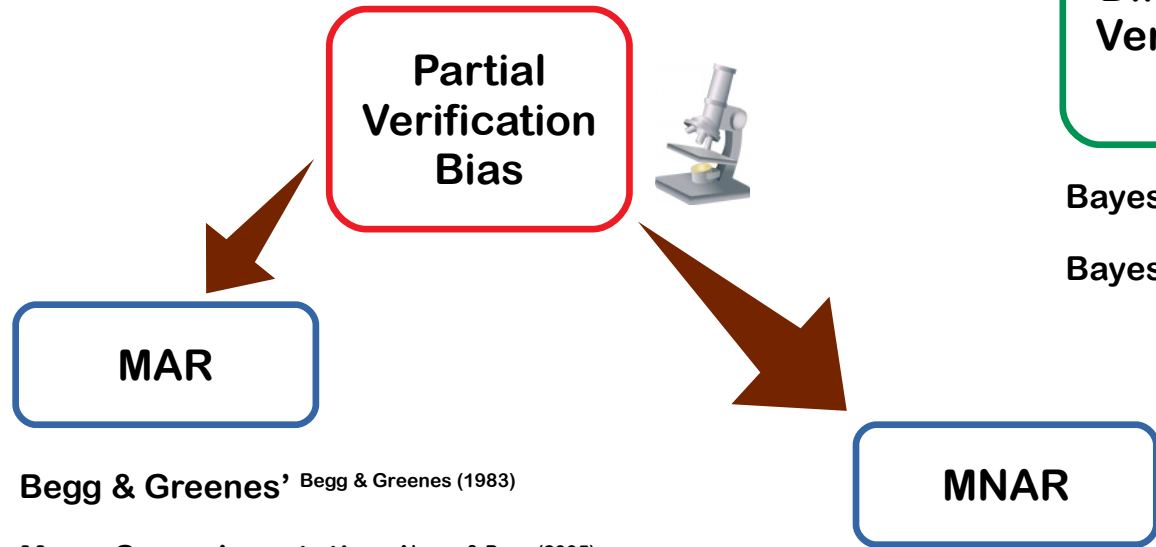
Specificity?

# Background

- VBs → Inaccurate estimates of accuracy measures<sup>Naaktkeboren et al. (2016), Hall et al. (2019)</sup>
- Impact on the clinical practice
  - Invalid diagnostic tests<sup>Chikere et al. (2019)</sup>
  - Clinical errors<sup>Hall et al. (2019)</sup>
- Cannot eliminate verification bias in medical data → Relies on methods to correct VB<sup>O'Sullivan (2018)</sup>

# Literature Review

## Existing Correction Methods



**Partial Verification Bias**



**MAR**

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- Begg & Greenes'** Begg & Greenes (1983)
- Mean Score Imputation** Alonzo & Pepe (2005)
- Inverse Probability Weighting** Alonzo & Pepe (2005)
- Semi-parametric Efficient Estimator** Alonzo & Pepe (2005)
- Multiple Imputation** Harel & Zhou (2006)
- Propensity Score Stratification** He & McDermott (2012)

**Differential Verification Bias**



- Bayesian Approach** Lu et al. (2010)
- Bayesian Latent Class Approach** de Groot et al. (2011c)

- Zhou's Begg & Greenes' Extension** Zhou (1993)
- Logistic Regression** Kosinski & Barnhart (2003a)
- Global Sensitivity Analysis** Kosinski & Barnhart (2003b)
- Neural Networks** Ünal & Burgut (2014)
- Log-Linear Regression** Rochani et al. (2015)
- Bayesian Approaches** Martinez et al. (2006), Buzoianu & Kadane (2008), Pennello (2011), Hajivandi et al. (2018)

**Thank You**

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