The Immune System and Immunisation



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The Immune System



ALBERT EINSTEIN

"Make things as simple as possible (but not simpler)."

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Outline of Talk



- An overview of the immune system
- The journey of a pathogen and the obstacles it meets
- How vaccines work and how soon we are protected
- Types of vaccine live vs inactivated
- Vaccine components
- Timing of vaccinations, timing of adverse reactions
- Herd immunity

What is the immune system?

 The body's defense against disease causing organisms, malfunctioning cells, and foreign particles







Pathogens

- Pathogens: disease causing agents- such asbacteria, virus, and fungi
- Patho- sickness agent
- Gen- to create



The Immune System – Defence Against Pathogens

• In other words, how to stop David from killing Goliath...



The Immune System

- Array of organs, cells and chemicals that:
- Determine self from "non-self"
- Identify potential dangers to the body
- Eliminate them by mounting an immune response



The Infant's Immune system...

- Naive needs exposure to foreign antigen in order to develop normally
- Maternally acquired immunity is temporary and does not protect against all infections.
- The infant immune system has the capacity to cope with a vast array of antigens at any one time.



Immune System Components



Source: <u>http://www.webmd.com/a-to-z-guides/components-of-the-immune-system</u>

Organisation of the Immune System



Immunity: Active and Passive





The Pathogen's Journey





Will Sickness Occur?

Getting sick or remaining well when exposed to a germ is a balance. It depends on the virulence of the germ and the ability of the person to resist the infection.

Both of these things are variable.

www.nrvs.info

The Consequences of Infection...

- Lifelong immunity (most of the time!)
- May be innocuous

BUT....

- May cause serious disease
- May cause permanent damage to the host
- May cause death



The Pathogen soon encounters the first level of defence

- Physical barriers (intact skin, intact mucous membranes, cilia etc....)
- Physiological factors (eg pH, temp)
- Protein secretions (complement, interferons)
- Phagocytes macrophages and PMNLs

Defining characteristic of innate immunity -**NO MEMORY PERSISTS.**



Macrophages – part of the first level of defence



Macrophage surrounds/"eats" the virus

- Digest most of the micro-organism
- Regurgitate the antigens
- Display antigens on their surface so that another type of white blood cell (lymphocytes) can take over.

What is an antigen?



Microbe

Fragments of Microbe = antigen

"Anything that can be bound by an antibody"

The Pathogen Next Encounters the Second Level of Defence

- Adaptive immunity
- The foreign agent is recognised in a specific manner e.g
- B Cells
- T Cells

THE IMMUNE SYSTEM ACQUIRES MEMORY



Cell-Mediated Immune Response



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What is an Antibody?



- Produced to one specific epitope (ie is antigen specific)
- Neutralises toxins
- Blocks adhesion/ cell entry
- Kills via complement
- Neutralises viral infectivity and prevents replication.

Immune System Superheroes!



Memory Cells

- Once infection has been eliminated, some B and T cells become memory cells
- These retain memory of the antigen
- On re-exposure, powerful immune response.
- This ability of the immune system to have a memory for previous antigens is the basis for vaccination.



How Soon After Exposure to an Antigen are we Protected?

- Immune response is generated after 4-7/7
- >7/7 get Primary immune response (IgM), lasts 3 weeks, memory cells made.
- Secondary/subsequent immune response, IgG, faster
- It takes 2 weeks to get optimum immune response after vaccination.



In Simple Terms....

Vaccines work by making us produce antibodies to fight disease without actually infecting us with the disease. If the vaccinated person then comes into contact with the disease itself, their immune system will recognise it and immediately produce the antibodies they need to fight it.



The Ideal Vaccine

- Produces the same immune protection as an infection without causing disease
- Generates long-lasting immunity
- Interrupts spread of infection



Vaccines can be broadly divided into two types

• Live attenuated

Inactivated









Basic Differences

Live Attenuated –"Weak Pathogen"

- A version of the living microbe that has been weakened in the lab so it can't cause disease.
- Vaccines are longer lasting and require fewer boosters
- However the disease could mutate back to the pathogenic strain
- eg BCG/ MMR/ Rotavirus/ Varicella/ Yellow fever

Inactive – "Dead Pathogen"

- Produced by killing the disease-causing microbe with chemicals, heat, or radiation.
- Cannot cause disease
- Cannot replicate
- Immune response mainly antibody based
- Antibody titre falls with time
- 3-5 doses required
- Classified as inactivated/conjugate/recombi nant/subunit

Inactivated Vaccines

Whole

- viruses
- bacteria

Fractional

- protein-based
 - toxoid
 - subunit
- polysaccharide-based
 - pure
 - conjugate

Live Attenuated Vaccines

 Produced by weakening a live pathogen and removing its ability to cause disease

PROS		CONS	
•	Large immune response Good "teachers" of the immune system Generally only 1-2 doses needed	•	Need strict refrigeration Could mutate back to disease-causing strain

Inactivated Vaccines

Produced by killing the pathogen

PROS	CONS	
 They may not have to be stored as carefully. They will never come back to life and cause the disease. 	• They usually require booster shots because they only weakly stimulate the immune system to make antibodies.	

Toxoid Vaccines

• Produced by inactivating the toxin produced by some pathogens eg tetanus and diphtheria

PROS	CONS	
• Generally only need one or two shots	Require refrigeration	
• They will never come back to life and cause the disease.		

Subunit Vaccines

• Produced by extracting the antigenic part of a micro-organism. Eg hep B and strep pneumo

PROS		CONS	
•	They will never come back to life and cause the disease.	•	They are more difficult to make and require new, expensive technology.

How Vaccines Are Made





Microbes grown on suitable medium



Purified to remove compounds that could cause allergic reactions

(not always possible)

Vaccine Components

- Suspension fluid (water, saline etc)
- Preservatives, stabilisers, antimicrobial agents
- a) Trace amounts
- b) May cause allergic reaction
- Adjuvants
- a) Aluminium salts to increase immunogenicity
- b) Eg hep B, tet, diphth



True or False?

Several childhood vaccines contain mercury, which is toxic to the nervous system.



Thiomersal

- Mercury containing compound used as a preservative and an inactivating agent
- In 1999 EU and US manufacturers decided to decrease thiomersal levels in vaccines as a precaution and to retain trust in vaccine supply
- WHO state that there is no evidence of toxicity
- All vaccines in the infant immunisation programme are thiomersal free

Timing of Vaccinations



Why are Gaps Needed Between Doses?

- To allow each immune response to develop eg primary immunisation
- To avoid immune interference if another live vaccine is given while the immune system is making a primary immune response, the activation of the innate immune system may neutralise the second vaccine. Hence we wait 4 weeks.

Timing of Primary Immunisation Course

- Maternal IgG is transferred across the placenta
- Passively acquired IgG can suppress response to DTP, Polio, Men C and Hib for 2 months
- Maternal antibody to measles may interfere for 1 year.

True or False?

Vaccinations can "overload" the immune system



Can Vaccines Overload the Immune System?

- We are exposed to countless antigens every day, in the food we eat, in the air we breathe, in the water we drink.
- The human body contains 100 trillion bacteria.
- The immune system is capable of responding to 100,000,000,000 antigens at a given time
- The MMR contains only 24 antigens.

NO EVIDENCE THAT VACCINES OVERLOAD THE IMMUNE SYSTEM

Arguing that vaccines will overwhelm a child's immune system



is like arguing that a tablespoon will make an Olympic swimming pool overflow thelogicofscience.com

Vaccine Failures and Reactions



When it's a joke for everyone, except the guy next in line.

Vaccine failures

- Primary failure when an individual fails to respond to the initial vaccine (eg 10% MMR)
- Secondary failure responds initially but response wanes over time (most inactivated vaccines)



Timing of Vaccine Reactions

- Inactivated generally within 48h
- Live vaccine according to time taken for virus to replicate
- Eg MMR vaccine
- a) Reactions to measles (malaise, fever, rash) occur in 1st week
- b) Rubella (pain, joint swelling) in 2nd week
- c) Mumps (parotid swelling) in 3rd week

Adverse events

- Live vaccine frequency of adverse events falls with number of doses
- Inactivated vaccines frequency of adverse events increases with number of doses...

if antibody levels are good, this binds to antigen in subseq dose, producing an Immune response which, if big enough, is inflammatory



HERD IMMUNITY

- When most people in community are immune to a particular infection that is spread from *person to person*, the natural transmission of the infection is effectively inhibited
- Vaccine uptake rates >90% (measles 95%)
- Not tetanus!



Herd Immunity

HERD IMMUNITY

- If enough of the population is immunized, even those that aren't are protected
- Who relies on herd immunity?
 - Infants
 - Elderly
 - Those with weakened immune systems
 - Those who are allergic to the vaccine



http://www.vaccines.gov/basics/protection/

