

## HUMORAL AND CELLULAR IMMUNE RESPONSE INDUCED BY NOVEL SARS-COV-2 VACCINES

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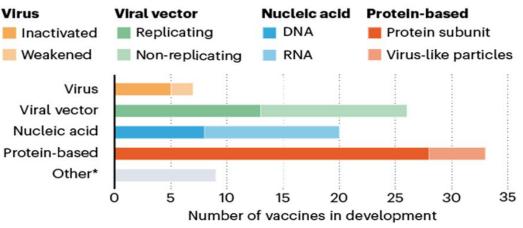


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### Background – COVID pandemic

- Severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2)
- 2020- global public health crisis
- Respiratory symptoms, multi-organ manifestations
- No effective treatment
- Vaccine development different platforms

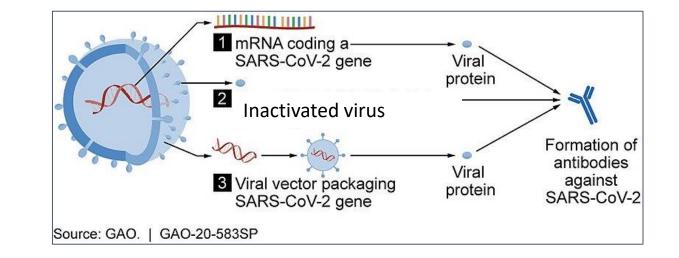
#### AN ARRAY OF VACCINES



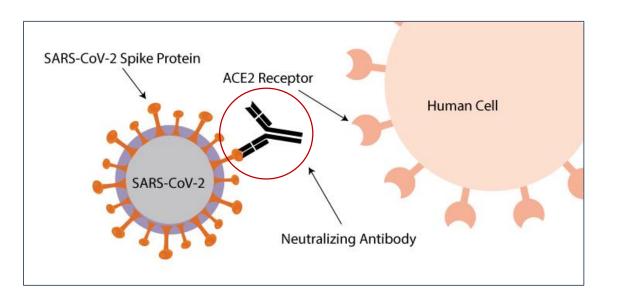
\* Other efforts include testing whether existing vaccines against poliovirus or tuberculosis could help to fight SARS-CoV-2 by eliciting a general immune response (rather than specific adaptive immunity), or whether certain immune cells could be genetically modified to target the virus.

### COVID-19 vaccines used in Hungary

- Inactivated  $\rightarrow$  whole virus
- mRNA and vector  $\rightarrow$  spike protein



- Spike protein binds to ACE2 receptor on the target cells
- Neutralizing antibodies are directed against Spike1 protein RBD



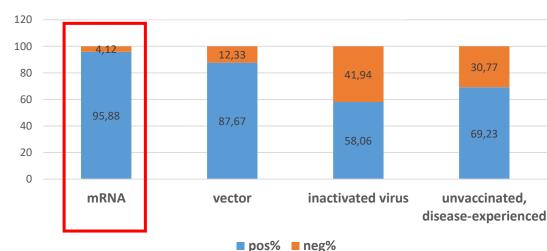
### Investigation of the immune response after vaccination

Investigated groups	<u>Number of</u> <u>collected</u> <u>samples</u>
mRNA-based vaccine (Pfizer-Biontech)	106
vector vaccines (AstraZeneca, Sputnik V)	77
inactivated virus vaccine (Sinopharm)	34
unvaccinated healthy	9
unvaccinated PCR-confirmed disease-experienced	29
TOTAL	255

Peripheral blood samples were taken 2-3 months after second vaccination/infection

## Measurement of the IgG antibodies directed against the spike protein of SARS-CoV-2

(p<0.001) (p=0.004)(p<0.001) (p<0.001 .0000 (p=0.004) Anti-SARS-CoV-2 lgG 0 ,0000 .0000 .0000 ,0000 .0000 unvaccinated, healthy disease-experienced inactivated virus unvaccinated, mRNA vector

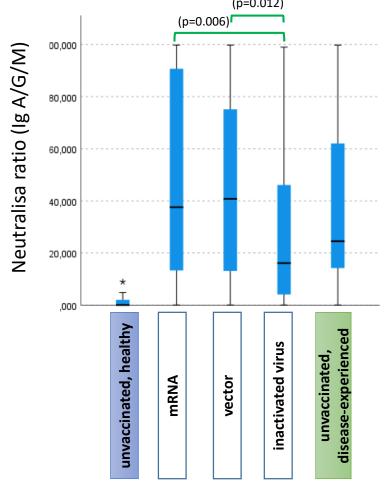


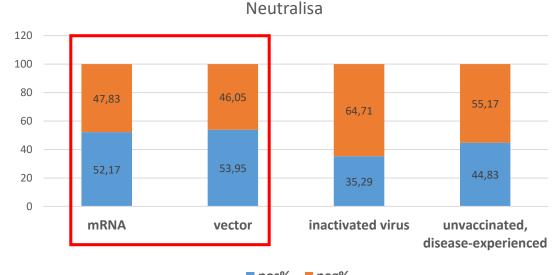
Anti-SARS-CoV-2 lgG

The mRNA vaccine induced the highest anti-SARS-CoV-2 S1 IgG antibody production

Anti-SARS-CoV-2 S1 IgG antibody Euroimmun ELISA

### Examination of the neutralizing antibody against SARS-CoV-2 spike protein



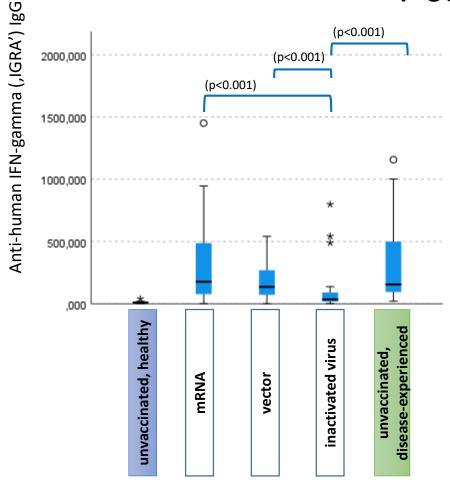


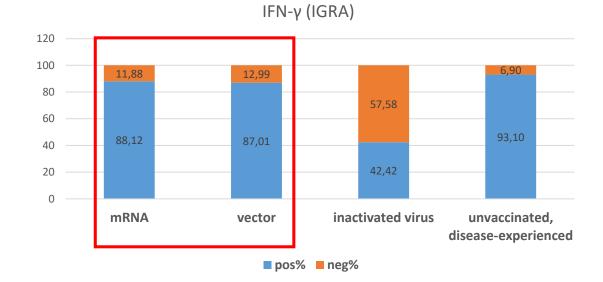
pos% neg%

The mRNA and vector vaccine groups had the highest amount of neutralizing antibody

NeutraLISA – ACE2 receptor S1 RBD inhibition assay (Euroimmun)

# Investigation of the SARS-CoV-2 S1 protein specific T cell immune response





The mRNA and vector vaccines induced as high IFN-gamma production as the natural infection

SARS-CoV-2 S1 protein specific T cell IFN-gamma release assay (IGRA) (Euroimmun)

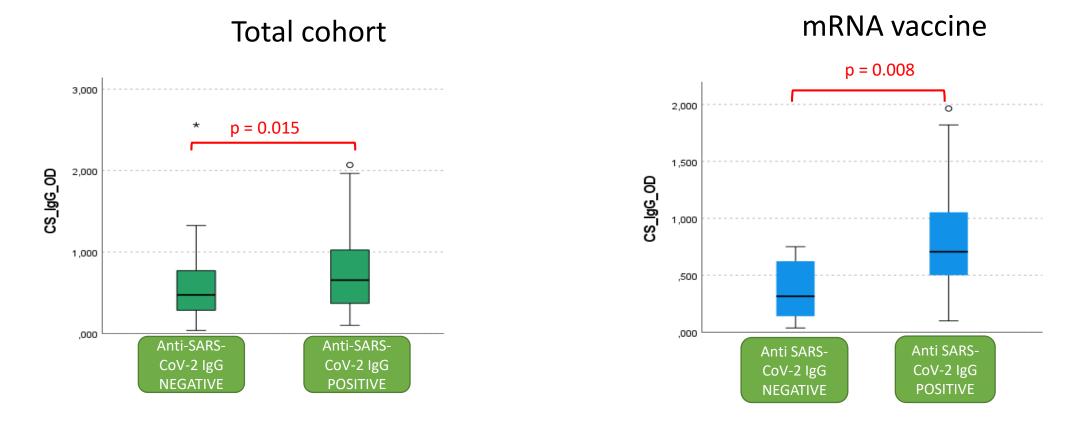
### Conclusion I.

- The mRNA-based vaccine induced the highest anti-SARS-CoV-2 antibody level
  - The mRNA and vector vaccines showed similarly good results in the cellular response and neutralizing antibody production
- Lower effectiveness of the inactivated virus vaccine may be due to the vaccine components and older age of the recipients

### Background – natural autoantibody

- Personalized natural autoantibody network
- First line of defense, maintanence of the immune homeostatis
- Recognizing evolutionarily conserved structures  $\rightarrow$  protection against autoimmunity
- Autoimmune phenomena  $\rightarrow$  molecular mimicry
- Possible connection between vaccine/infection induced humoral response and natural autoantibodies

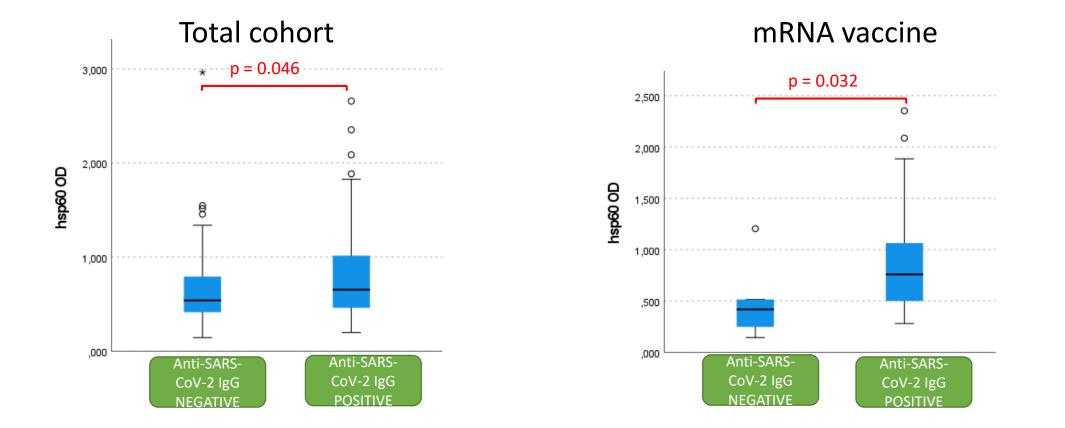
Measurement of anti-citrate synthase (CS) natural autoantibody in anti-SARS-CoV-2 IgG positive and negative goups



SARS-CoV-2 IgG positive group had higher anti-CS IgG autoantibody level

Anti-CS IgG autoantibody in-house ELISA

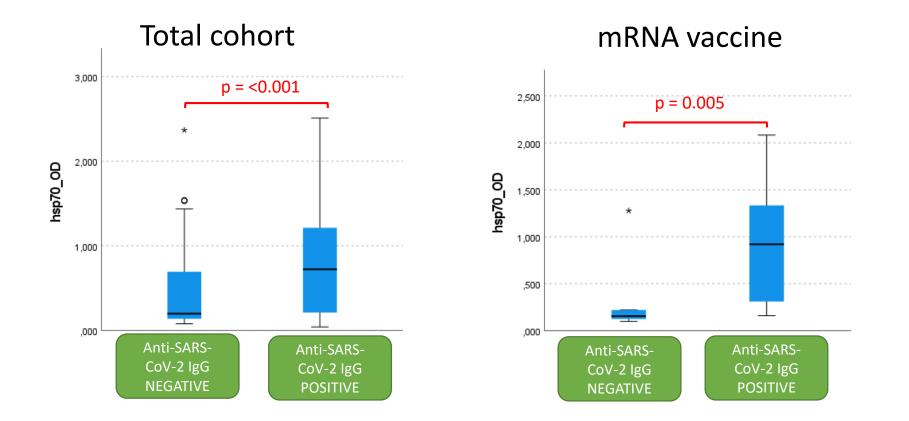
Level of natural autoantibody against Heat shock protein (Hsp) 60 in anti-SARS-CoV-2 IgG positive and negative goups



In the SARS-CoV-2 IgG positive group anti-Hsp60 IgG autoantibody level was higher

#### Anti-Hsp60 IgG autoantibody in-house ELISA

Investigation of natural autoantibody against Hsp70 in anti-SARS-CoV-2 IgG positive and negative goups



In the SARS-CoV-2 IgG positive group showed higher anti-Hsp70 IgG autoantibody level

Anti-Hsp70 IgG autoantibody in-house ELISA

### **Final conclusion**

- The mRNA vaccine elicited the strongest humoral and cellular immune responses against the spike protein of SARS-CoV-2
- COVID-19 vaccine or SARS-CoV-2 infection induced IgG isotype antibodies positively correlated with natural autoantibody levels
- The natural autoantibodies may serve as potential screening targets to predict the strength of antigen-induced immune response



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