

Supplementary Material

Immune Responses to SARS-CoV-2 Infection and Vaccination in Dialysis Patients and Kidney Transplant Recipients

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Baseline characteristics	BNT162b2 vaccinated dialysis patients, n=475	mRNA-1273 vaccinated dialysis patients, n=138	p-value
Age, yrs (± SD)	70 (± 14)	61 (± 16)	<0.001
Sex, n (%)			
Female	182 (38.3)	51 (37.0)	0.772
Pre-existing kidney disease, n (%)			
Glomerulonephritis	85 (17.9)	27 (19.6)	0.655
Diabetic nephropathy	68 (14.3)	35 (25.4)	0.002
Hypertensive nephropathy	118 (24.8)	33 (23.9)	0.824
ADPKD	31 (6.5)	11 (8.0)	0.554
Other genetic nephropathy	62 (13.1)	10 (7.2)	0.062
Unknown / other	111 (23.4)	22 (15.9)	0.062
Haemodialysis, n (%)	449 (94.5)	138 (100.0)	
Years of Haemodialysis (IQR)	4 (2-6)	4 (1-8)	0.387
CAPD, n (%)	26 (5.5)	0	0.005
Years of CAPD (IQR)	2 (1-4)	0	n. a.

Supplemental Table S1a: Baseline characteristics of SARS-CoV-2 naïve dialysis patients vaccinated with BNT162b2 or mRNA-1273.

Abbreviations: ADPKD: Autosomal Dominant Polycystic Kidney Disease; CAPD: Continuous Ambulatory Peritoneal Dialysis; IQR: Interquartile

Range; n. a.: not applicable; SARS-CoV-2: Severe Acute Respiratory Syndrome Coronavirus Type 2; SD: Standard Deviation; yrs: Years.

	Sum of squares	Df	F	p-value
Regression	450259224.4	3	19.770	<0.001
Residual	4198147851	553		
Total	4648407075	556		

Coefficients	Standard Error	t	p-value
Vaccine used	281.992	3.248	0.001
Assay used	220.439	-1.379	0.169
Time between sampling and vaccination	6.194	0.973	0.331

Supplemental Table S1b: Differences in SARS-COV2 IgG in SARS-COV2 naïve dialysis patients vaccinated with BNT162b2 or mRNA-1273 adjusted for used Immunoassay (RBD vs. S1) and time between sampling and vaccination by a linear multiple regression model (corrected $R^2=0,092$) Two vaccinations with mRNA-1273 are correlated with higher SARS-COV2 IgG compared to two vaccinations with BNT162b2

Baseline characteristics	mRNA-vaccinated dialysis patients, n=613	Convalescent dialysis patients receiving a BNT162b2 booster shot, n=38	p-value
Age, yrs (\pm SD)	68 (\pm 15)	69 (\pm 16)	0.526
Sex, n (%)			
Female	233 (38.0)	18 (47.4)	0.197
Pre-existing kidney disease, n (%)			
Glomerulonephritis	112 (18.3)	3 (7.9)	0.104
Diabetic nephropathy	103 (16.8)	8 (21.1)	0.499
Hypertensive nephropathy	151 (24.6)	10 (26.3)	0.816
ADPKD	42 (6.8)	6 (15.8)	0.041
Other genetic nephropathy	72 (11.7)	4 (10.5)	0.820
Unknown / other	140 (22.8)	7 (18.4)	0.527
Haemodialysis, n (%)	587 (95.8)	37 (97.4)	0.629
Years of Haemodialysis (IQR)	4 (2-6)	3 (2-6)	0.693
CAPD, n (%)	26 (4.2)	1	0.629
Years of CAPD (IQR)	2 (1-4)	13	0.444

Supplemental Table S2a: Baseline characteristics of SARS-CoV-2 naïve dialysis patients vaccinated with BNT162b2 or mRNA-1273 compared to COVID-19 convalescent dialysis patients receiving a BNT162b2 booster shot. Abbreviations: ADPKD: Autosomal Dominant Polycystic Kidney Disease; CAPD: Continuous Ambulatory Peritoneal Dialysis; IQR: Interquartile Range; SARS-CoV-2: Severe Acute Respiratory Syndrome Coronavirus Type 2; SD: Standard Deviation; yrs: Years.

	Sum of squares	Df	F	p-value
Regression	671958428.9	3	28.647	<0.001
Residual	4917378248	585		
Total	5589336677	588		

Coefficients	Standard Error	t	p-value
Booster vs. 2 mRNA vaccinations	525.822	4.749	< 0.001
Assay used	242.729	1.692	0.091
Time between sampling and vaccination	6.268	1.563	0.119

Supplemental Table S2b: Differences in SARS-COV2 IgG SARS-CoV-2 naïve dialysis patients vaccinated with BNT162b2 or mRNA-1273 compared to COVID-19 convalescent dialysis patients receiving a BNT162b2 booster shot after adjusting for used assay (RBD vs. S1) and time between sampling and vaccination in days by a linear multiple regression model (corrected $R^2=0,116$) *Receiving a single Booster Vaccination with BNT162b2 after infection is associated with higher SARS-COV2 IgG compared to receiving two mRNA vaccinations in dialysis patients adjusted for used Assays und time between sampling and vaccination*

Anti-RBD (BAU/ml), median (IQR), n=29	Anti-S1 (BAU/ml), median (IQR), n=29	Spearman Correlation Coefficient	p-value
390.0 (793.0)	493.5 (875.7)	0.99	<0.0001

Supplemental Table S3a: Correlation between SARS-COV2 IgG measured by S1-Antigen and RBD Immunoassay in 29 dialysis patients 30 days after vaccination with BNT162b2 (BioNTech / Pfizer). Anti-SARS-CoV-2 immunoglobulin G (IgG) targeting the spike (S) protein were detected by the semiquantitative Euroimmun anti-SARS-CoV-2 IgG ELISA on the Euroimmun Analyzer I (Euroimmun Diagnostik, Lübeck, Germany) and the quantitative IDK® anti-SARS-CoV-2 IgG ELISA (Immundiagnostik AG, Bensheim, Germany) on the DYNEX DSX® (Dynex Technologies, Chantilly, VA, USA), both using the recombinant S1 antigen of the spike protein, and the chemiluminescent microparticle immunoassays by Abbott for the quantitative detection of anti-spike RBD (receptor binding domain) IgG (SARS-CoV-2 IgG II Quant) on the automated system Alinity i (Abbott, Abbott Park, IL, United States). Quantitative IgG values were measured in BAU (binding antibody unit/ml).

N 29	Mann-Whitney-U- Test	Z	Sig. (2- tailed)
BAU/ml	392.00	-0.433	0.658

Supplemental Table S3b: Comparisons of mean SARS-COV2 IgGs measured by S1-Antigen and RBD Immunoassay in 29 dialysis patients 30 days after vaccination with BNT162b2 (BioNTech / Pfizer). Anti-SARS-CoV-2 immunoglobulin G (IgG) targeting the spike (S) protein were detected by the semiquantitative Euroimmun anti-SARS-CoV-2 IgG ELISA on the Euroimmun Analyzer I (Euroimmun Diagnostik, Lübeck, Germany) and the quantitative IDK® anti-SARS-CoV-2 IgG ELISA (Immundiagnostik AG, Bensheim, Germany) on the DYNEX DSX® (Dynex Technologies, Chantilly, VA, USA), both using the recombinant S1 antigen of the spike protein, and the chemiluminescent microparticle immunoassays by Abbott for the quantitative detection of anti-spike RBD (receptor binding domain) IgG (SARS-CoV-2 IgG II Quant) on the automated system Alinity i (Abbott, Abbott Park, IL, United States). Quantitative IgG values were measured in BAU (binding antibody unit/ml). Sig.: Significance

Spearman Correlation	
Coefficient	p-value
0.265	0.05

Supplemental Table S4a: Correlation of timespan between transplantation and vaccination in years and SARS-COV-2 IgG: *Years after transplantation and SARS-COV2 IgG are barely positive correlated*

	Sum of squares	Df	F	p-value
Regression	1767049.688	4	4.180	0.004
Residual	8667150.479	82		
Total	10434200.167	86		

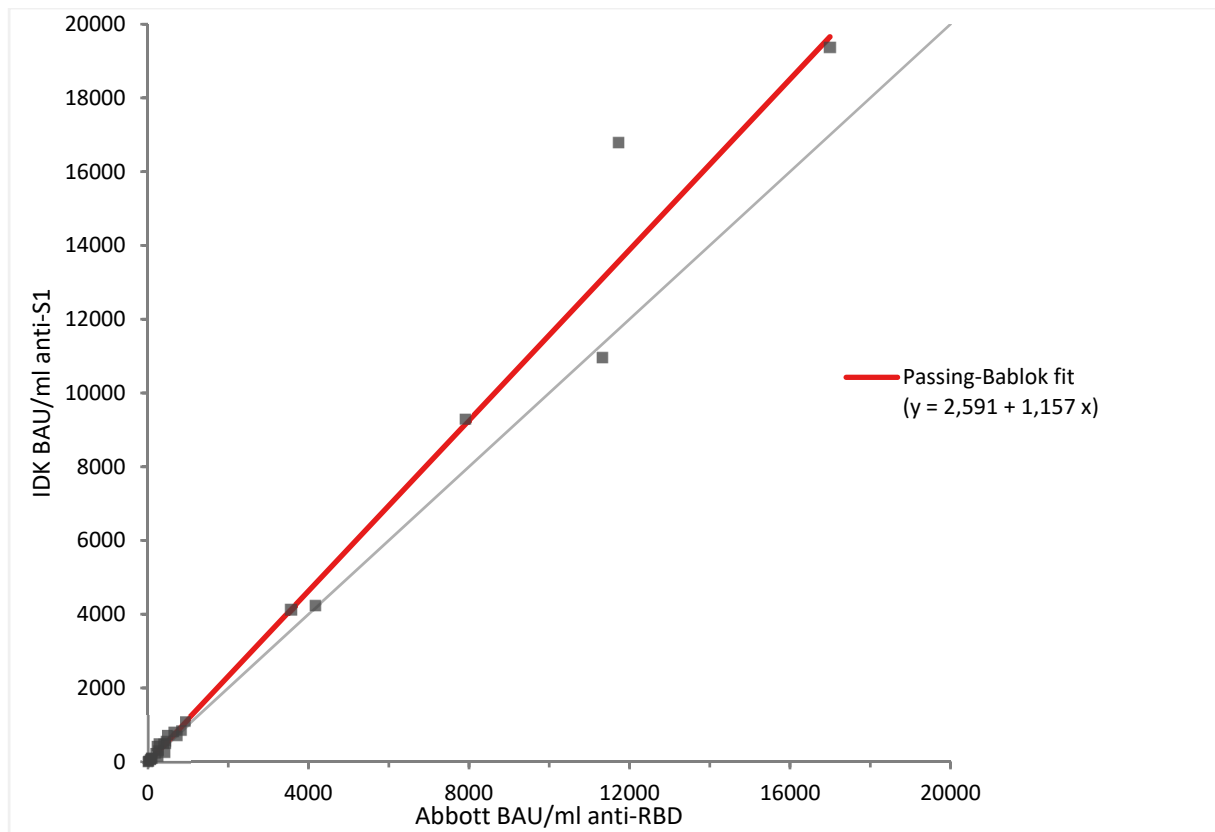
Coefficients	Standard Error	t	p-value
Assay used	68.481	2,089	0.04
MPA / MMF	79.968	-2.433	0.017
Age (yrs)	2.005	0.011	0.992
Time between sampling and vaccination	1.536	0.732	0.466

Supplemental Table S4b: Linear regression model of lacking seroconversion and immunosuppressive regimens with Mycophenolate Mofetil / Mycophenolic Acid adjusted for assay used, age and time between sampling in days and vaccination.

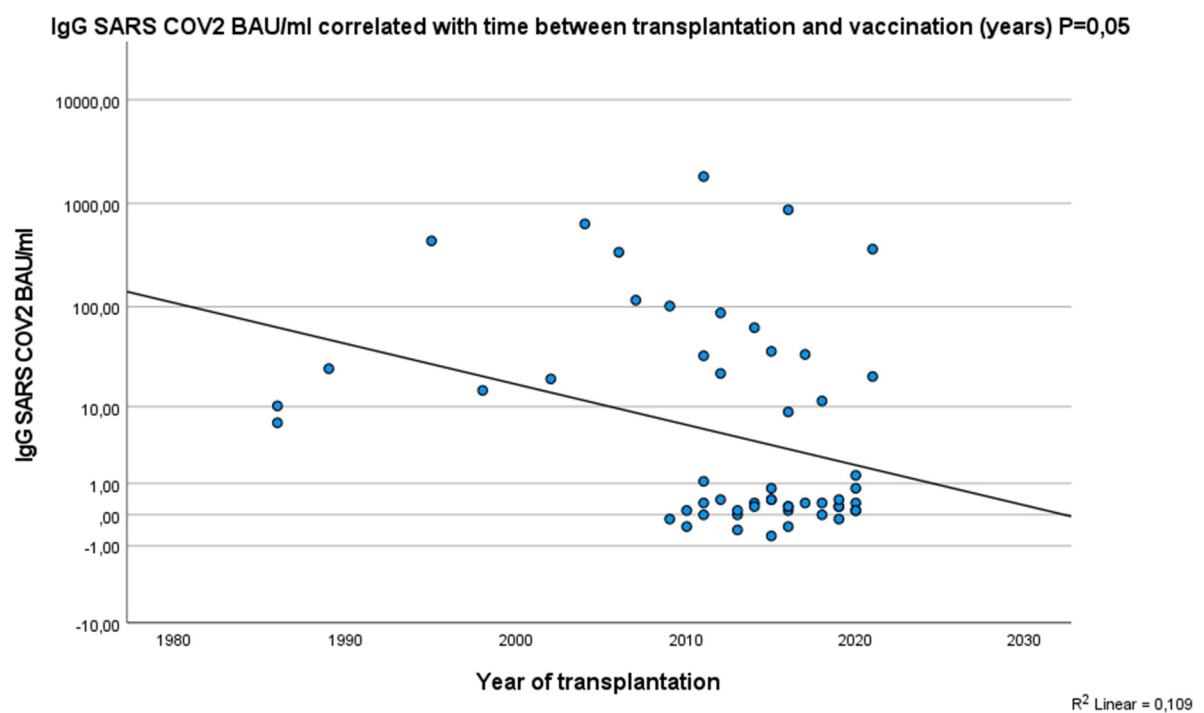
Abbreviations: MMF: Mycophenolate Mofetil; MPA: Mycophenolic Acid; yrs: Years. *MPA/MMF therapy is negatively correlated with SARS-COV2 IgG after two BNT162b2 vaccination adjusted for Age, time between sampling and vaccination and assay used*

Baseline characteristics	Kidney transplant recipients, n=32
Age, yrs (\pm SD)	55 (\pm 13)
Sex, n	
Female	11
Male	22
Immunosuppression, n	
Tacrolimus	32
Cyclosporin A	0
Belatacept	0
mTOR	0
Azathioprine	1
MMF / MPA	27
Steroid	30
Underlying kidney disease, n	
Glomerulonephritis	10
Diabetic nephropathy	1
Hypertensive nephropathy	2
ADPKD	10
Other genetic nephropathy	2
Unknown / other	7
Kidney transplantation mode, n	
Deceased	9
Living donor	23
Living donor AB0i	5
Vaccine used, n	
BNT162b2 (BionTech / Pfizer)	29
mRNA-1273 (Moderna)	3

Supplemental Table S5: Baseline characteristics of SARS-CoV-2 naive kidney transplant recipients with prospectively monitored for vaccine-induced humoral and cellular immunity. Abbreviations: ADPKD: Autosomal Dominant Polycystic Kidney Disease; IQR: Interquartile Range; MMF: Mycophenolate Mofetil; MPA: Mycophenolic Acid; mTOR: mechanistic Target Of Rapamycin; SD: Standard Deviation; yrs: Years.



Supplemental Figure S1: Passing-Bablok-Regression for SARS-COV2 IgG Assays targeting targeting S1 or RBD: Intercept 2,6 (95%-CI: -10,7 to 14,7); Slope 1,16 (95%-CI: 1,0 to 1,3).



**Supplemental Figure S2: IgG SARS COV2 BAU/ml correlated with time between transplantation and vaccination: X: logarithmic representation of IgG levels in BAU/ml
Y: Year of Transplantation**