Immune effector mechanisms

Intracellular pathogens

## T CD4 – helper - differentiation



Dynamics

Response profile – focus on CD4 Th1

Effector mechanisms- CD8, CD4 Th1, humoral



## **TH1 RESPONSE**



lgG

## **INDUCTION OF IL-12 EXPRESSION**



# Th1 differentiation



Pathogen

#### EFFECTOR MECHANISMS – CELLULAR RESPONSES



TNFR family and cell death induction



## Perforin/Granzyme B



Contents of the dense core	Specialized function
Perforin	Pore formation; disruption of endosomal trafficking
Granulysin	Microbicidal agent
Calreticulin	Calcium and perforin binding
Granzymes	A to M
Dipeptidylpeptidase I (cathepsin	
C)	Activation of granzymes
Chondroitin sulfate proteoglycans	Complexes to granzymes

Resident lysosomal proteins (periphery of granule)

Mannose 6-phosphate receptor	Protein trafficking
H+-ATPase	Granule acidification
Cathepsin D	Protease
Cathepsin L	Protease
Arylsulfatase	
β-Hexosamidase	
β-Glucuronidase	
CD63	
Lamp 1	
Lamp 2	

#### CITOTOXIC IMMUNE SYNAPSE



#### CD4 Th1 CELLS ACTIVATE MACROPHAGES



## IFN $\gamma$ EFFECTS ON MACROPHAGES



#### GRANULOMAS – RESULT OF CD4 Th1 response against Tb



Nature Reviews | Immunology

#### DELAYED TYPE HYPERSENSITIVY RESPONSE



Cancer Immunol Immunother. 2012 Sep; 61(9): 1485–1492.

## HUMORAL RESPONSES – IgG Th1 – IFN $\gamma$ – isotype switch to IgG

Antibody dependent citotoxicity

Antibody dependent phagocytosis

Complement activation

Neutralization

ADCC

Signaling events upon antibody-dependent cellular cytotoxicity



#### ACTIVATION OF THE COMPLEMENT SYSTEM





#### NEUTRALIZATION



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