

Post vaccination febrile seizures: Clinical severity and outcome data is reassuring

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Outline

- Background on febrile seizures
- Study aims and methods
- Findings and discussions
- Conclusions and future direction



Background: Febrile seizures (FS)

- Caused by sudden change in one's body temperature
- Most common type of seizure in childhood
 - 1 in 30 children
 - 6 months and 6 years of age
 - Peak incidence in the second year of life
 - 30% will have a second episode
- No known long term neurological effect

Background: Post-vaccination febrile seizures (PVFS)

- Associated with
 - Whole cell pertussis vaccine
 - Measles containing vaccine
 - Trivalent influenza vaccine
- Decrease parent and provider confidence on vaccine safety



Clinical question

- Child presents with FS following 12 month old MMR vaccine
 - Is this child's FS any different to a FS due to another cause (i.e respiratory illness)?
 - Will they have another FS with subsequent vaccinations?

Aims

- Describe the epidemiological profile of children with post-vaccination FS (PVFS) and non-PVFS
- Describe the clinical severity and outcomes of PVFS and non-PVFS cases
- Describe the recurrence rates of FS

Methods: Participant recruitment

- 1 May 2013 to 30 June 2014
- Children < 6 years
- Presenting with their *first FS* at PAEDS sites
 - Children's Hospital at Westmead, Sydney
 - Royal Children's Hospital, Melbourne
 - Princess Margaret Hospital, Perth
 - Women's and Children's Hospital, Adelaide
 - Lady Cilento Children's Hospital, Brisbane

Methods: Case definition

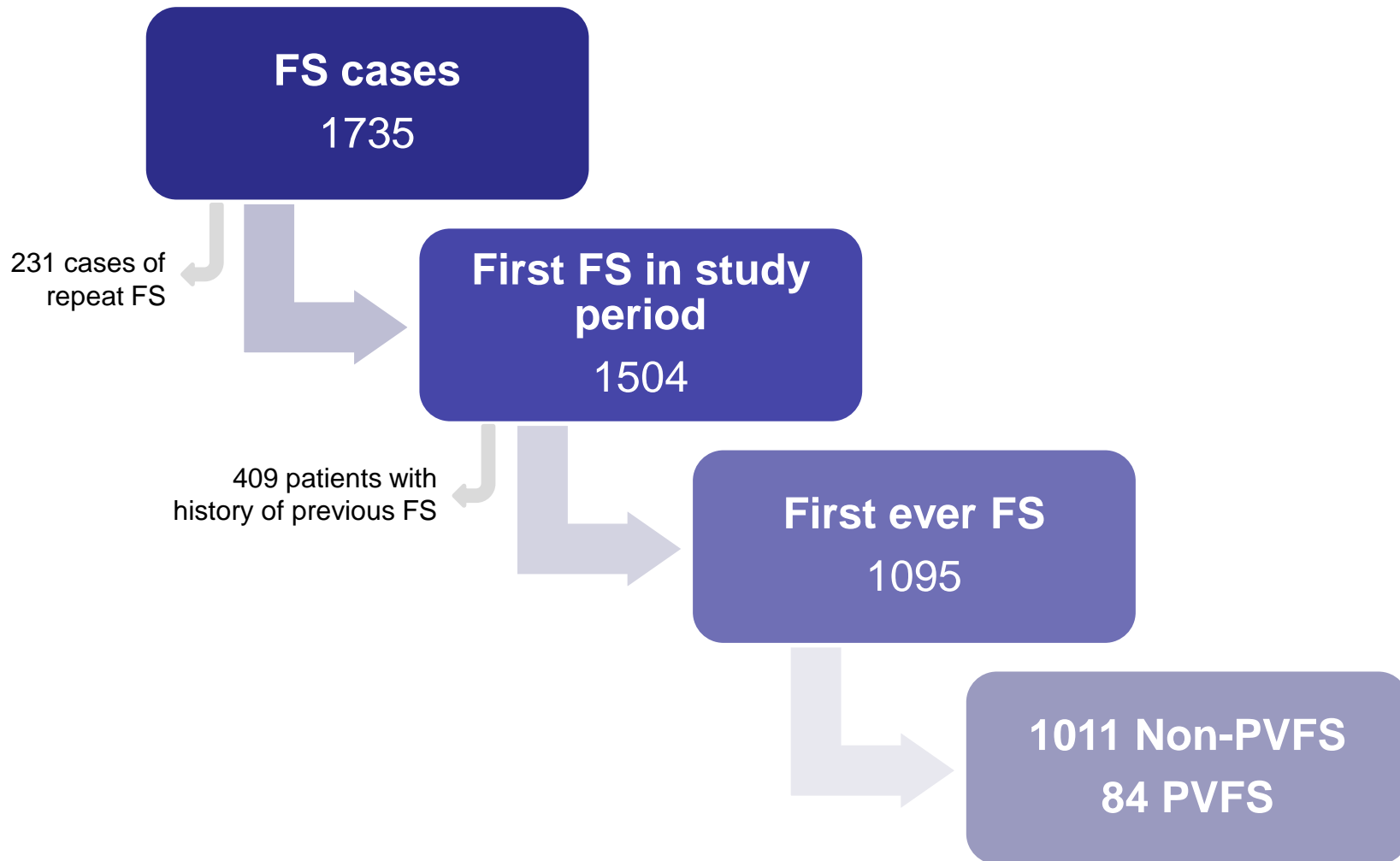
- PVFS
 - Seizure, associated with fever documented either by a parent and/or health provider, occurring within 14 days of any vaccine

- Non-PVFS
 - Febrile seizure outside the abovementioned period following vaccination

Methods: Data collection

- Medical and vaccination history
- Clinical features on presentation/admission
- Investigations
- Management
- Clinical outcome

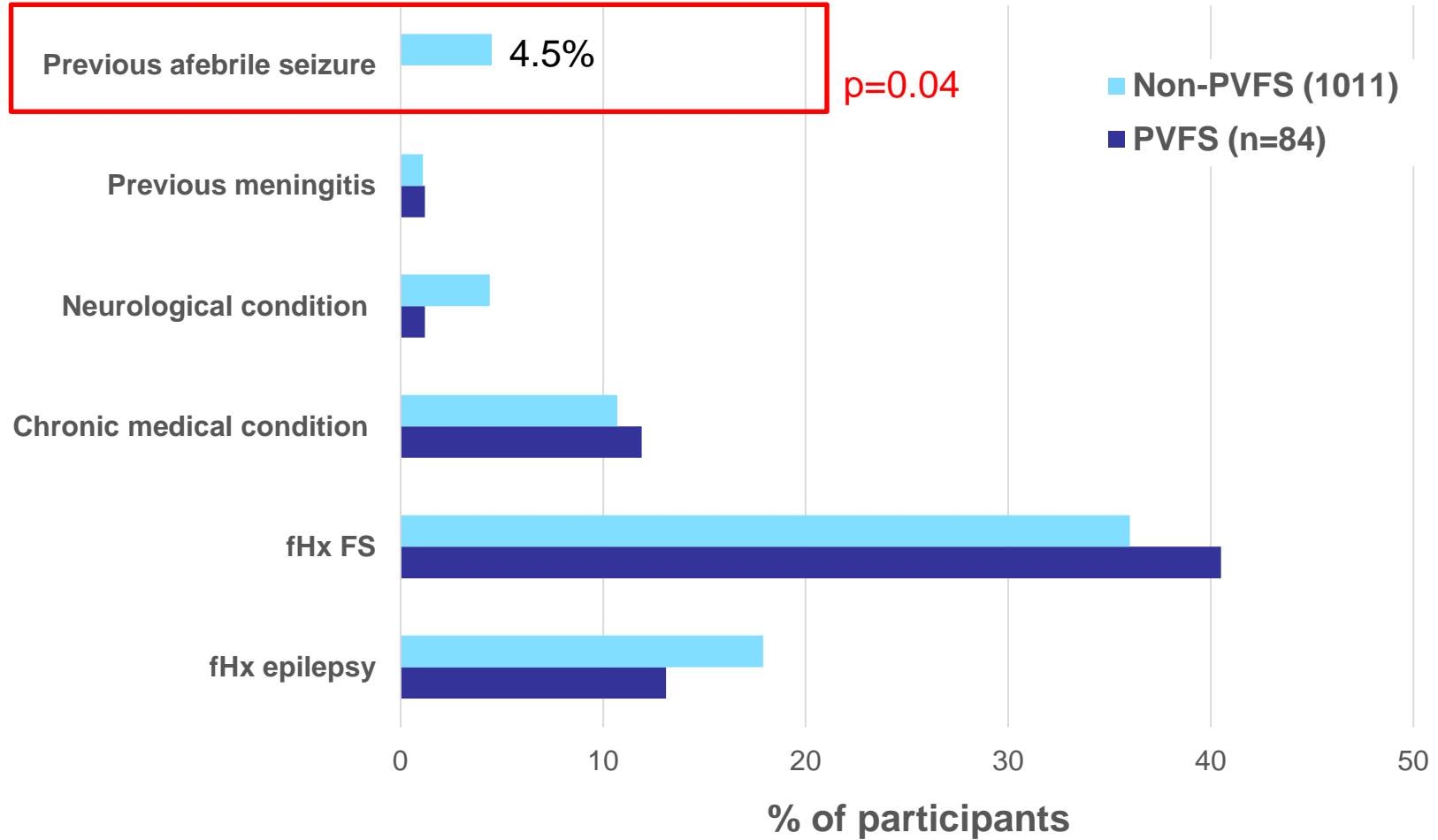
Results: Study population



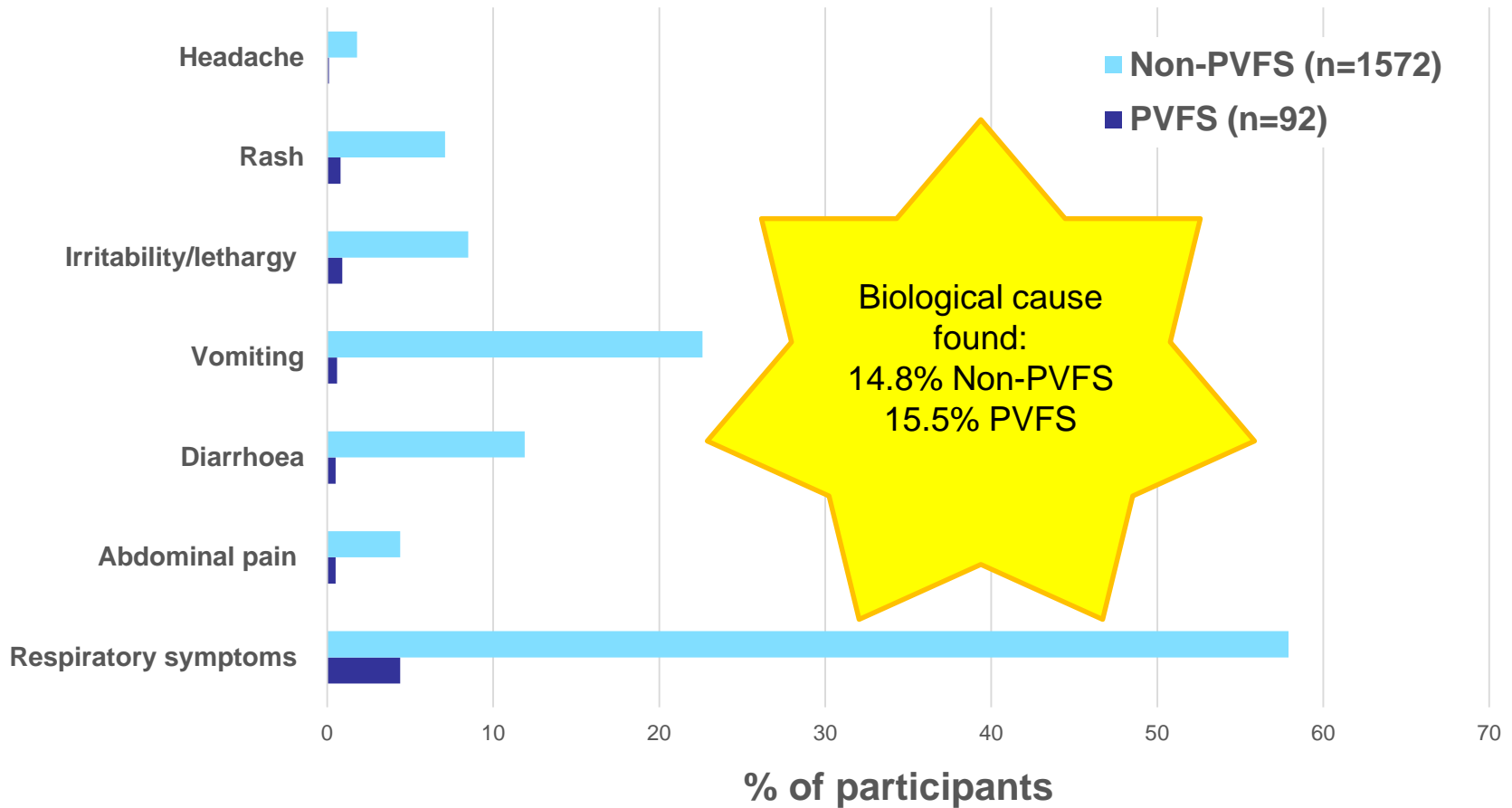
Results: Participant details

| | Non-PVFS (%) | PVFS (%) | |
|--------------|------------------|------------------|--------|
| n | 1011 | 84 | |
| Age (months) | 20.5 (14.2-28.9) | 13.4 (12.4-18.3) | <0.001 |
| Sex (male) | 549 (54.3%) | 38 (45.2%) | 0.11 |
| Indigenous | 29 (2.9%) | 1 (1.2%) | 0.73 |
| Birthweight | | | 0.58 |
| <1500g | 16 (1.6%) | 1 (1.2%) | |
| 1500-2000g | 50 (4.9%) | 3 (3.6%) | |
| 2500-4000g | 709 (70.1%) | 68 (81.0%) | |
| >4000g | 98 (9.7%) | 9 (10.7%) | |
| Gestation | | | 0.44 |
| <28 weeks | 7 (0.7%) | 0 (0.0%) | |
| 28-31 weeks | 5 (0.5%) | 1 (1.2%) | |
| 32-36 weeks | 78 (7.7%) | 4 (4.8%) | |
| >36 weeks | 860 (85.1%) | 78 (92.9%) | |

Results: Participant medical history



Results: Clinical features



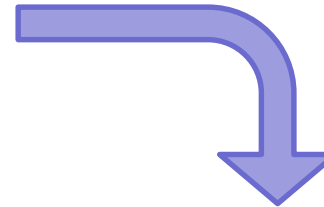
Results: Seizure severity and outcome

| | Non-PVFS (%) | PVFS (%) | Unadjusted OR | p |
|-------------------------------------|--------------|------------|------------------|------|
| n | 1011 | 84 | | |
| Seizure duration > 15 minutes | 264 (26.1%) | 25 (29.8%) | 1.14 (0.70-1.86) | 0.6 |
| Repeat seizure within 24h admission | 105 (10.4%) | 8 (9.5%) | 0.89 (0.41-1.84) | 0.7 |
| Medications | | | | |
| During admission | 92 (9.1%) | 11 (13.1%) | 1.51 (0.77-2.93) | 0.23 |
| On discharge | 44 (4.4%) | 3 (3.6%) | 0.81 (0.25-2.68) | 0.74 |
| Length of stay >1 day | 146 (14.4%) | 16 (19.0%) | 1.40 (0.79-2.47) | 0.25 |
| ICU admission | 23 (2.3%) | 2 (2.4%) | 1.01 (0.26-4.38) | 0.99 |
| Death | 1 (0.1%) | 0 (0.0%) | | |
| Readmission within 48h with FS | 10 (1.0%) | 0 (0.0%) | | |

NO difference in severity or outcome

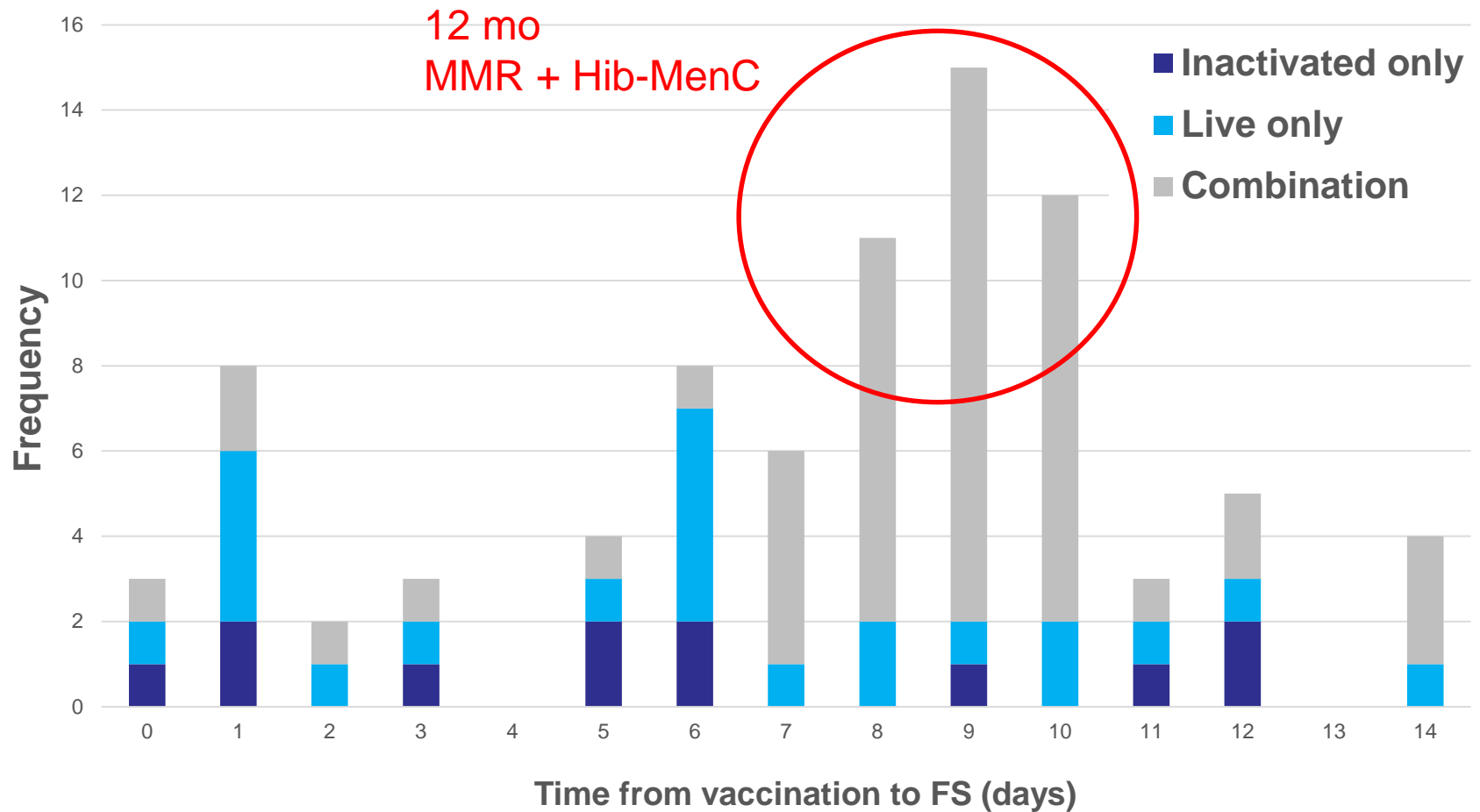
Results: Vaccines involved

| PVFS | n |
|--|------------|
| Total | 84 |
| PVFS following inactivate vaccine only | 12 (14.3%) |
| PVFS following live vaccine | 72 (85.7%) |



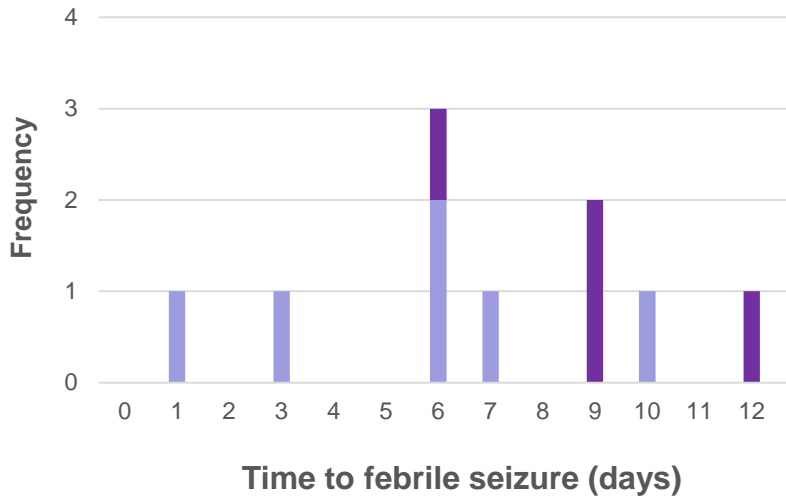
| Live vaccine involved | n |
|-----------------------|------------|
| MMR | 44 (61.1%) |
| MMRV | 18 (25%) |
| Varicella | 2 (2.8%) |
| Rotavirus | 8 (11.1%) |

Results: Timing of PVFS by vaccine type



Results: PVFS by age and biological cause found

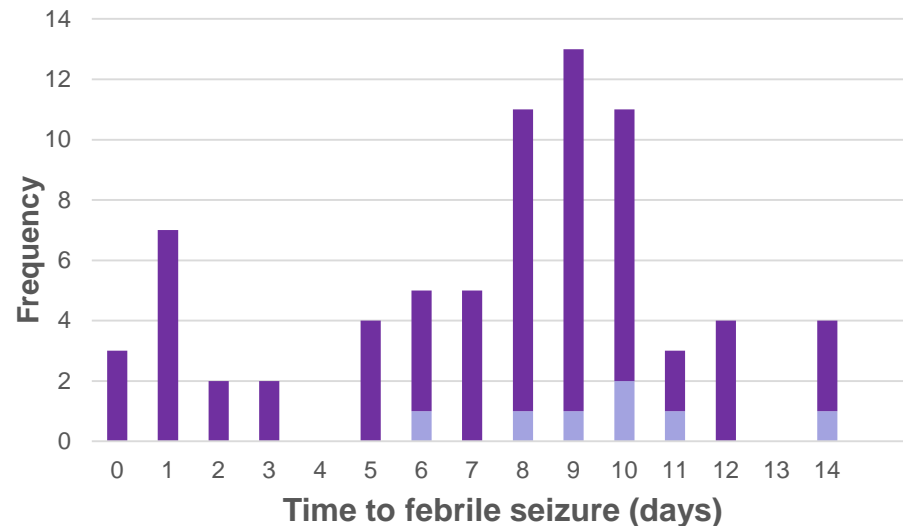
Children aged < 10 months (n=10)



6wks, 4mo, 6mo:
Hep B, DTPa, HiB, IPV, 13vPCV
6 wks, 4mo: + Rotavirus

12mo: MMR + Hib-MenC
18mo: MMR + V / MMRV
4yo: DTPa-IPV

Children > 10 months (n=74)



- Biological cause found
- Biological cause not found

Conclusions

- PVFS account for small proportion of first FS presenting to hospitals in Australia
- No difference in clinical severity or outcomes between PVFS and non-PVFS
- Majority of PVFS are simple FS requiring hospital stay 1 day or less, with no medications required
- Majority of PVFS are associated with a live-attenuated vaccine



Discussion

- Strengths
 - PAEDS network
 - Active surveillance
 - Prospective case ascertainment
 - Detailed clinical data

- Challenges
 - Short study period
 - Recurrence rates difficult to determine
 - Limited to paediatric hospitals only

Future studies

- Recurrence rates of FS following PVFS
- Genetic markers
- Developmental outcomes



PAEDS Paediatric Active Enhanced Disease Surveillance