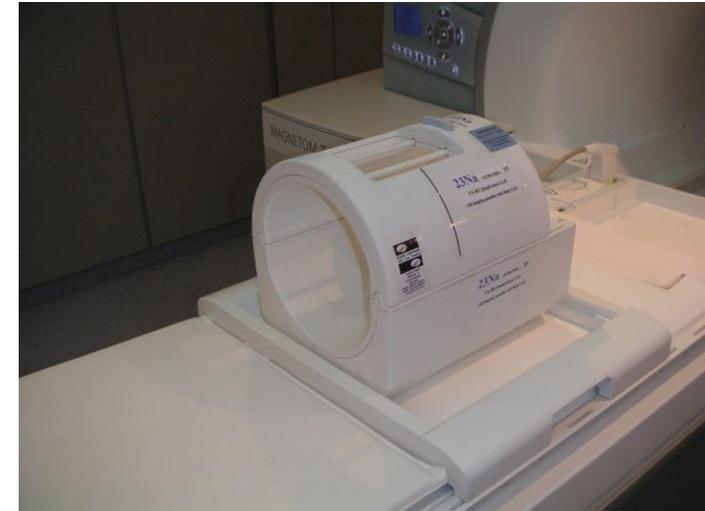
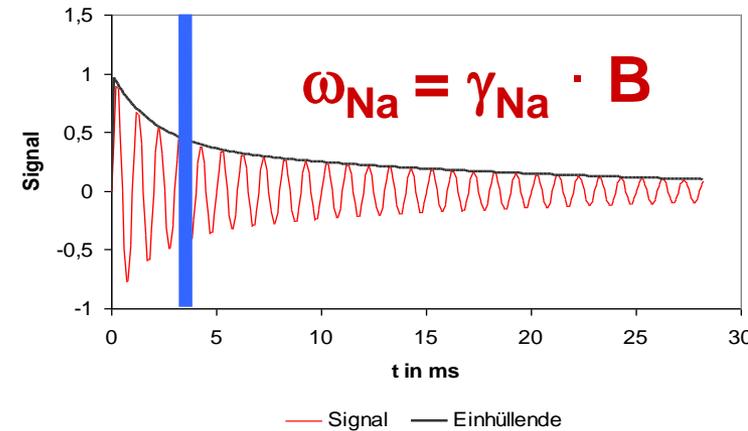


# Na-MRI can help us find the salt



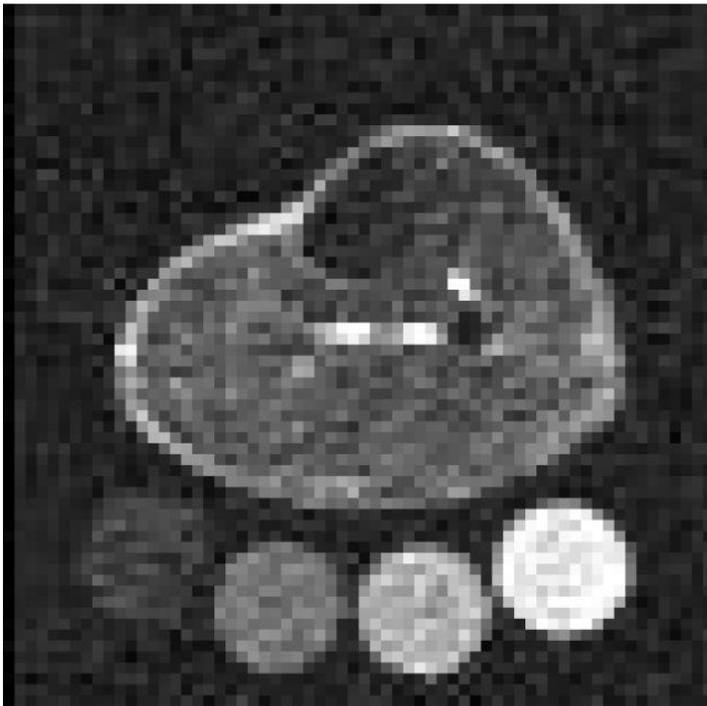
TE = 2...3 ms



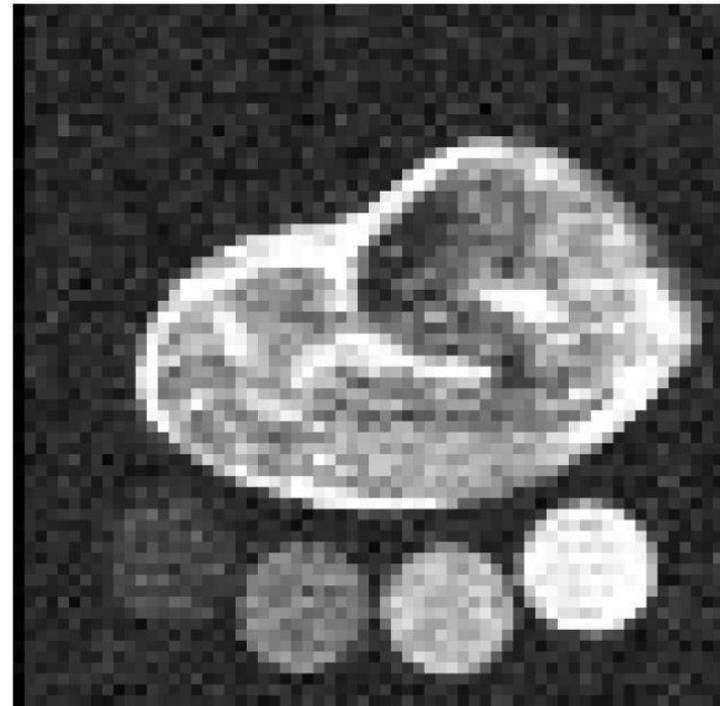
# Sodium research – quo vadis?

## $^{23}\text{Na}$ MRI of tissue $\text{Na}^+$ content

Man, 24 y, healthy



Man, 85 y, hypertension



# Na<sup>+</sup> storage and cardiovascular outcome

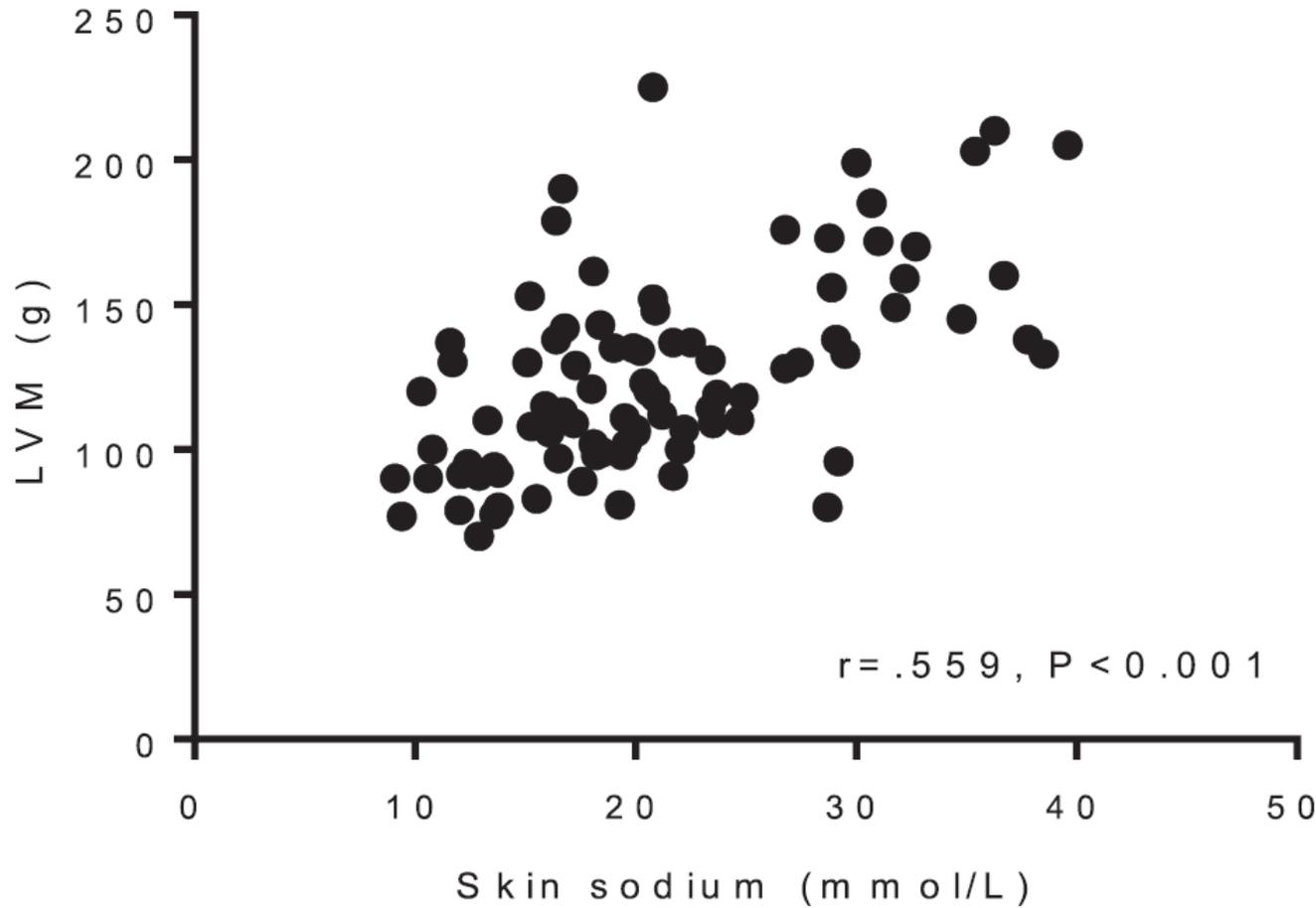
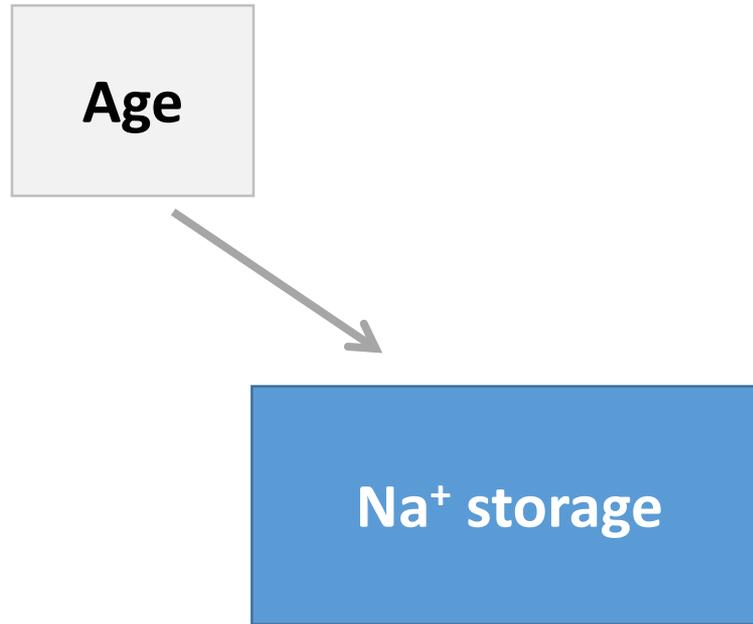


Figure 4. Relationship between skin sodium content and LVM.



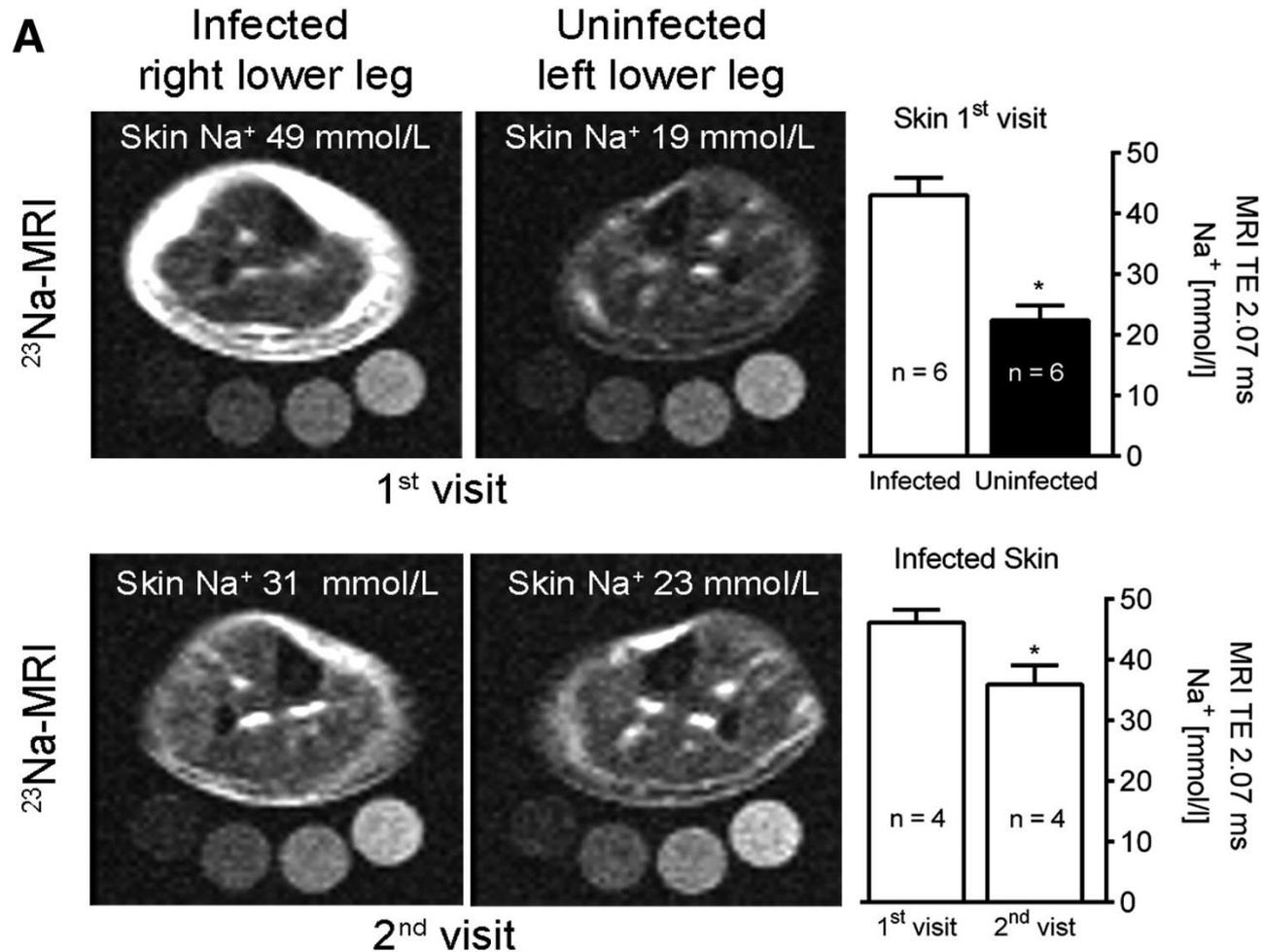
Why would we kill ourselves?

ENCEPHALOMYELITIS

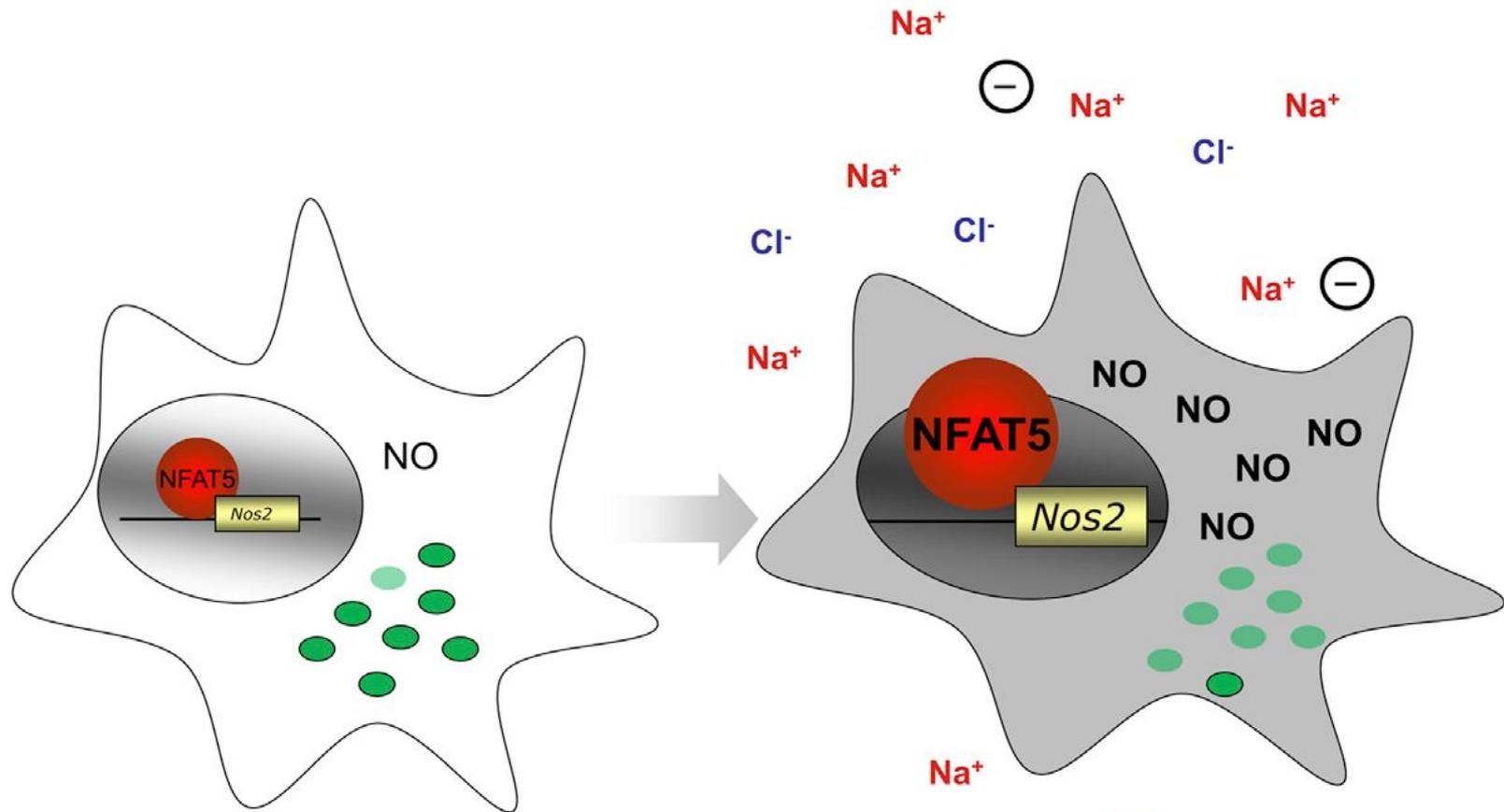
HYPERTENSION

CVD

# Salt barrier formation to ward off infections



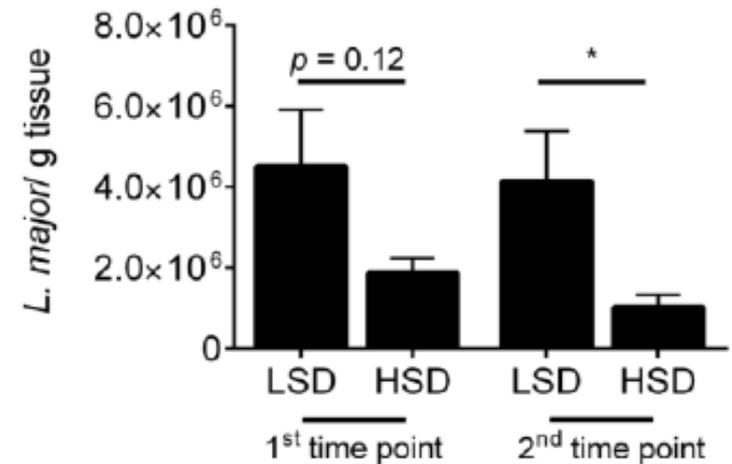
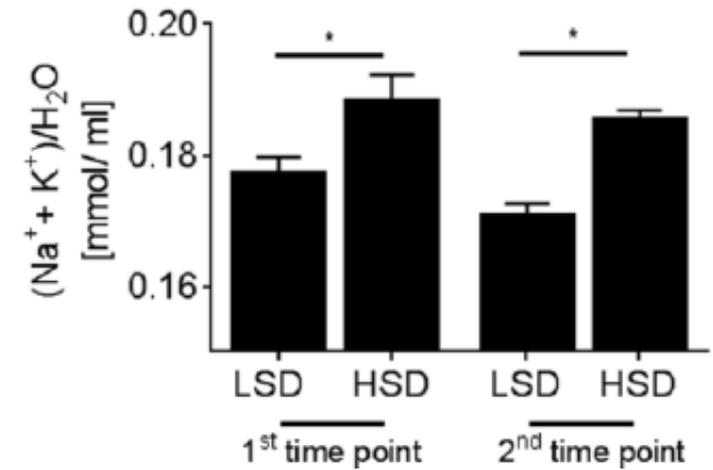
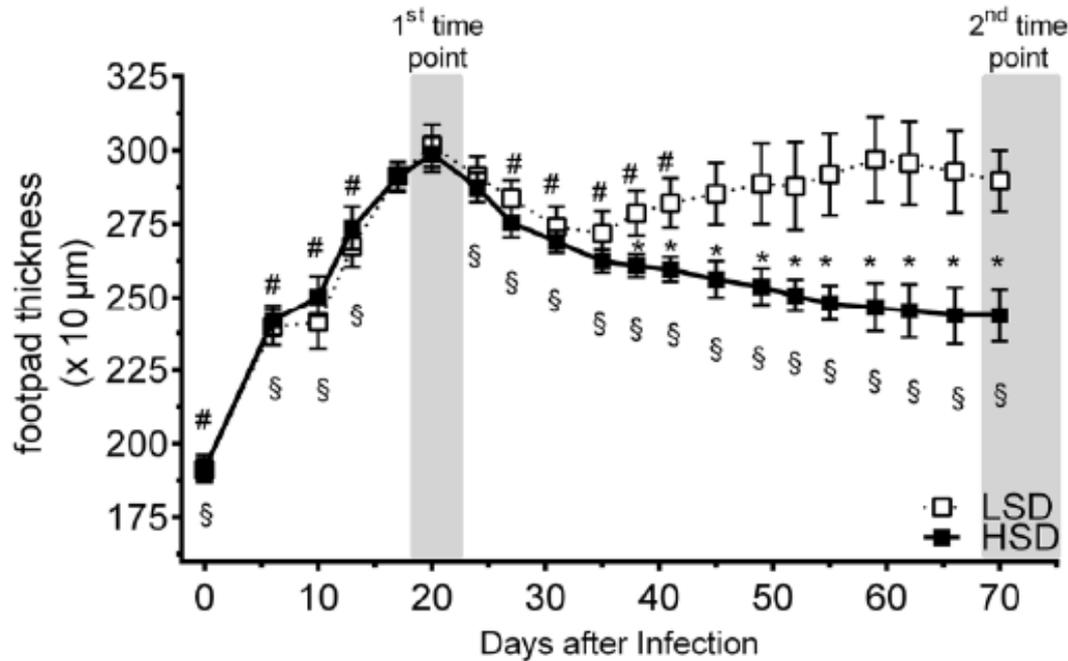
# Salt barrier formation to ward off infections

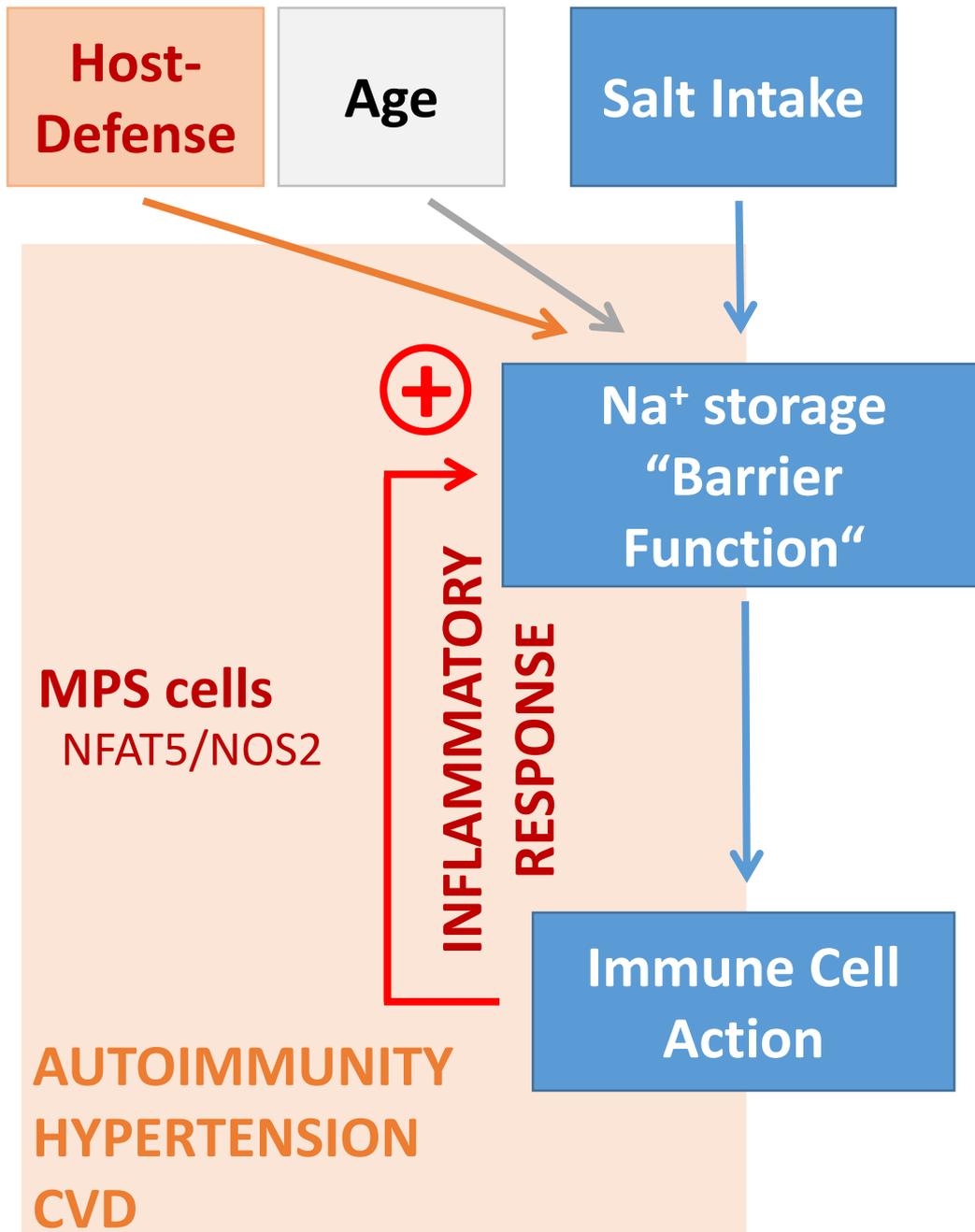


● *Leishmania major*

⊖ Glycosaminoglycans

# Salt barrier formation to ward off infections





# Autoimmunity – a mislead host defense.

## LETTER

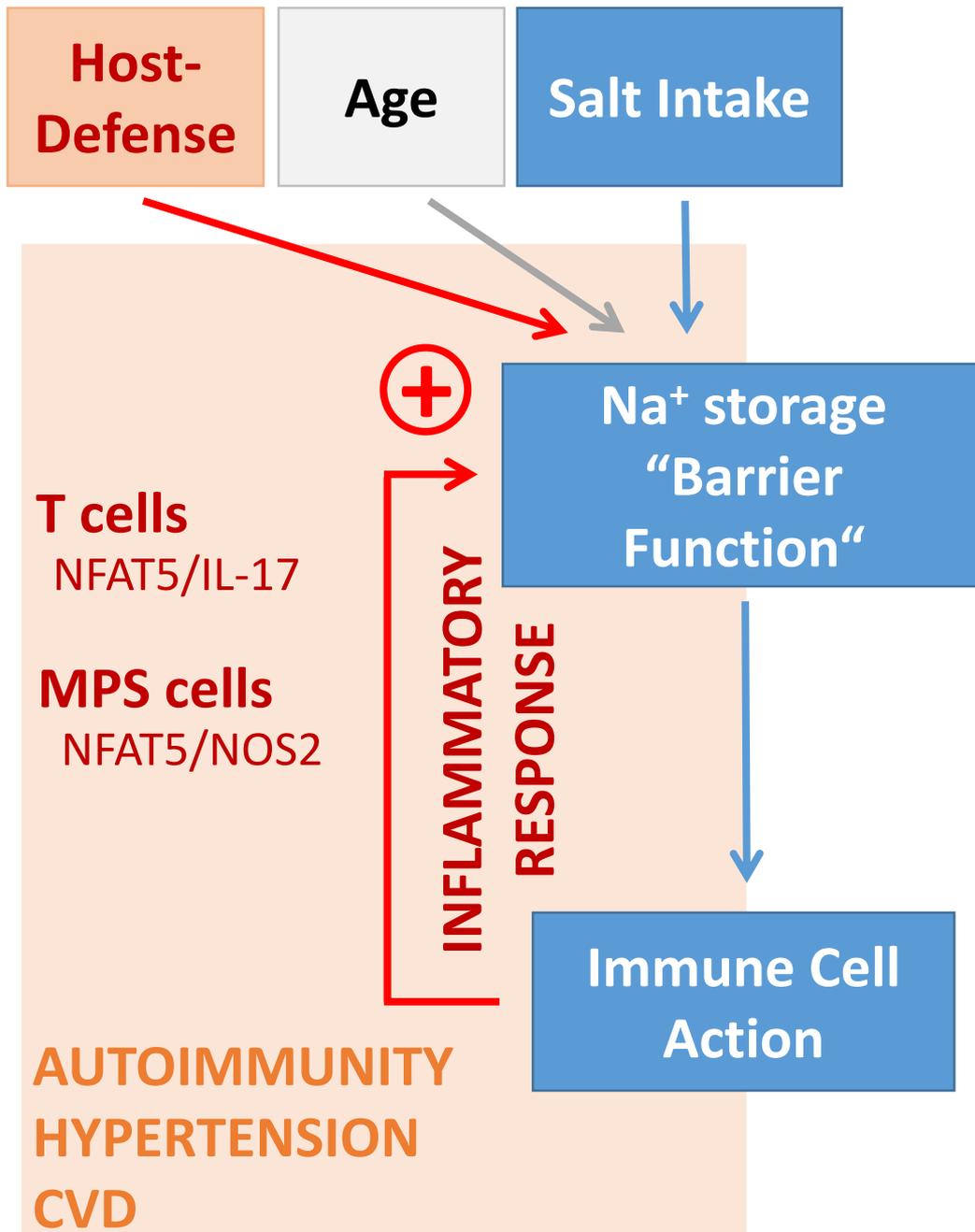
doi:10.1038/nature11868

---

---

### **Sodium chloride drives autoimmune disease by the induction of pathogenic T<sub>H</sub>17 cells**

Markus Kleinewietfeld<sup>1,2</sup>, Arndt Manzel<sup>3,4</sup>, Jens Titze<sup>5,6</sup>, Heda Kvakan<sup>7,8</sup>, Nir Yosef<sup>2</sup>, Ralf A. Linker<sup>3</sup>, Dominik N. Müller<sup>7,9\*</sup> & David A. Hafler<sup>1,2\*</sup>



# **Local control of salt metabolism by homeostatic immune cells**

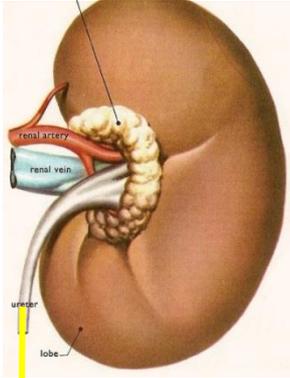
# What I have learned at school

Skin

Equilibrium

Blood

Kidney



Clearance

$\text{Na}^+$

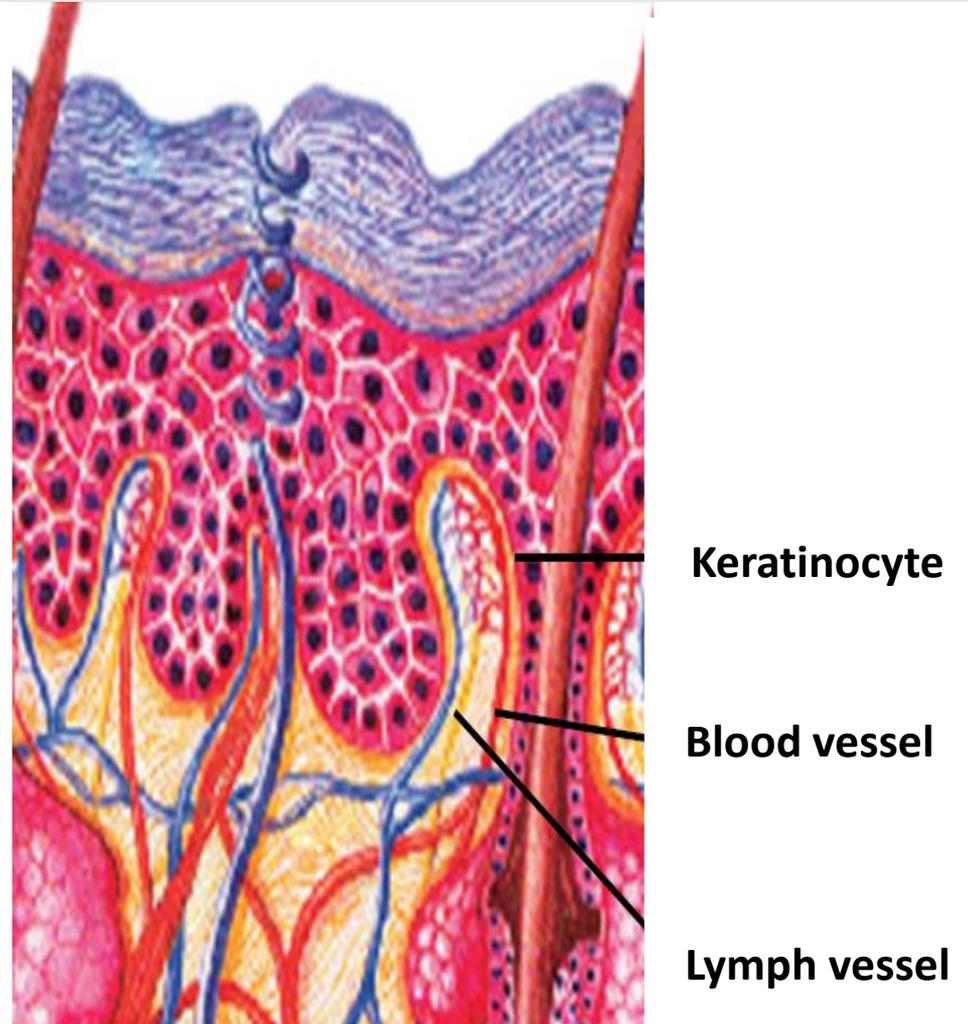
$\text{Cl}^-$

Urine

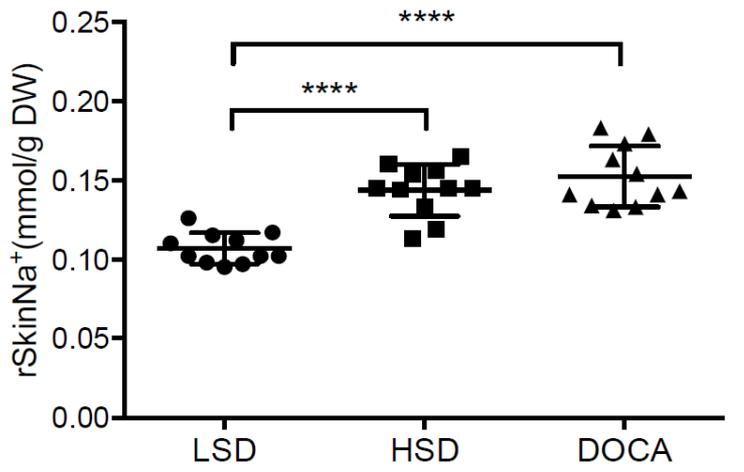
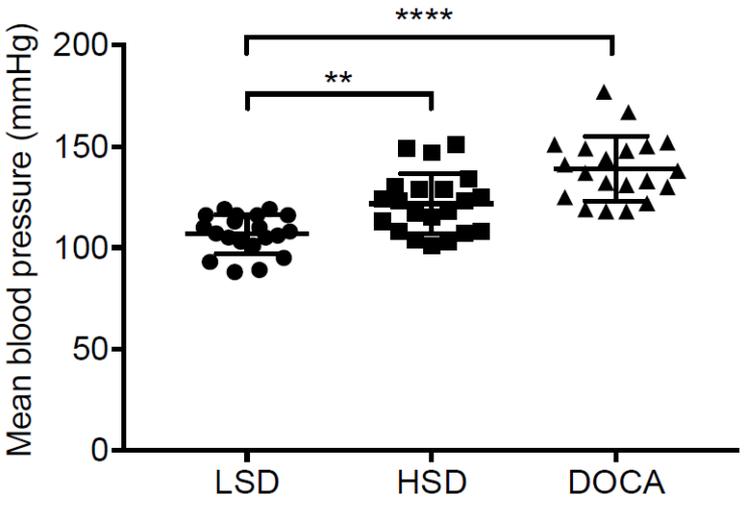
# Skin sodium storage in guinea pigs and rats

Stratum Corneum	30 mmol/kg
Stratum Granulosum	160 mmol/kg
Stratum Spinosum	374 mmol/kg
Stratum Germinativum	313 mmol/kg
Dermis	530 mmol/kg

J Investig Dermatol 79(3):167–169, 1982

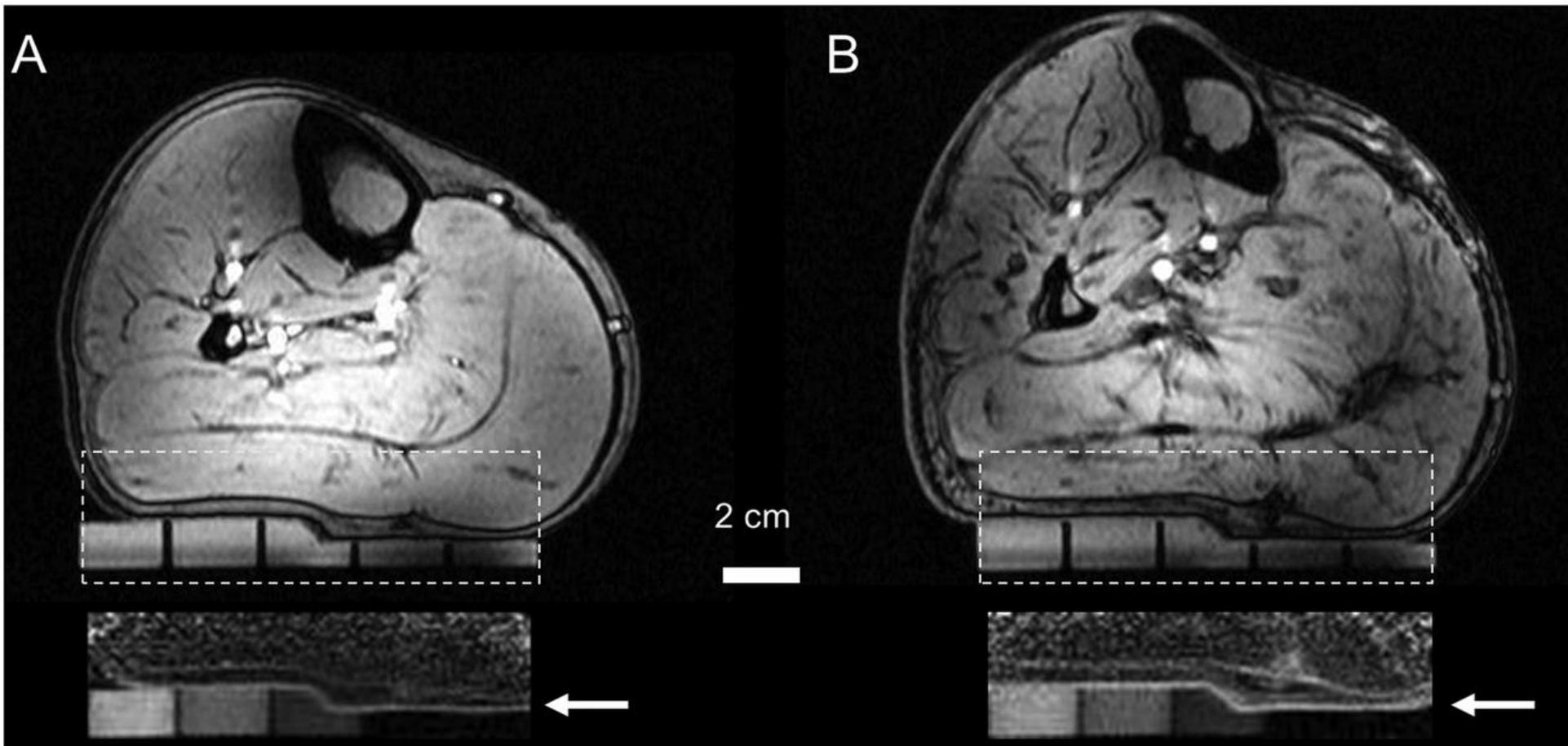


# Skin sodium storage and hypertension in rats



Keratinocyte  
Blood vessel  
Lymph vessel

# 7T Na-MRI can help us find the salt much better



# The problem with what I have learned at school

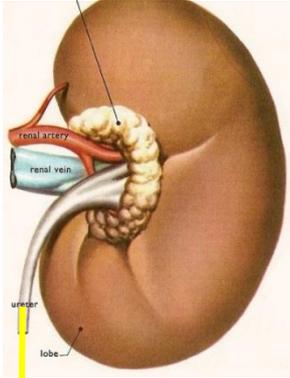
Skin

Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup>  
Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup> Na<sup>+</sup>

Equilibrium

Blood

Kidney

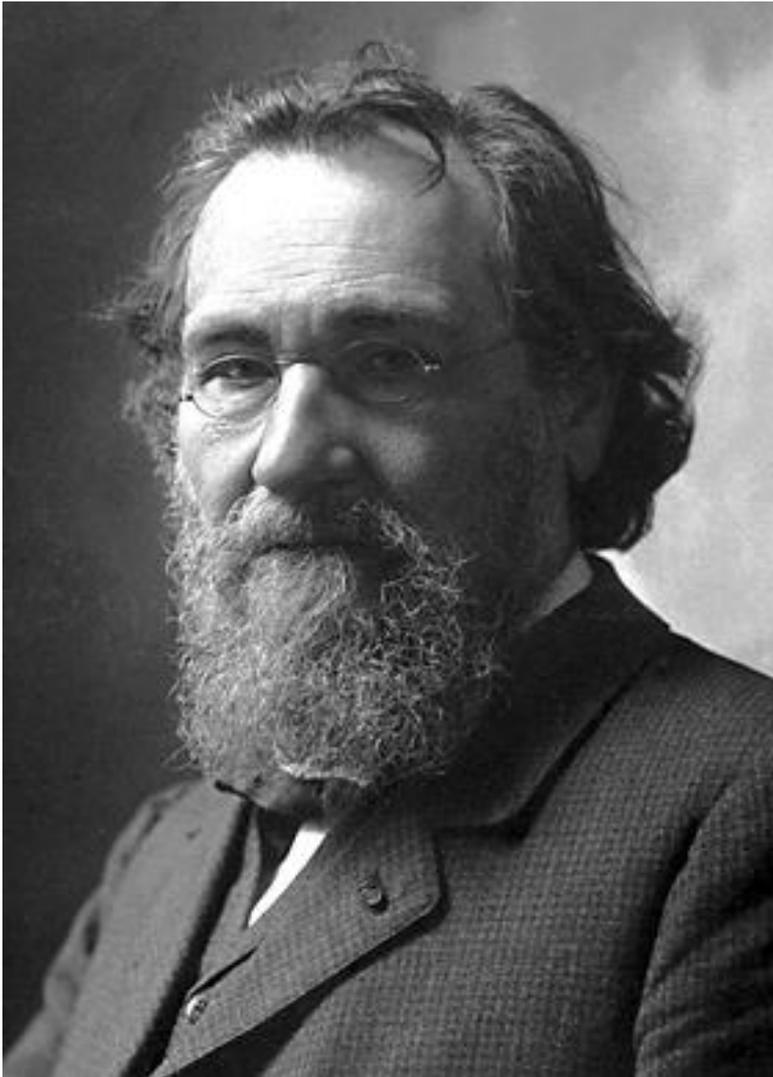


Clearance

Na<sup>+</sup>  
Cl<sup>-</sup>

Urine

# What is immune function?



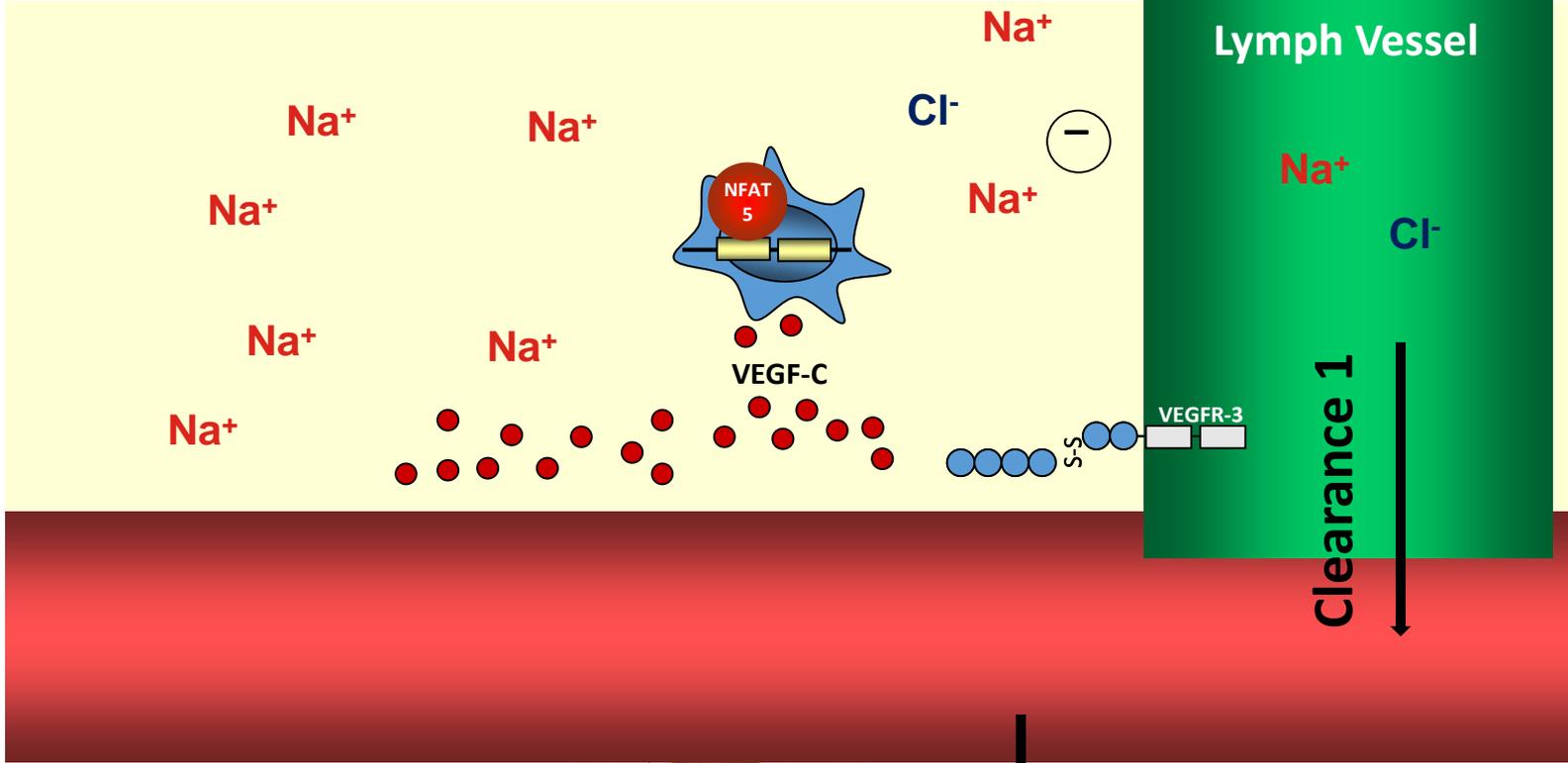
## **Physiological inflammation:**

Phagocytes are engaged in essentially the same process — clearing the body of dysfunctional elements (endogenous ‘other’) and unwanted external intruders.

Tauber AI.

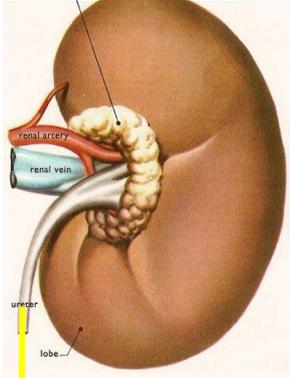
Nat Rev Mol Cell Biol. 2003 Nov;4(11):897-901

**Skin**  
**Interstitial**



**Blood**

**Kidney**

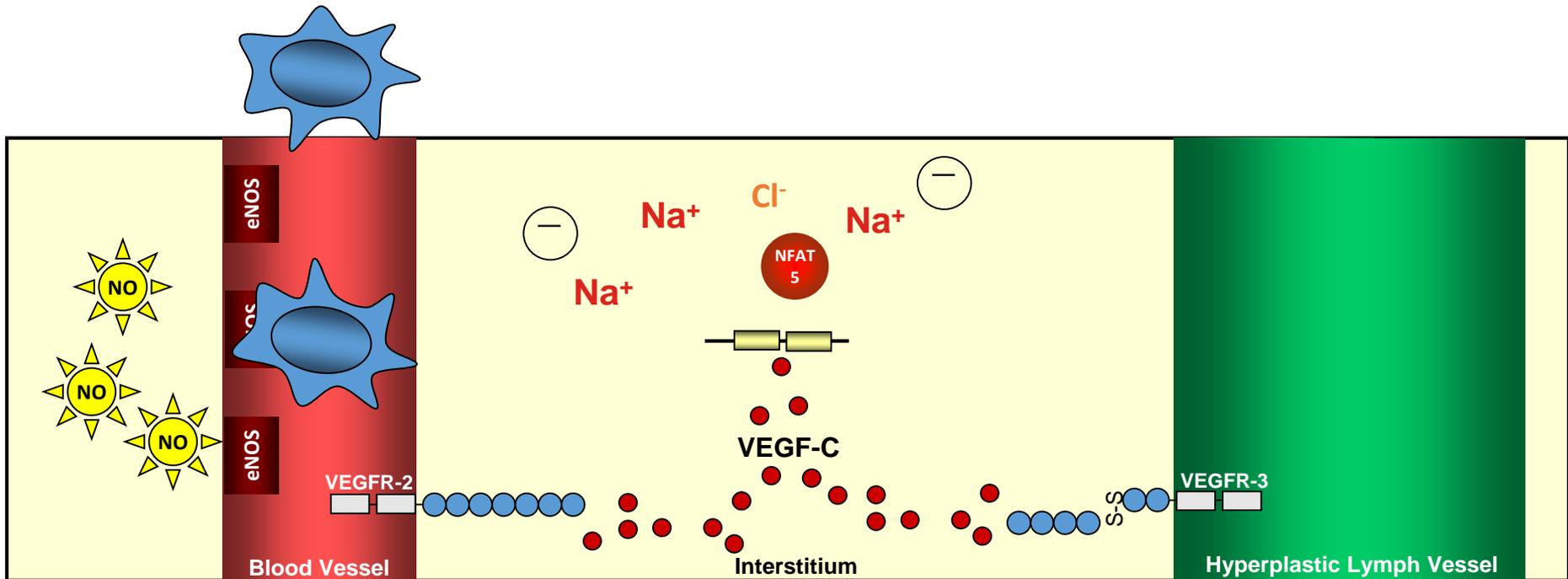
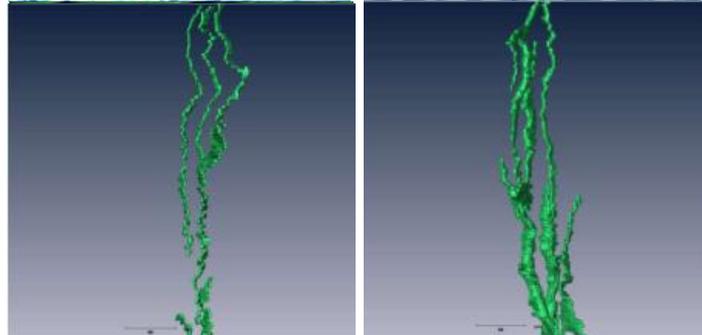


**Clearance 2**

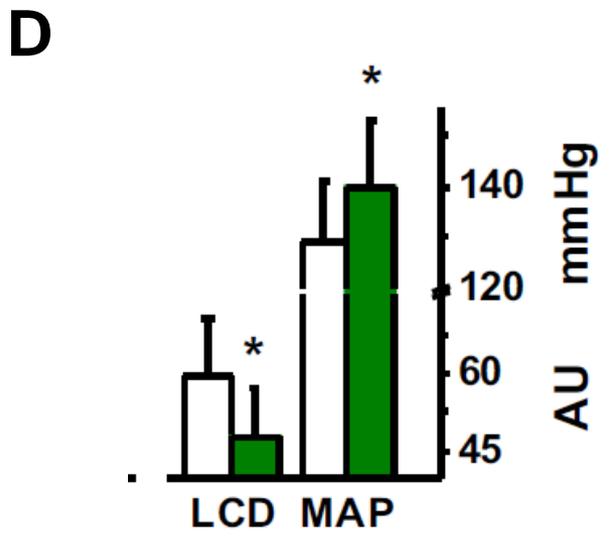
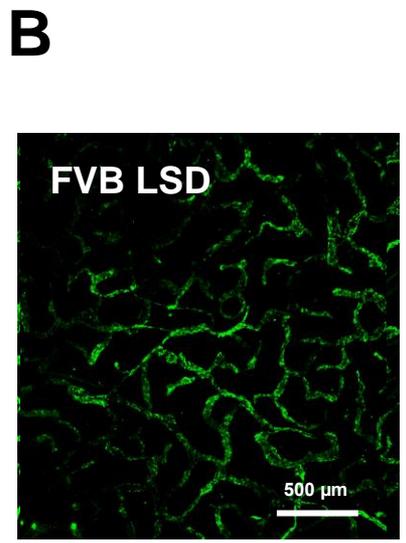
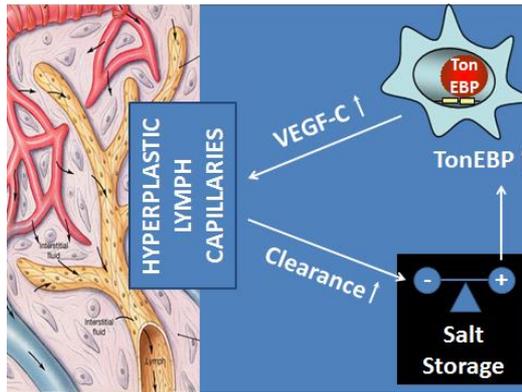
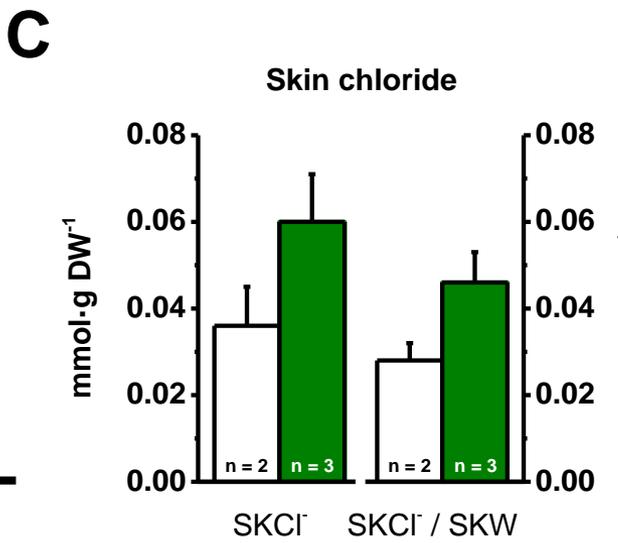
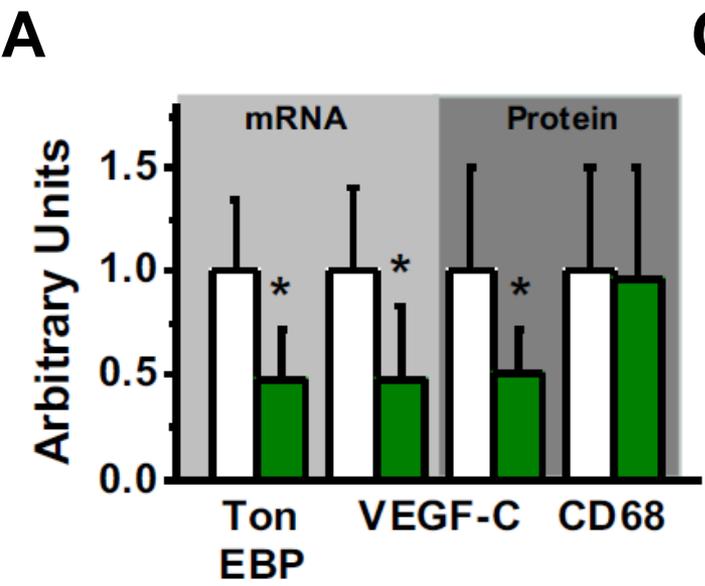


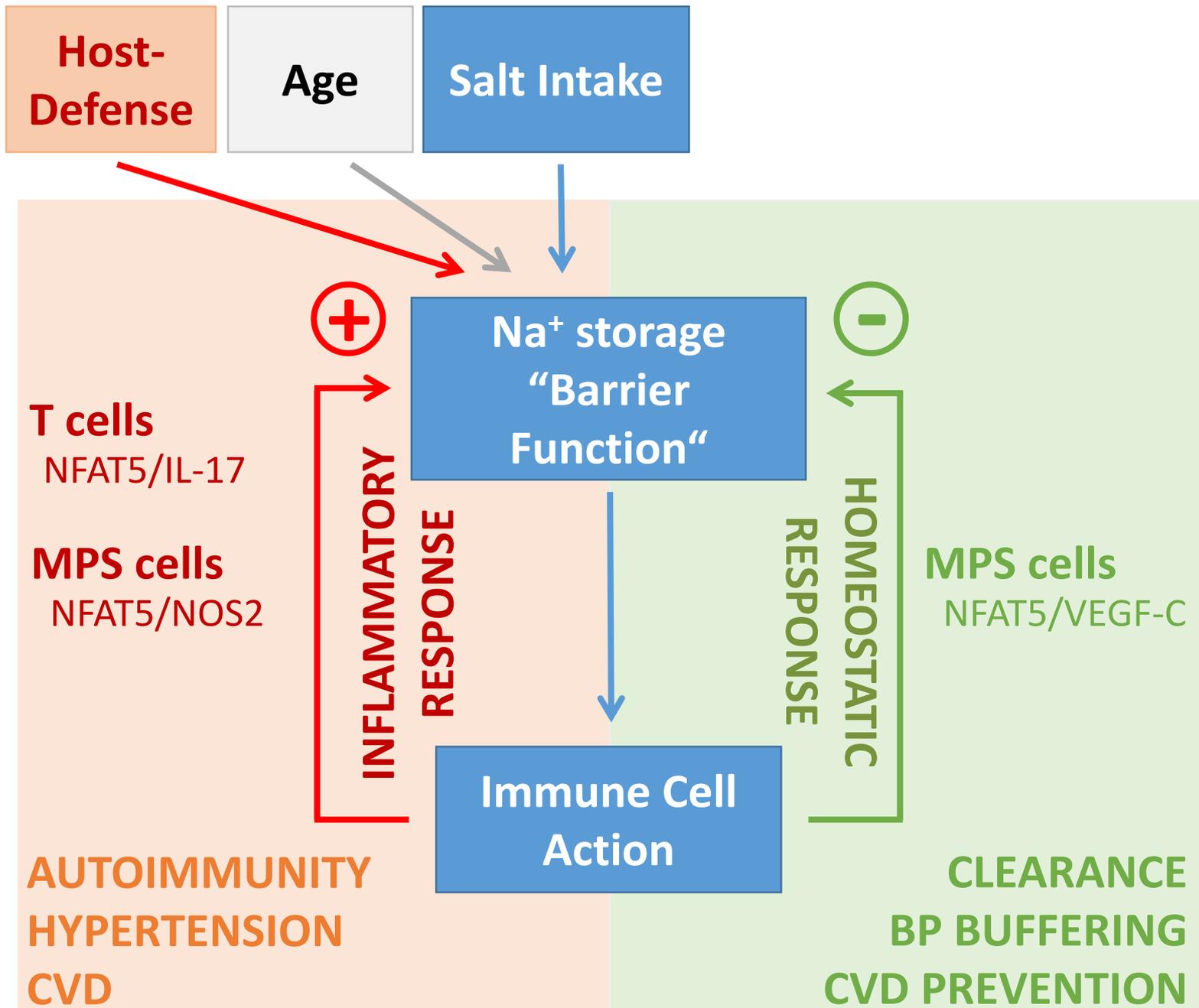
**Urine**

	LSD	HSD
Blood: [Na <sup>+</sup> + K <sup>+</sup> ]	143 ± 2 mM	145 ± 2 mM
Skin: [Na <sup>+</sup> + K <sup>+</sup> ]	177 ± 8 mM	191 ± 7 * mM



# Macrophages and skin electrolytes and pressure





**Host-Defense**

**Age**

**Salt Intake**

**(+)**

**Na<sup>+</sup> storage  
"Barrier  
Function"**

**(-)**

**T cells**  
NFAT5/IL-17

**MPS cells**  
NFAT5/NOS2

**INFLAMMATORY  
RESPONSE**

**HOMEOSTATIC  
RESPONSE**

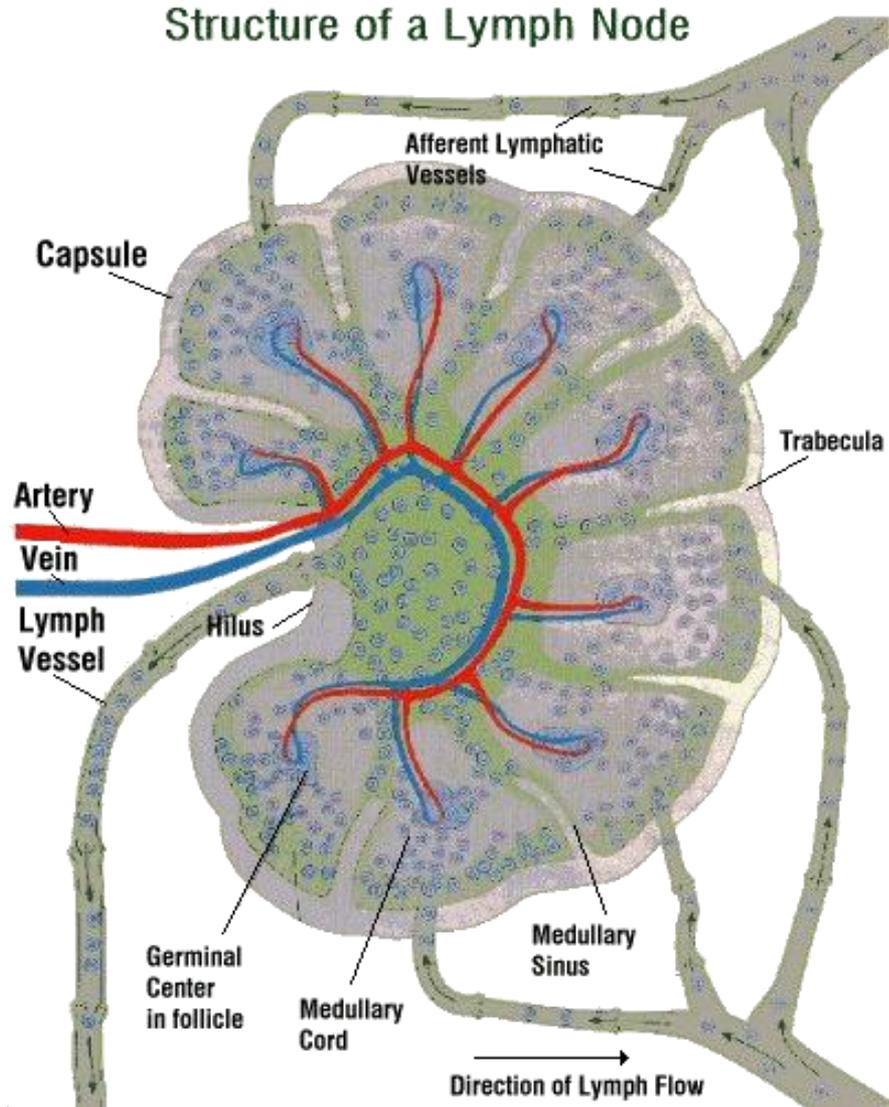
**Immune Cell  
Action**

**MPS cells**  
NFAT5/VEGF-C

**AUTOIMMUNITY  
HYPERTENSION  
CVD**

**CLEARANCE  
BP BUFFERING  
CVD PREVENTION**

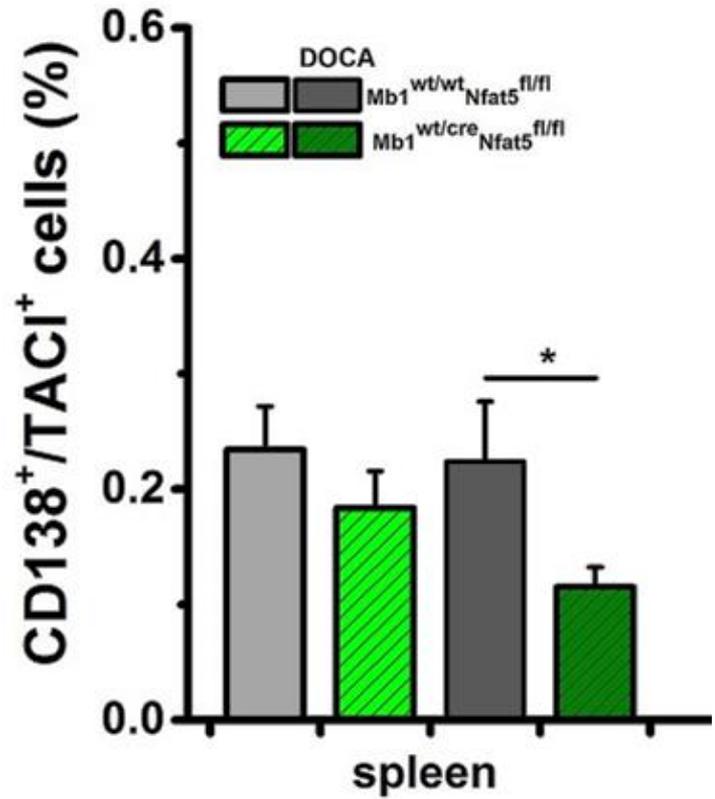
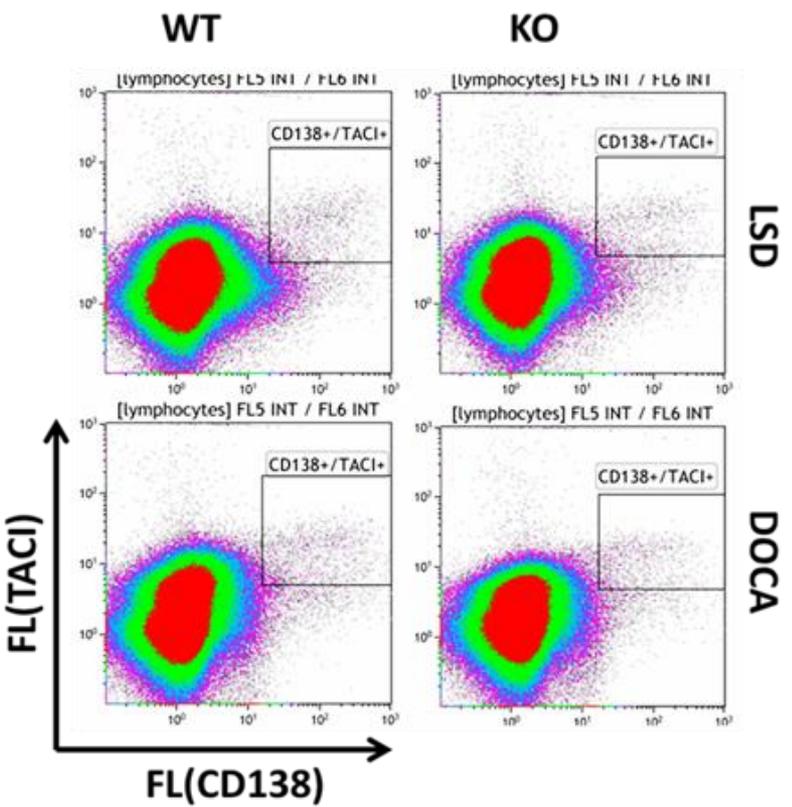
# B cells in salty microenvironments



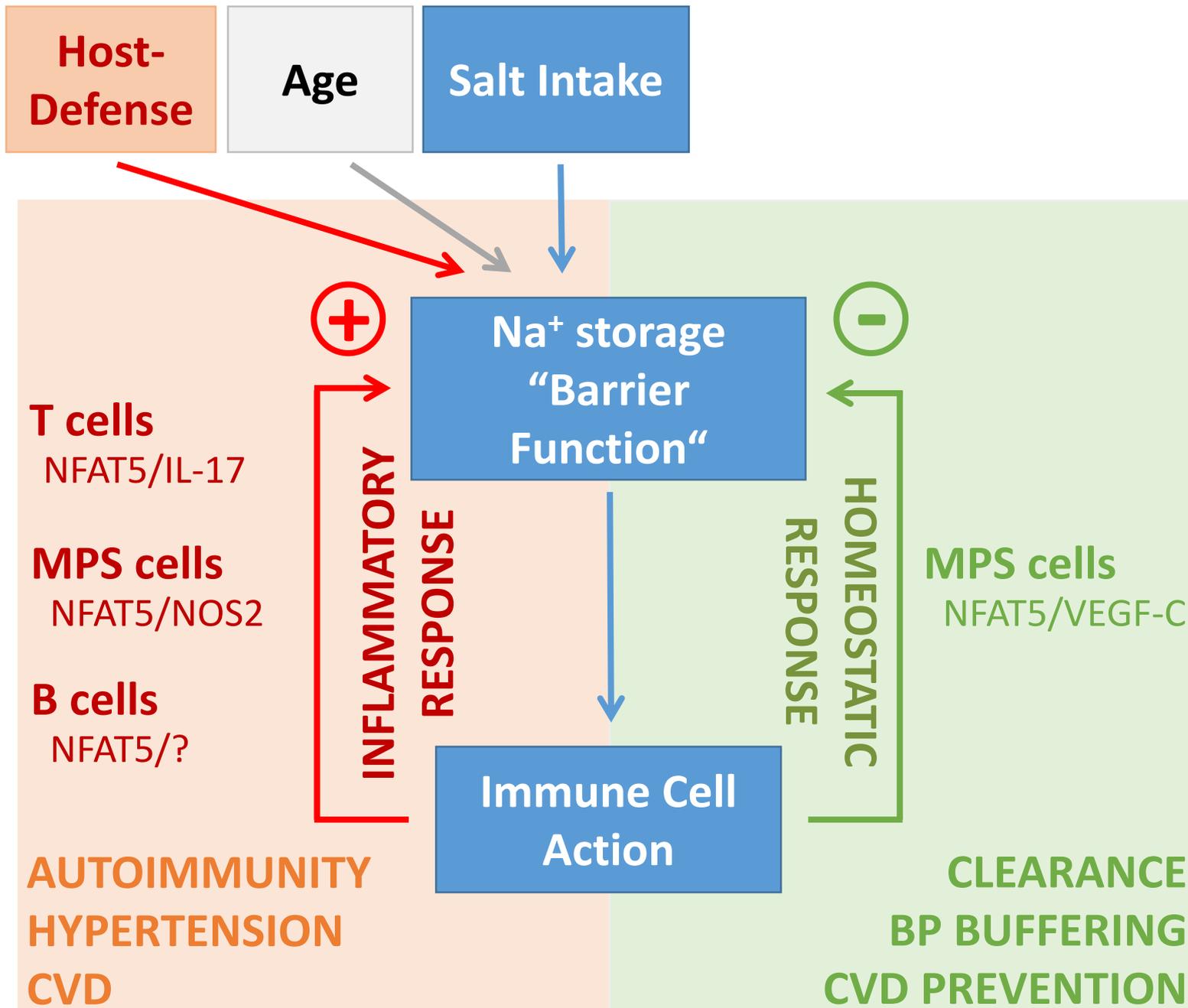
# B cells in salty microenvironments

Lilijana Rokvic  
 Wolfgang Schuh  
 Hans-Martin Jäck

## Spleen plasma cells/blasts

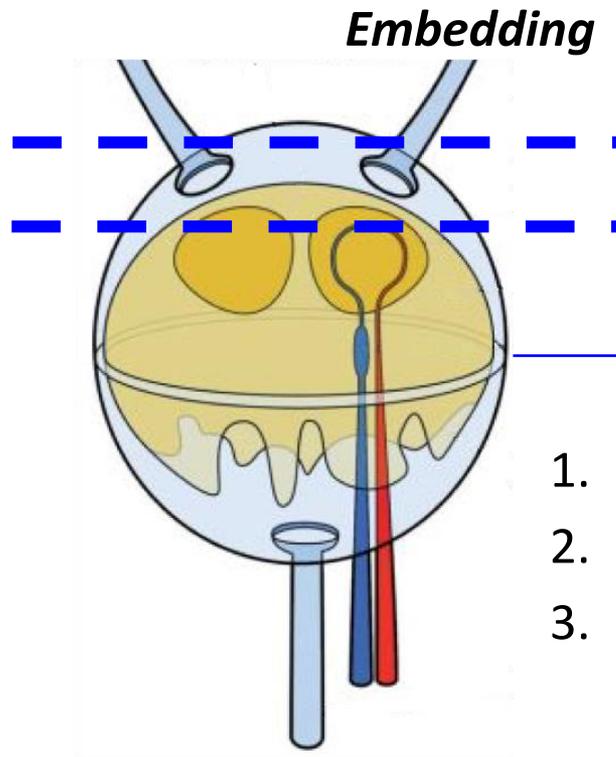


work in progress



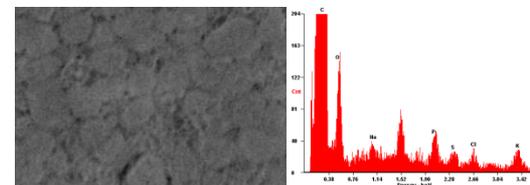
# Salty microenvironments: a need to bring material sciences into medicine

Stojan Perisic  
Joachim Spatz



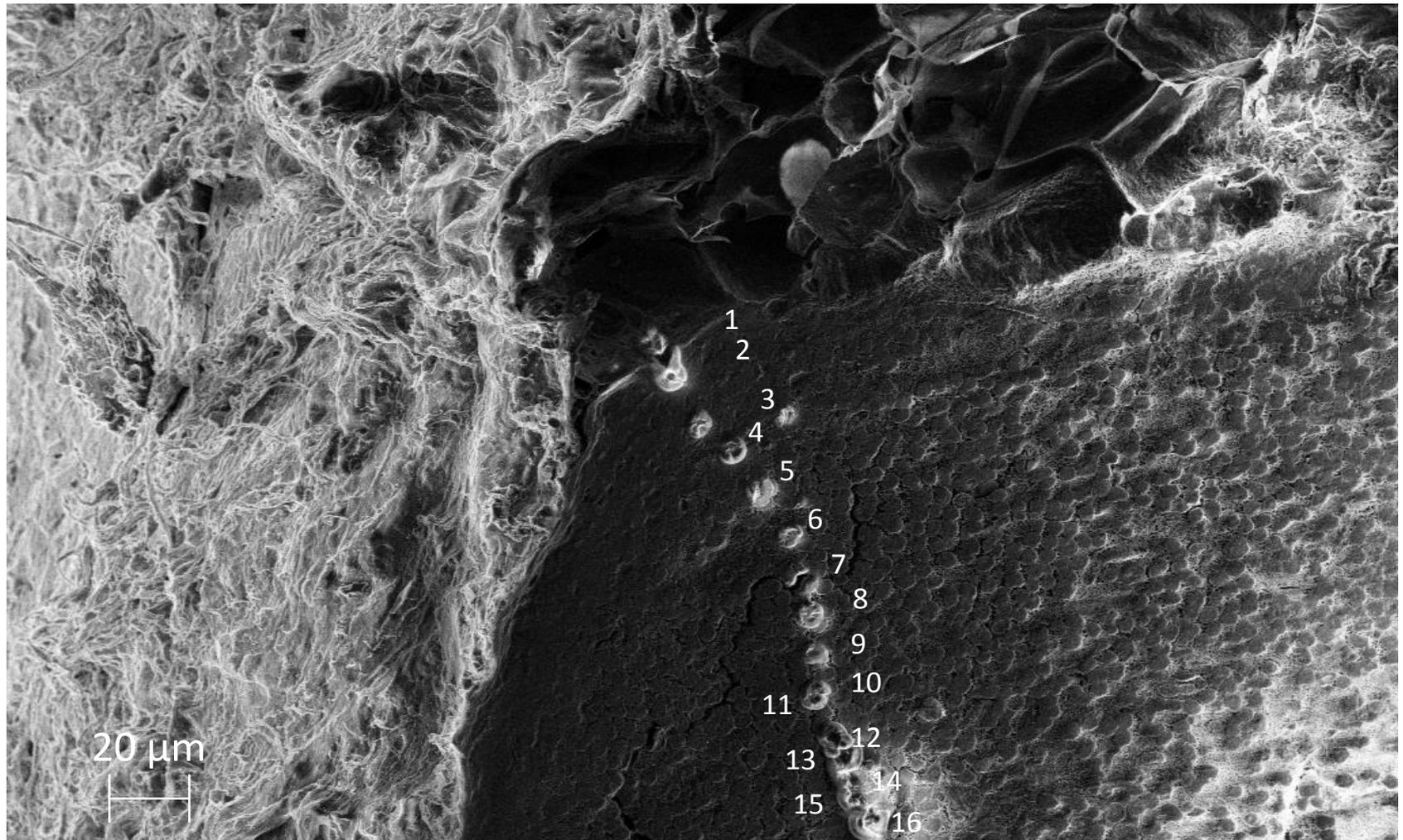
1. Planing
2. Freeze-drying
3. SEM+EDX

Rosendaal et al 2009, modified



work in progress

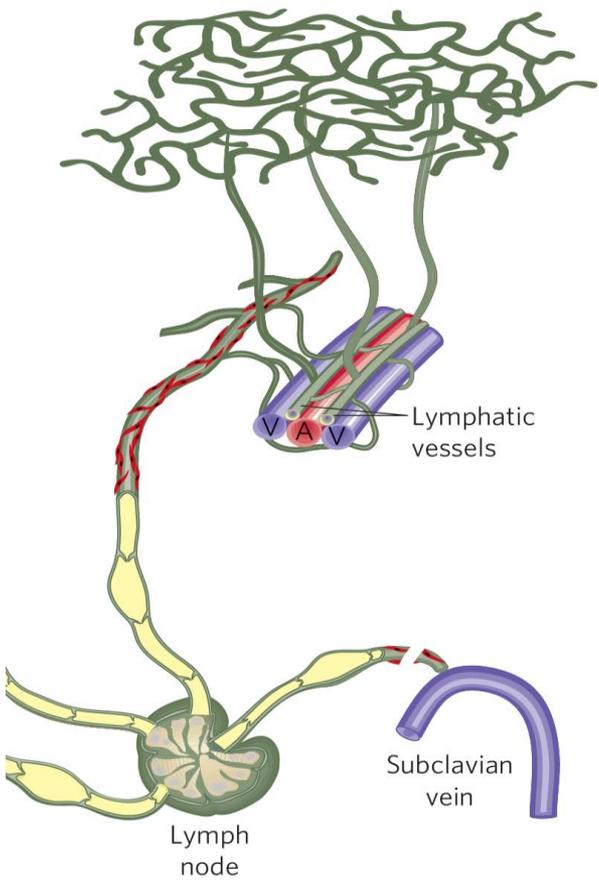
# Salty microenvironments: a need to bring material sciences into medicine



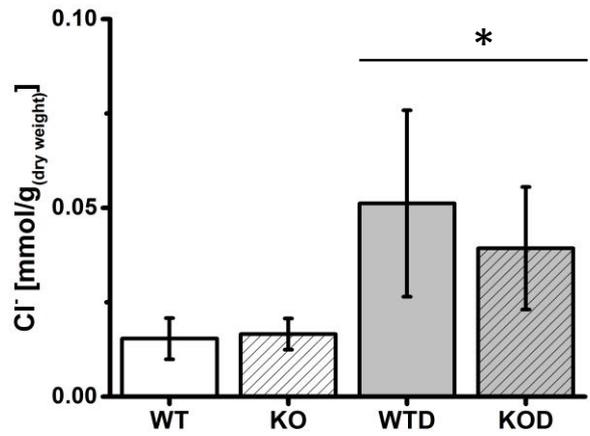
work in progress

# B cells in salty microenvironments

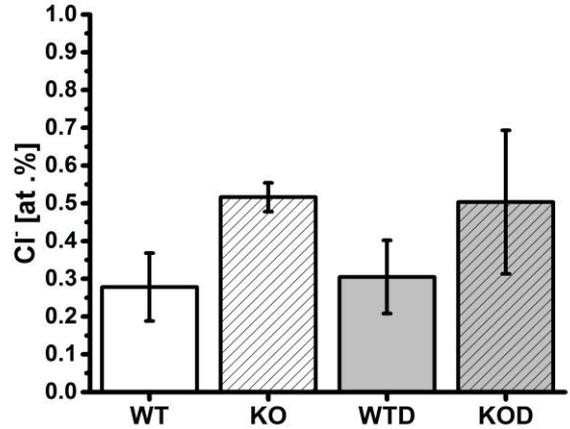
mb1<sup>cre</sup>Nfat5<sup>fllox</sup> mice:



Skin



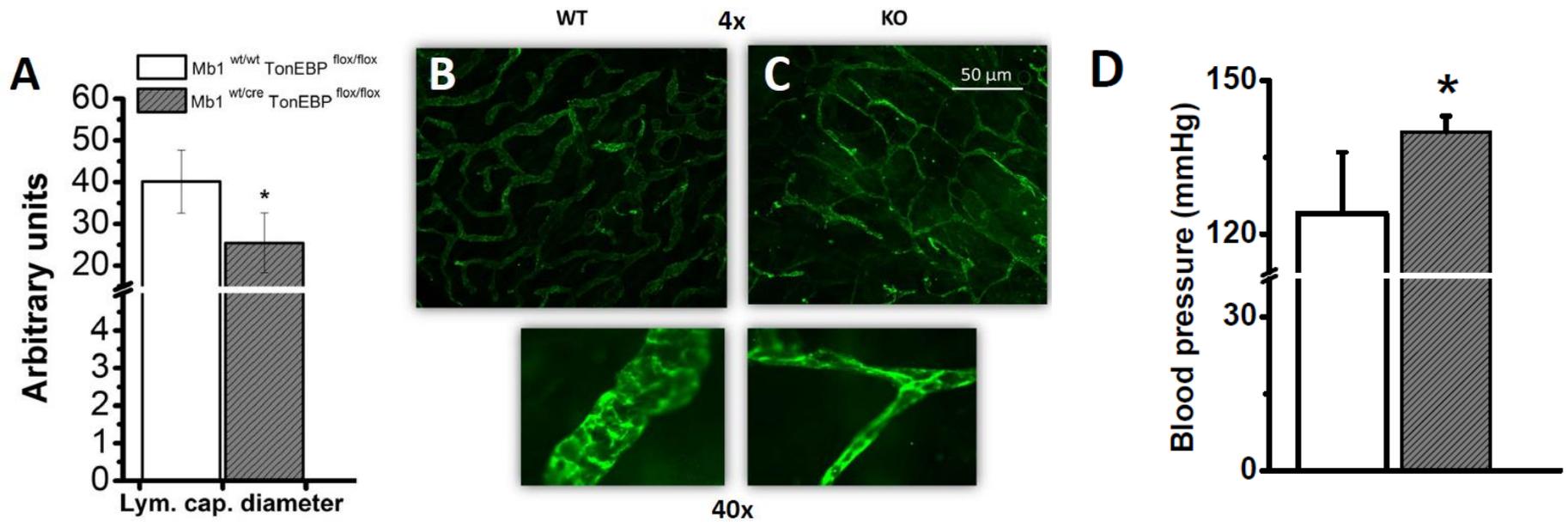
Lymph Node



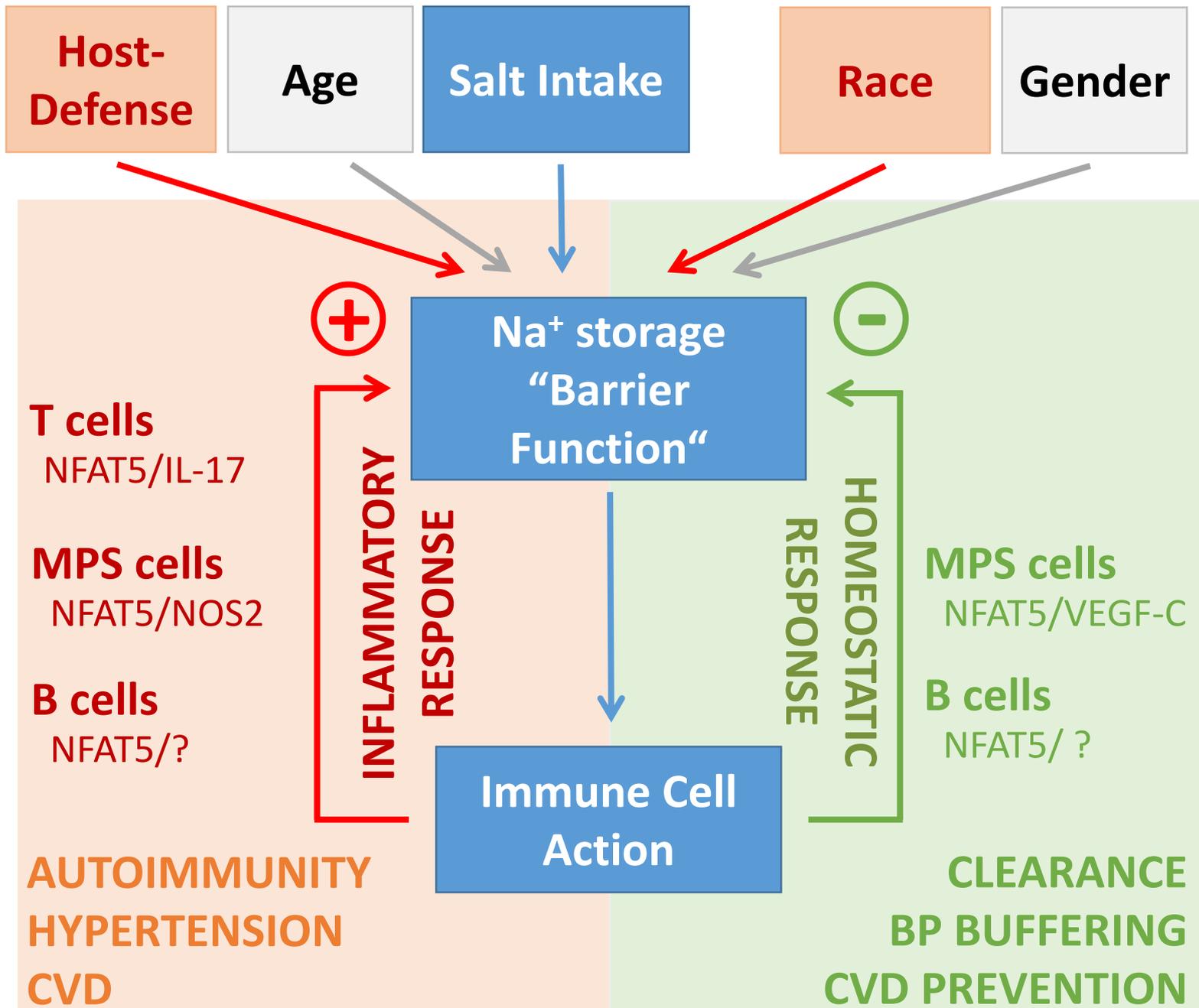
work in progress

# B cells in salty microenvironments

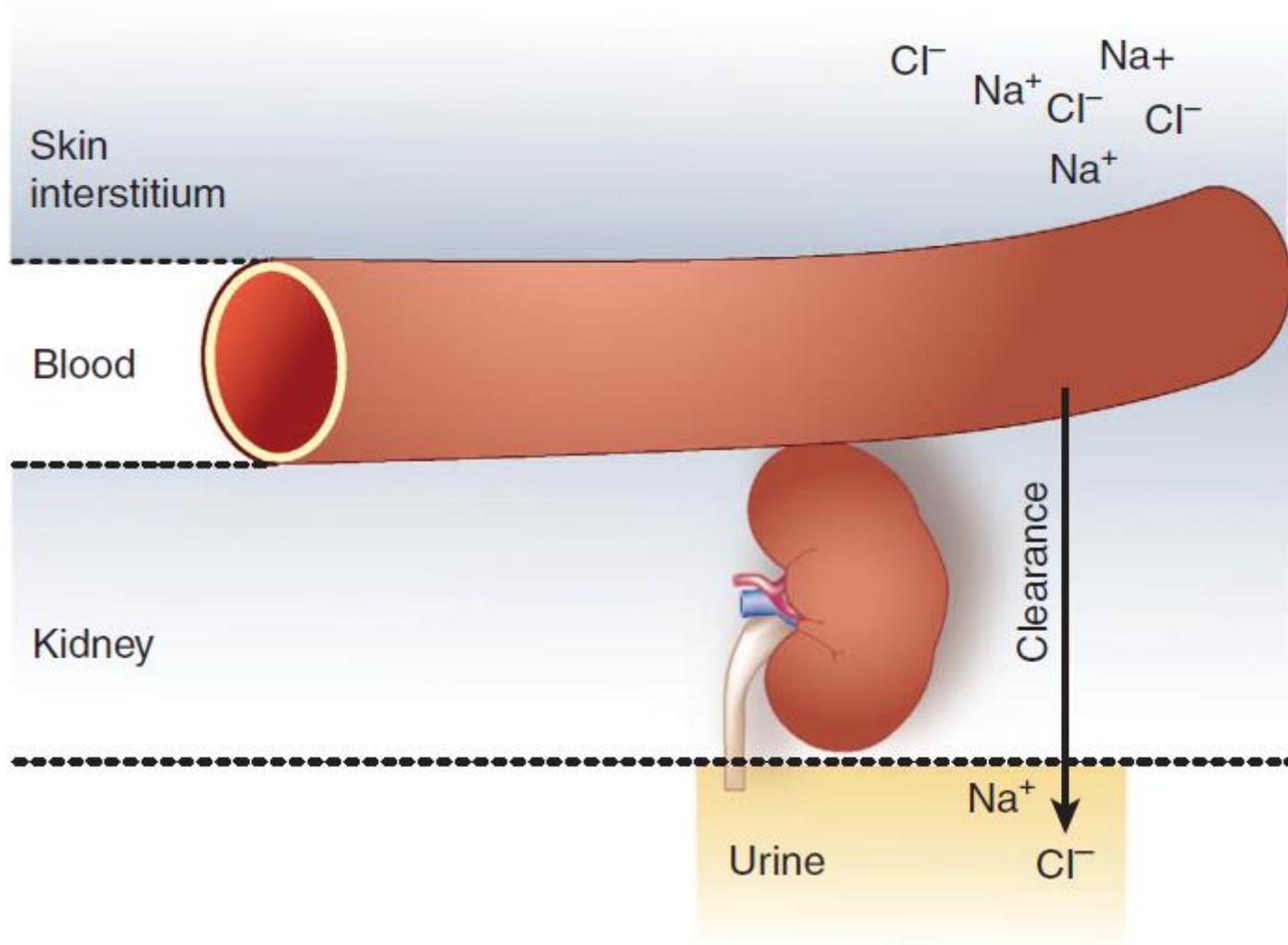
$mb1^{cre}Nfat5^{flox}$  mice: salt-sensitive hypertension



work in progress



# State of the art – and an alternative



# State of the art – and an alternative

