

# Using data linkage for vaccine safety surveillance: methods, learnings and impact

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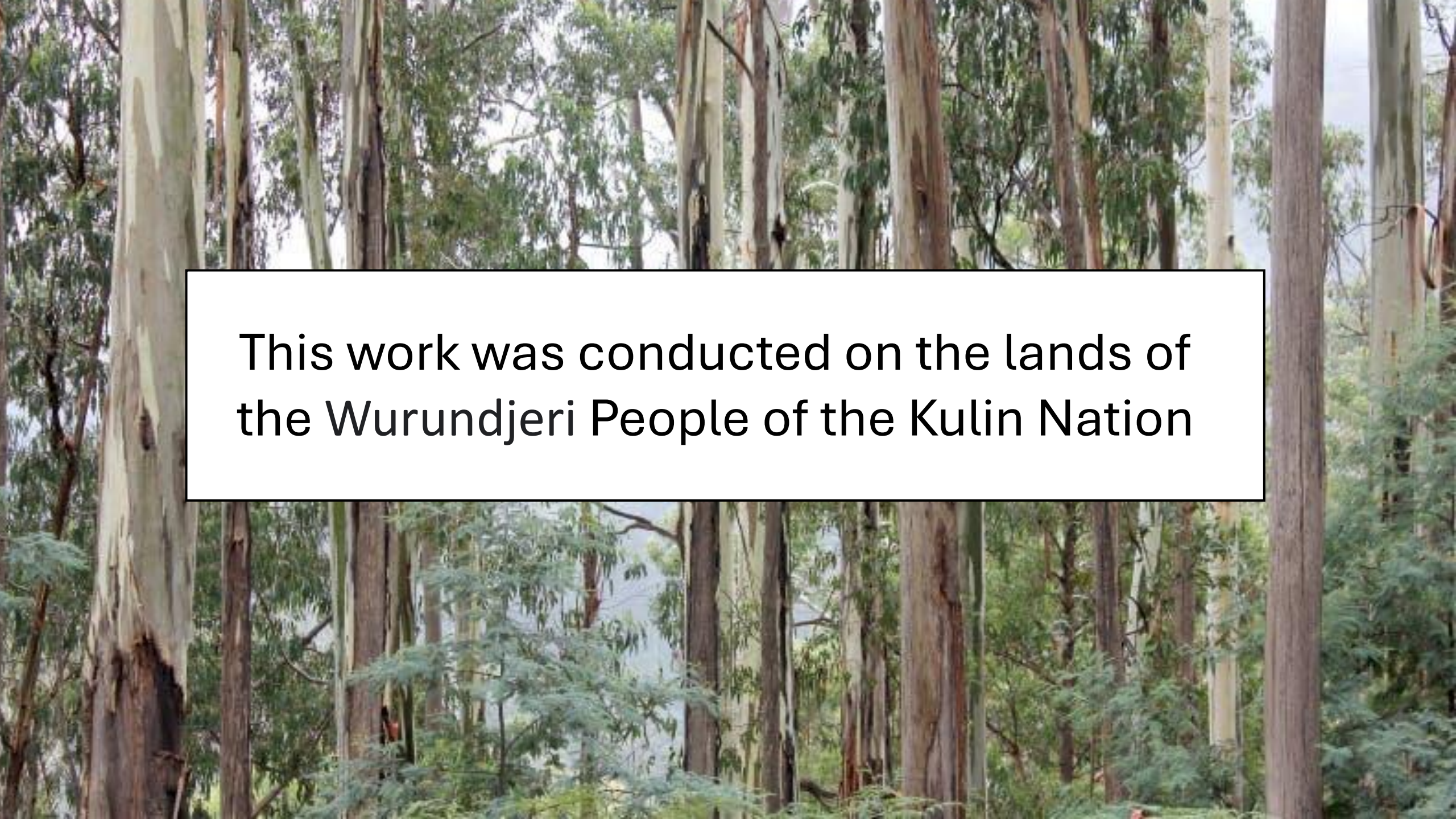
*27/09/24*

*Co-authors: Jim Buttery, James Boyd*

WCE

WORLD CONGRESS OF EPIDEMIOLOGY 2024



A dense forest of tall, slender trees with light-colored bark and green foliage. The trees are closely packed, and the background is filled with more trees, creating a sense of depth. The lighting is natural, suggesting an overcast day.

This work was conducted on the lands of  
the Wurundjeri People of the Kulin Nation



# Adverse Events Following Immunisation (AEFI) surveillance in Victoria, Australia

**Spontaneous surveillance** – reporting via SAEFVIC platform



**Active surveillance** – solicited reports via text and email

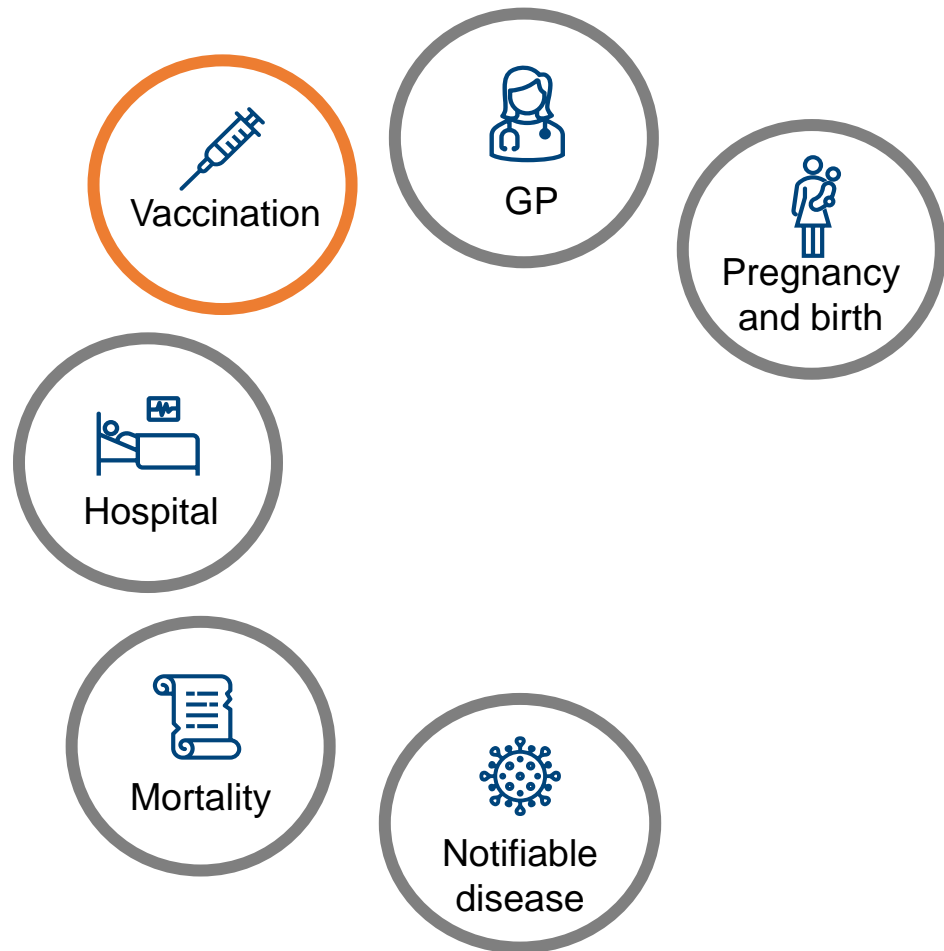


**VACCINE SAFETY  
HEALTH LINK**  
VSHL

- Wider population
- Lower data bias
- Relatively hands-free
  - Cost effective

# VSHL Design

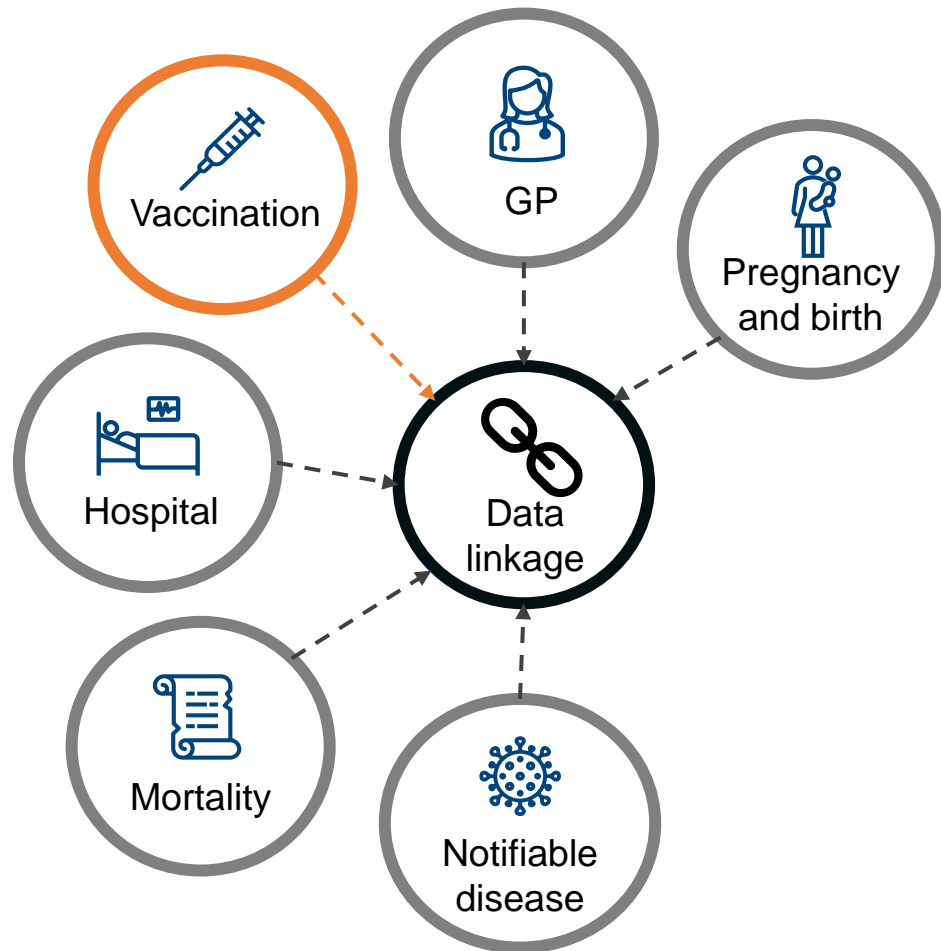
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## Steps of data linkage:

- 1) Standardisation
- 2) Enrichment

# VSHL Design

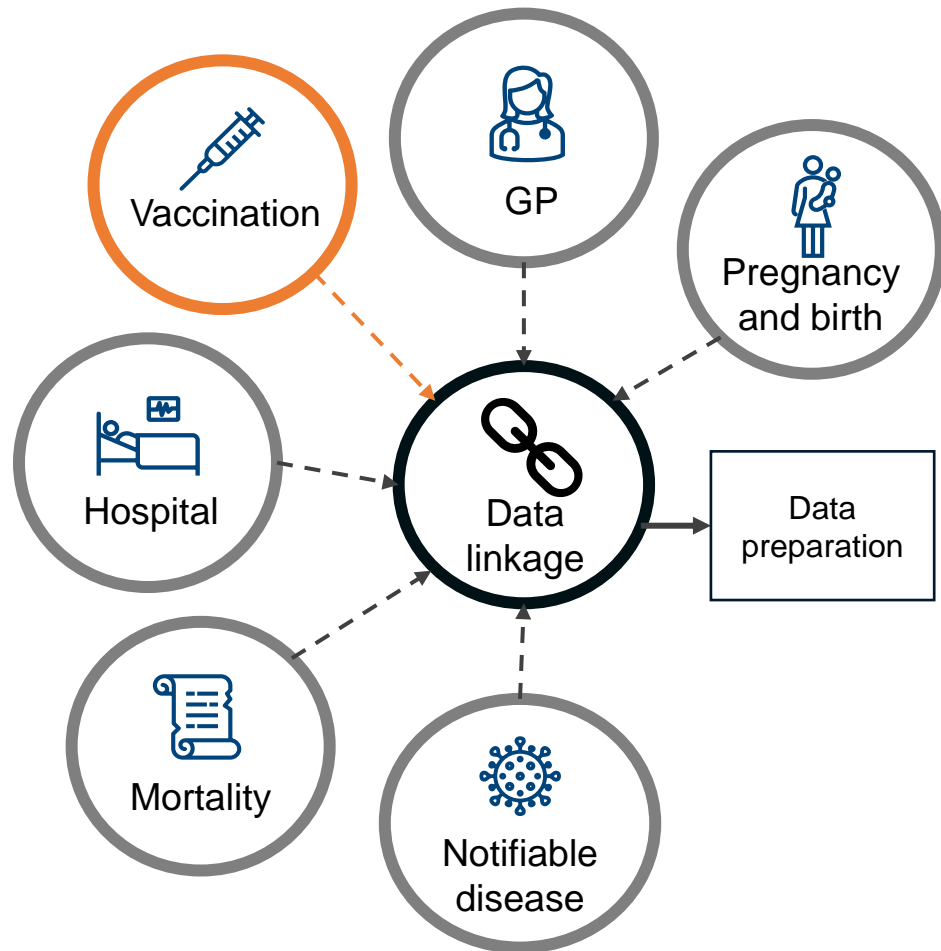


## Steps of data linkage:

- 1) Standardisation
- 2) Enrichment
- 3) Linkage using deterministic and probabilistic methods

Completed at the Centre for Victorian Data Linkage (CVDL) using privacy preserving methods

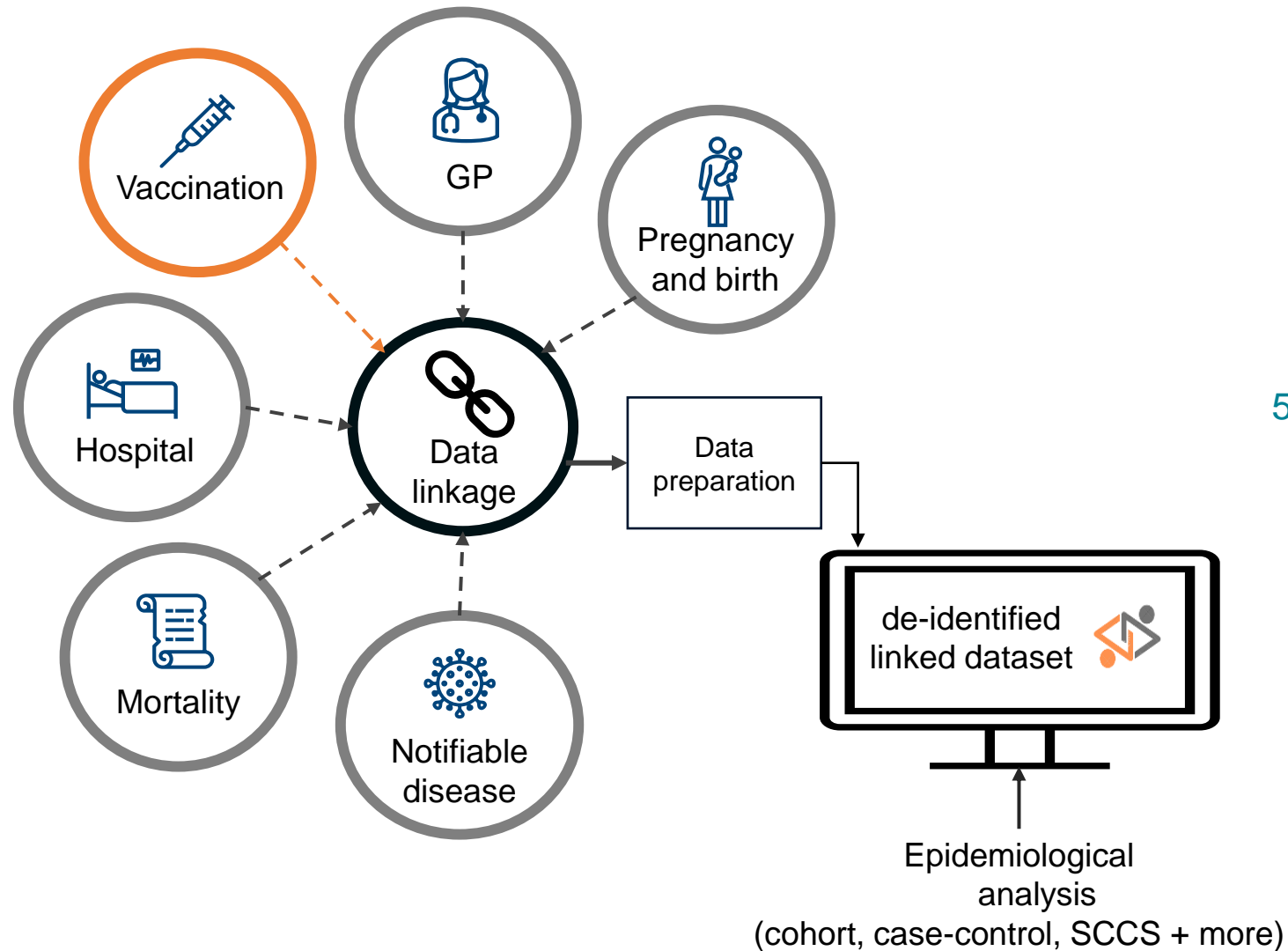
# VSHL Design



## Steps of data linkage:

- 1) Standardisation
- 2) Enrichment
- 3) Linkage using deterministic and probabilistic methods
- 4) Quality assurance
- 5) De-identification and data encryption







# VSHL Design

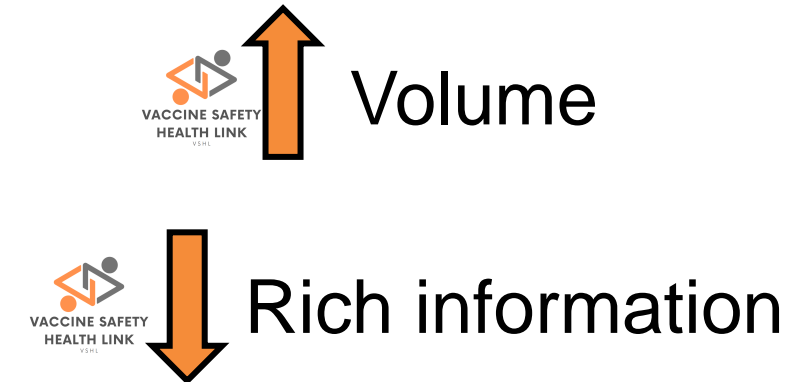


## Steps of data linkage:

- 1) Standardisation
- 2) Enrichment
- 3) Linkage using deterministic and probabilistic methods
- 4) Quality assurance
- 5) De-identification and date encryption
- 6) Data upload and warehousing
- 7) Monthly refresh

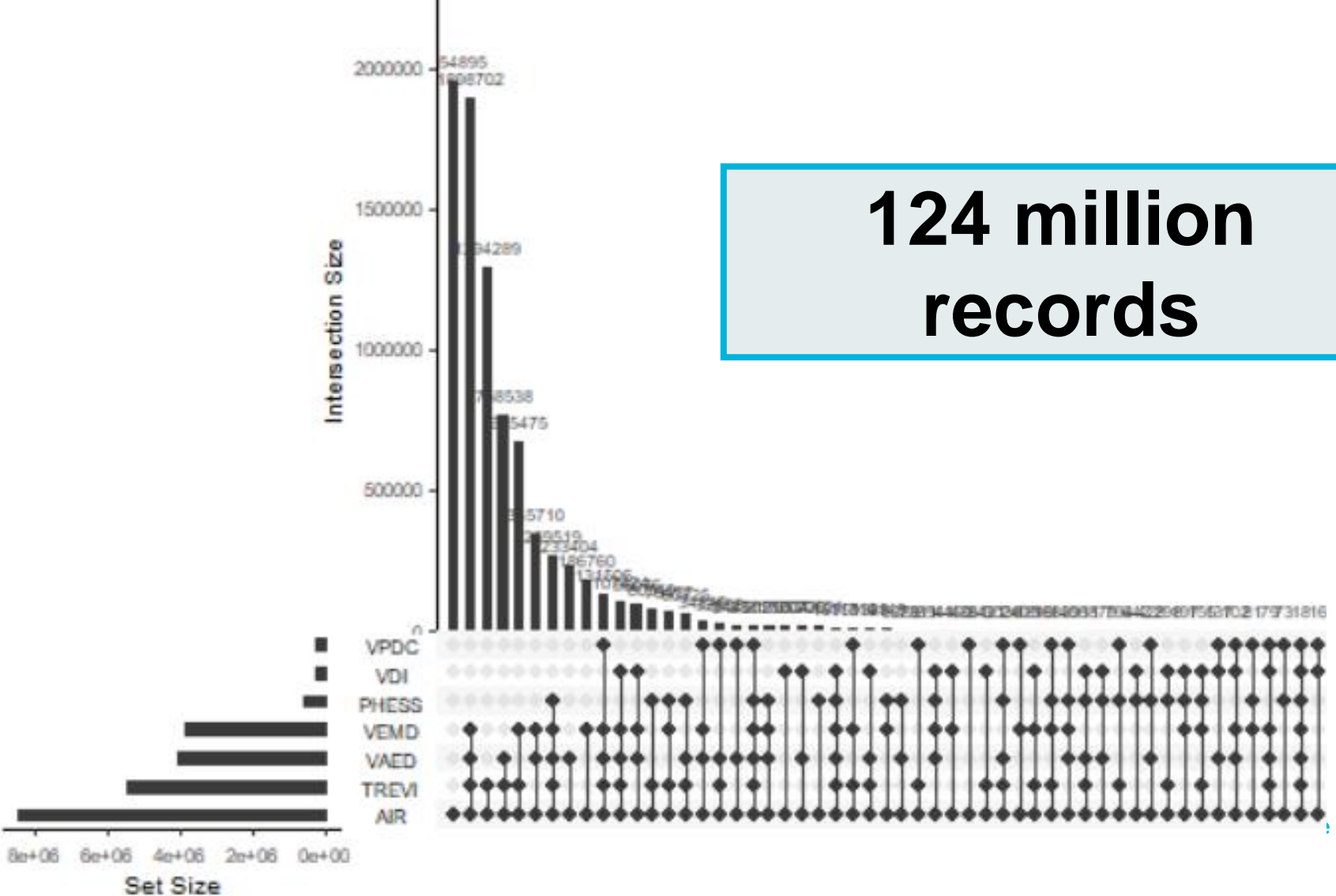
# Data availability

		Healthcare event	
		Yes	No
Vaccination	Yes	  (when notified)  VACCINE SAFETY HEALTH LINK VSHL	 VACCINE SAFETY HEALTH LINK VSHL
	No	 VACCINE SAFETY HEALTH LINK VSHL	 VACCINE SAFETY HEALTH LINK VSHL





# Data volume



# Information for action



Used for **8** local investigations



6 Continents



28 Countries

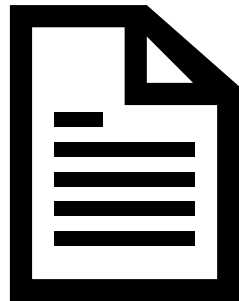


36 Sites



>250 Million People

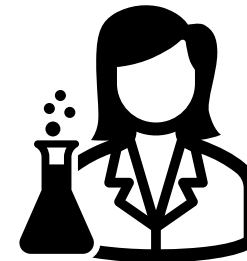
Used for **4** international investigations



Policy



Education



Development

# Lessons learned using VSHL

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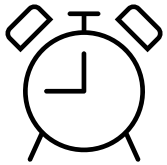
## •Allow enough time

- Governance processes
- Computer processing time with large datasets and many users
- Virtual machine hours
- Importing and exporting code/data

# Lessons learned using VSHL

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## • **Allow enough time**



- Governance processes
- Computer processing time with large datasets and many users
- Virtual machine hours
- Importing and exporting code/data



## • **Remain flexible**

- Different variables
- Encryption

# Lessons learned using VSHL

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## • Allow enough time

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- Computer processing time with large datasets and many users
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## • Remain flexible

- Different variables
- Encryption



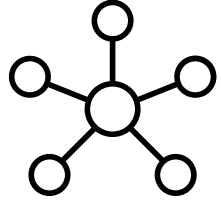
## • Working with large volumes of data

- Pre-processing in SQL
- Creating a playground of dummy data
- Group chat and regular team meetings.



# Lessons learned using VSHL

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## •Explore the data limitations

- Variable completeness and accuracy
- Variable agreement
- Possible data bias with incorrect links

# VSHL Impact

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- Enabling studies that were previously impossible due to underreporting and reporting bias.
- Recommended changes to vaccine policy or healthcare worker education.
- Rapidly integrated into Victoria's routine vaccine safety processes.
- Collaboration with the Global Vaccine Data Network™ (GVDN) extends impact internationally.
- Learnings applicable beyond the scope of vaccine safety.



**Stay in touch!**  
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**Read the detailed paper  
about establishing VSHL!**

# Vaccine safety data linkage in Australia

- 18 different agencies
- 21 separate authorisations
- 12 ethics approvals
- **Australian Government approval took 4 years from initial request**
- Linked data arrived 2 years after grant finished



ARC linkage grant  
2008-2013



## PREVENTABLE DISEASE

**Process trumps potential public good: better vaccine safety through linked cross-jurisdictional immunisation data in Australia**

Katherine M. Duszynski,<sup>1,2</sup> Nicole L. Pratt,<sup>3</sup> John W. Lynch,<sup>2</sup> Annette Braunack-Mayer,<sup>2,4</sup> Lee K. Taylor,<sup>5</sup> Jesla G. Berry,<sup>1</sup> Vicki Xafis,<sup>2</sup> Jim Buttery,<sup>6,7</sup> Michael S. Gold<sup>1</sup> on behalf of the Vaccine Assessment Using Linked Data (VALID) Working Group



# Collaborative Western Australia and Victoria analysis for Guillain Barre Syndrome

**Example:** self-controlled case series (SCCS) analysis conducted to measure association between adenovirus vector vaccine and Guillain Barre Syndrome.

**VIC RI:** 2.71 (1.01, 3.86)  $p < 0.0001$

**WA RI:** 1.04 (0.38, 2.87)  $p = 0.94$

## Meta-analysis approach

Independent analysis in each state and combine final result

**Collaborative meta-analysis approach:**

**2.45 (1.76, 3.41)  $p < 0.0001$**

## Combined de-identified linelist approach

Each state gets own linelist with exposure and outcome data, de-identifies the linelist and merges to create one large linelist for analysis

**Combined linelist approach:**

**2.18 (1.55, 3.07)  $< 0.0001$**

## Combined linkage approach

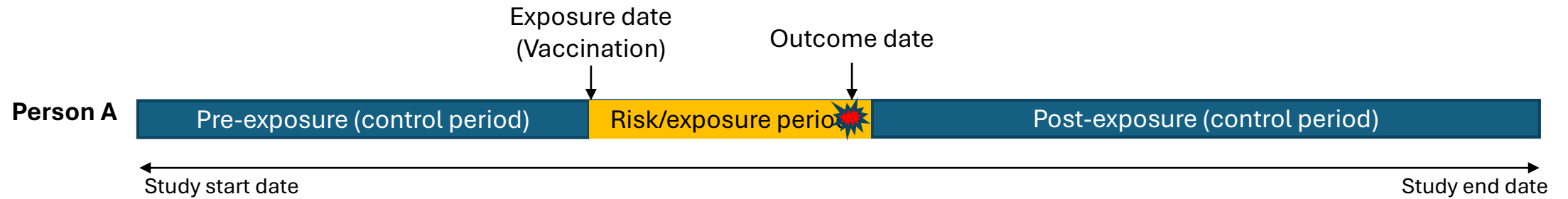
Secure repository of data from each state with common unique identifier to create one linelist where exposure and outcome records between jurisdictions can be connected. Analysis is conducted on the one large linelist

**Not testing**

**CONFIDENTIAL**

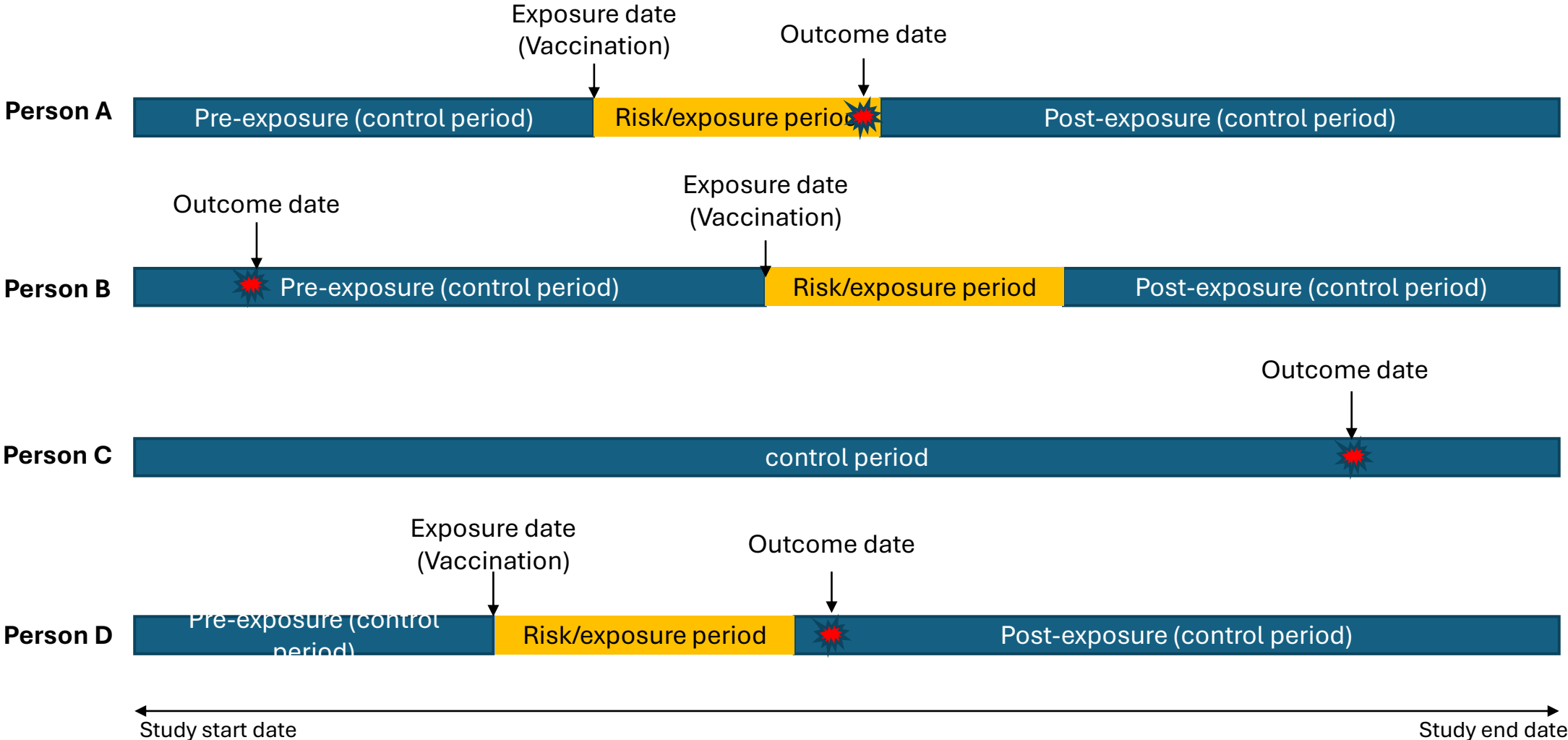
# Self-controlled case series (SCCS)

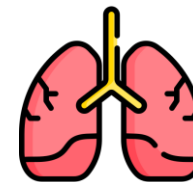
For every person with the outcome in study period



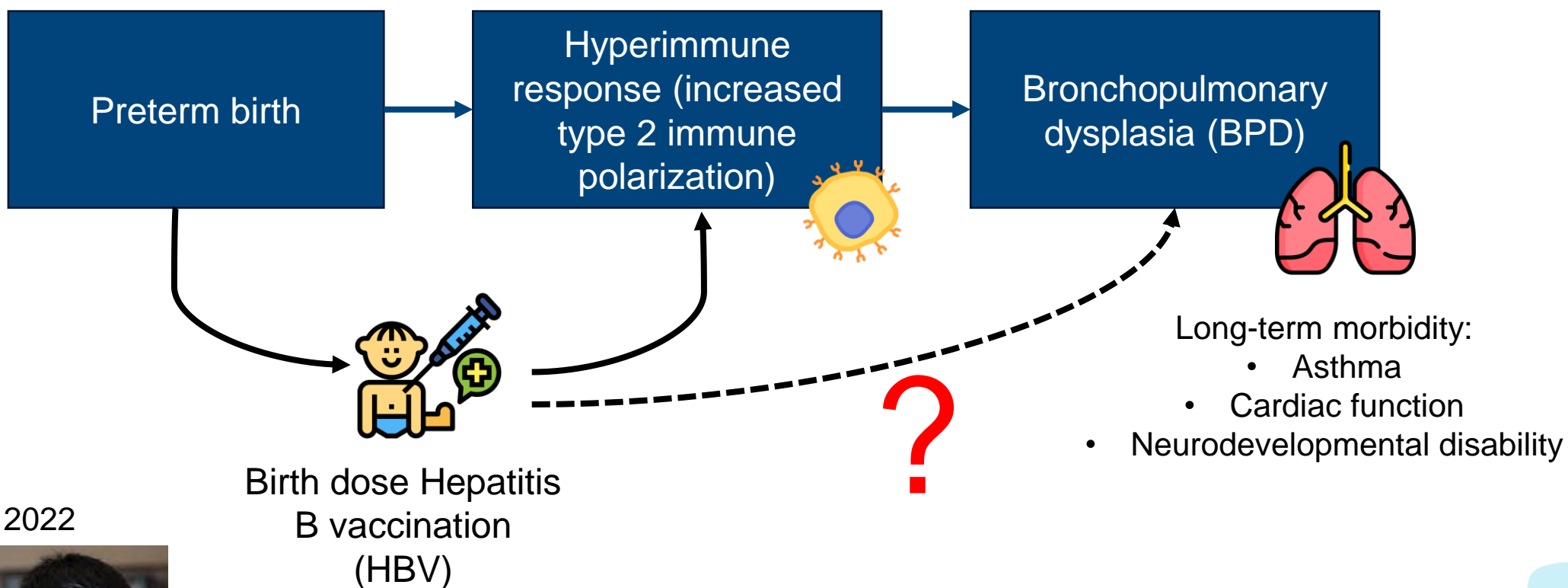


# Self-controlled case series (SCCS)





# Study background

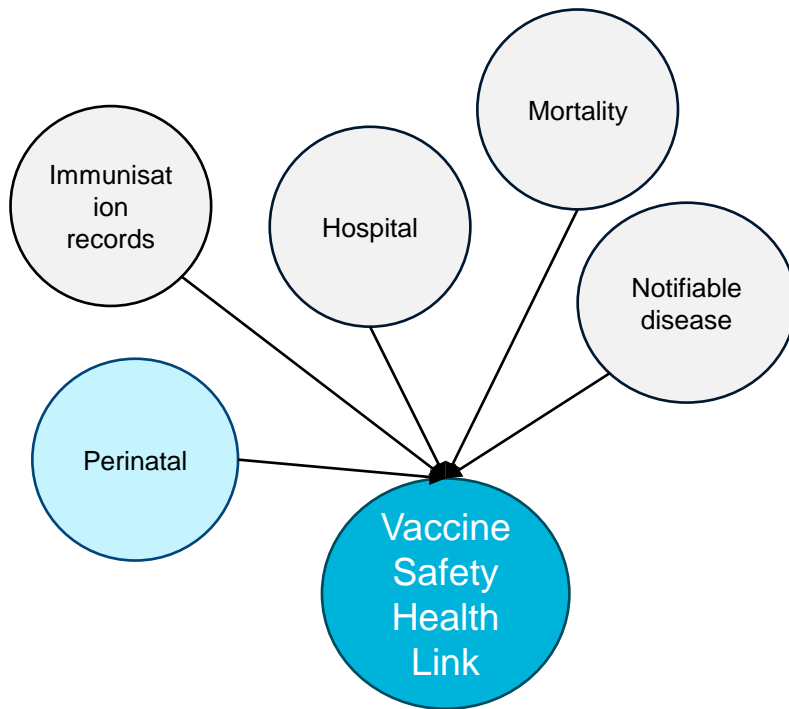


Lao et al. 2022



# Study methods

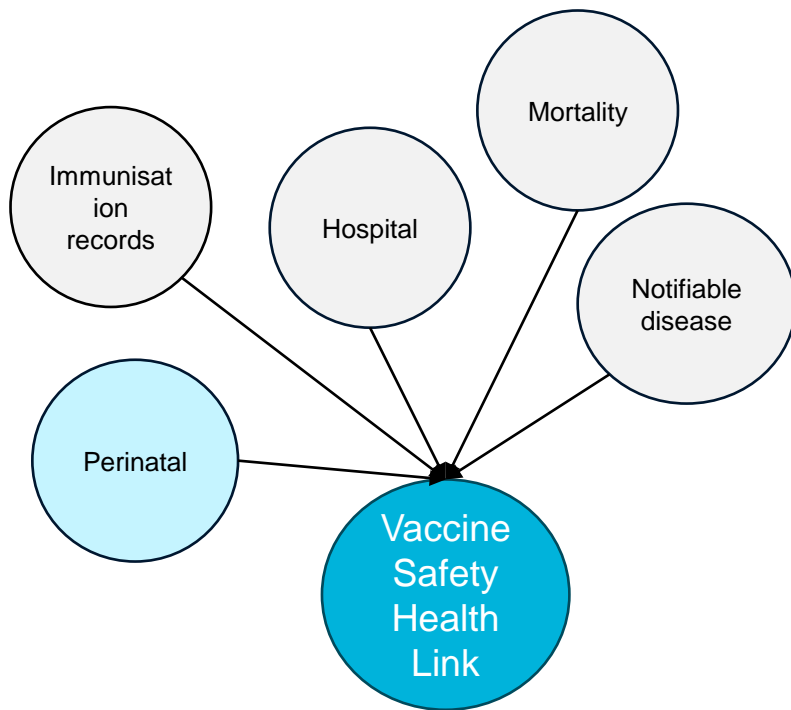
**Step 1) Cohort – preterm infants born alive at <29 weeks gestation between 2017 and 2021 as per the Victorian Perinatal Data Collection**



Identifier	Gestation	Date of birth encrypted
Infant A	26	15 May 2020
Infant B	24	17 Feb 2017
Infant C	28	2 Sep 2019
Infant D	28	3 Oct 2019

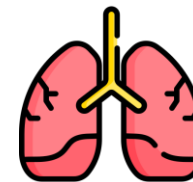
# Study methods

## Step 2) Exposure – HBVV vaccination less than 24 hours as per the Victorian Perinatal Data Collection



Identifier	Gestation	Date of birth encrypted	Vaccination status
Infant A	26	15 May 2020	1
Infant B	24	17 Feb 2017	0
Infant C	28	2 Sep 2019	1
Infant D	28	3 Oct 2019	1

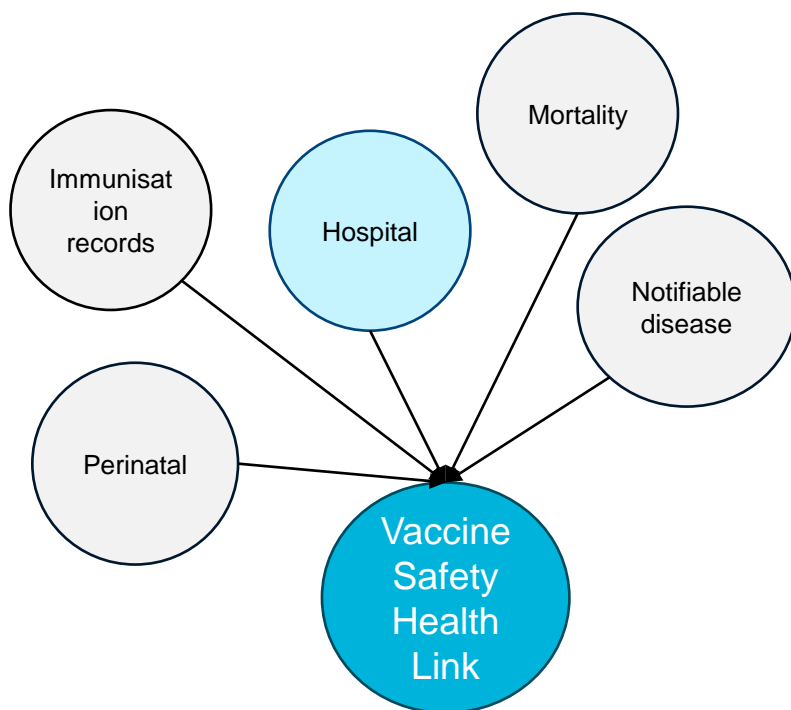
# Study methods



VSHL Example

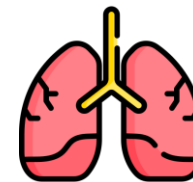
## Step 3) Outcome – Bronchopulmonary dysplasia ICD-10-AM code as per the Victorian Admitted Episodes Dataset

P27.1, P27.8, P27.9  
ICD-10-AM codes.  
Assigned where  
supplemental oxygen  
is required at 36  
weeks gestation  
corrected.



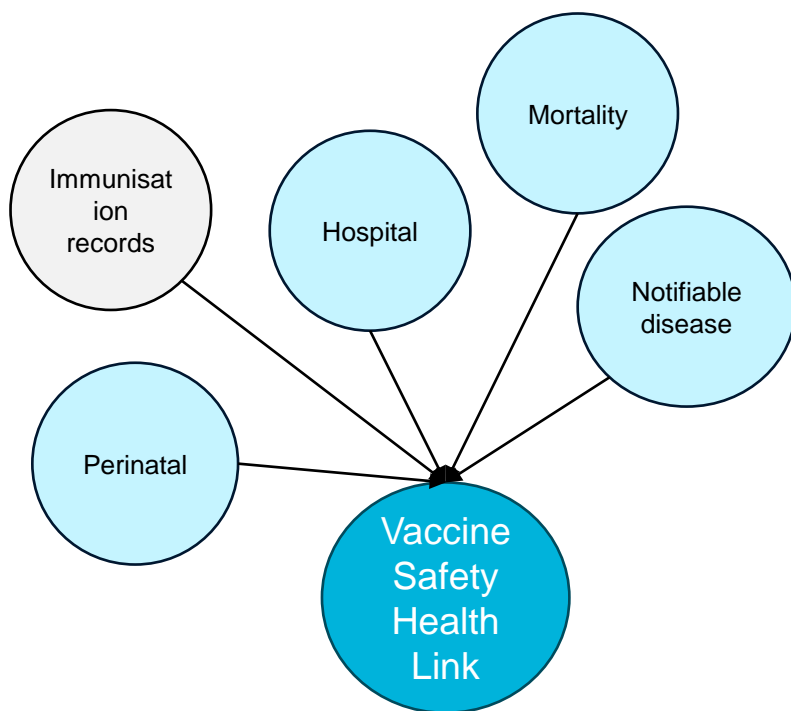
Identifier	Gestation	Date of birth encrypted	Vaccination status	BPD status
Infant A	26	15 May 2020	1	1
Infant B	24	17 Feb 2017	0	0
Infant C	28	2 Sep 2019	1	0
Infant D	28	3 Oct 2019	1	1



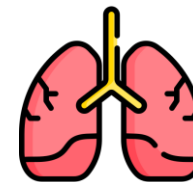


# Study methods

## Step 4) Confounders – Identified via directed acyclic graph in consultation with neonatologists



					Confounders					
Identifier	Gestation	Date of birth encrypted	Vaccination status	BPD status	A	B	C	D	E	F
Infant A	26	15 May 2020	1	1	x	x	x	x	x	x
Infant B	24	17 Feb 2017	0	0	x	x	x	x	x	x
Infant C	28	2 Sep 2019	1	0	x	x	x	x	x	x
Infant D	28	3 Oct 2019	1	1	x	x	x	x	x	x

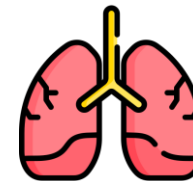


# Study results

		BPD		
		Yes	No	Total
Hepatitis B vaccine	Yes	155	151	<b>306</b>
	No	317	195	<b>512</b>
	Total	<b>472</b>	<b>346</b>	<b>818</b>

Unadjusted relative risk *0.81 (0.67,0.98)*  
Adjusted relative risk *0.83 (0.69,1.00)*

No evidence of increased risk of BPD in preterm infants who received HBVV birth dose compared to those who did not



# Study impact

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- National importance to address this question about risks of HBVV timing in pre-term infants
- Victoria with the Vaccine Safety Health Link is uniquely placed to investigate this potential association – large sample of real-world data.
- Findings support existing World Health Organization recommendations to immunise all infants against hepatitis B within 24 hours of birth
- Findings have been communicated to researchers and neonatologists. Helps address local practice.