Adverse events following immunization: Pathophysiology and Communication to

Parents and the Public

Simon Dobson Vaccine Evaluation Centre BC Children's Hospital, Vancouver, BC

Western Canadian Immunization Forum 2011

Where does the infformation come from?

The Parents

- **iPHIS**
- The referring doctor

The referral:

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Mentry on

Hives

- 10 month old
- 2 month immunizations fine
- 4 months: within 10 minutes hives and edema of the lower lip
- Adrenaline given s/c and im
- Rushed to ER

Hives

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- 2 month immunizations fine
- 4 months: within 10 minutes hives and edema of the lower lip
- Adrenaline given s/c and im
- Rushed to ER
- "But doctor, did he come to any harm because his 6 month vaccines were not given on time"

Report from Institute of Medicine 1991

Can it?

Report from Institute of Medicine 1991

Can it?

■ Did it?

Report from Institute of Medicine 1991

Can it?

Did it?

■ Will it?

Can it?

- Strength of association
- Analytic bias
- Biologic gradient
- Consistency
- Biologic plausability

Vaccine Adverse Events: Did it?

- Previous general experience
- Is there another etiology
- It happened before to this recipient
- Timing

After this:

- 4 year old boy
- Pre-school visit
- 2 hours later: widespread hives
- Soon after wheezing, itchy eyes
- Goes to the ER

Will it?

Rechallenge

After this:

- 4 year old boy
- Pre-school visit
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■ It is all in the story......

Mechanisms of Adverse Events: 1) The immunization process

Poor injection technique

- Vaso-vagal
 - **■** faints
 - **■** Funny turns
 - Hypotonic/Hyporesponsive Episode (HHE)

Mechanisms of Adverse Events: 1) The immunization process

Dermographism



Long term consequences:

- 15 year old girl
- At 2 months old had received a single immunization
- 2 hours later, found limp, grey and unresponsive
- Paramedics ++
- ER: "likely reaction to pertussis component"
- MHO: "Hypotonic/Hyporesponsive Episode", "will never recur"

Long term consequences 15 year old girl continued:

- It did!
- At 4 months:
- 2 hours later; eyes rolled back and went limp
- Pediatrician; "we are not doing that again"
- No immunizations since

Mechanisms of Adverse Events: 2) Direct effect of vaccine components

Viral replication

Local suppuration and adenitis (BCG)

Mechanisms of Adverse Events: 3) Host immune response

- Local inflammation results from the innate immune response
 - Toll-like receptors (TLRs)
 - Macrophages and dendritic cells move in
 - Results in redness and swelling

Mechanisms of Adverse Events: 3) Host immune response

- **■** The Arthus reaction
 - Antigen deposit is met with lots of antibody
 - Redness, swelling AND Pain

Mechanisms of Adverse Events: 3) Host immune response

- Systemic inflammatory response
 - **■** Fever
 - Irritability
 - Nausea
 - Vomiting
 - **■** Muscle aches

Mechanisms of Adverse Events: 3) Host immune response

- Systemic inflammatory response host factors:
 - Age younger less likely
 - Gender girls worse than boys
 - Genetics ??
 - **■** Previous infection
 - **■** Previous doses of vaccine

Mechanisms of Adverse Events: 3) Host immune response; Allergic reactions

- Type I hypersensitivity reactions
 - Urticaria
 - Angio-edema
 - anaphylaxis

Mechanisms of Adverse Events: 3) Host immune response; Allergic reactions

- Type IV hypersensitivity reactions
 - 72-96 hours after immunization
 - Especially thiomersal containing vaccines

Mechanisms of Adverse Events: 3) Host immune response

- "Other"
 - Thrombocytopenia
 - **■** Autoimmune
 - **■** Guillain-Barré syndrome
 - Etc, etc, etc.

The future:

- CISA (and similar)
- Clinical Immunization Safety Assessment Network
- Goals:
 - Clinical evaluation, diagnosis and management of AEFIs
 - Understand AEFIs at the individual level
 - Evidence based algorithms for the next shot
 - **■** Experts for inquiries

Summary

- It is all in the story
- Parents remember
- There are individual and family consequences to what you say
- Give a clear account of why you think what happened happened
- If you don't know say so