

## Tables and Figures

**Table 1: Comparison of early-stage and late-stage pleural effusions of LTX recipients.**

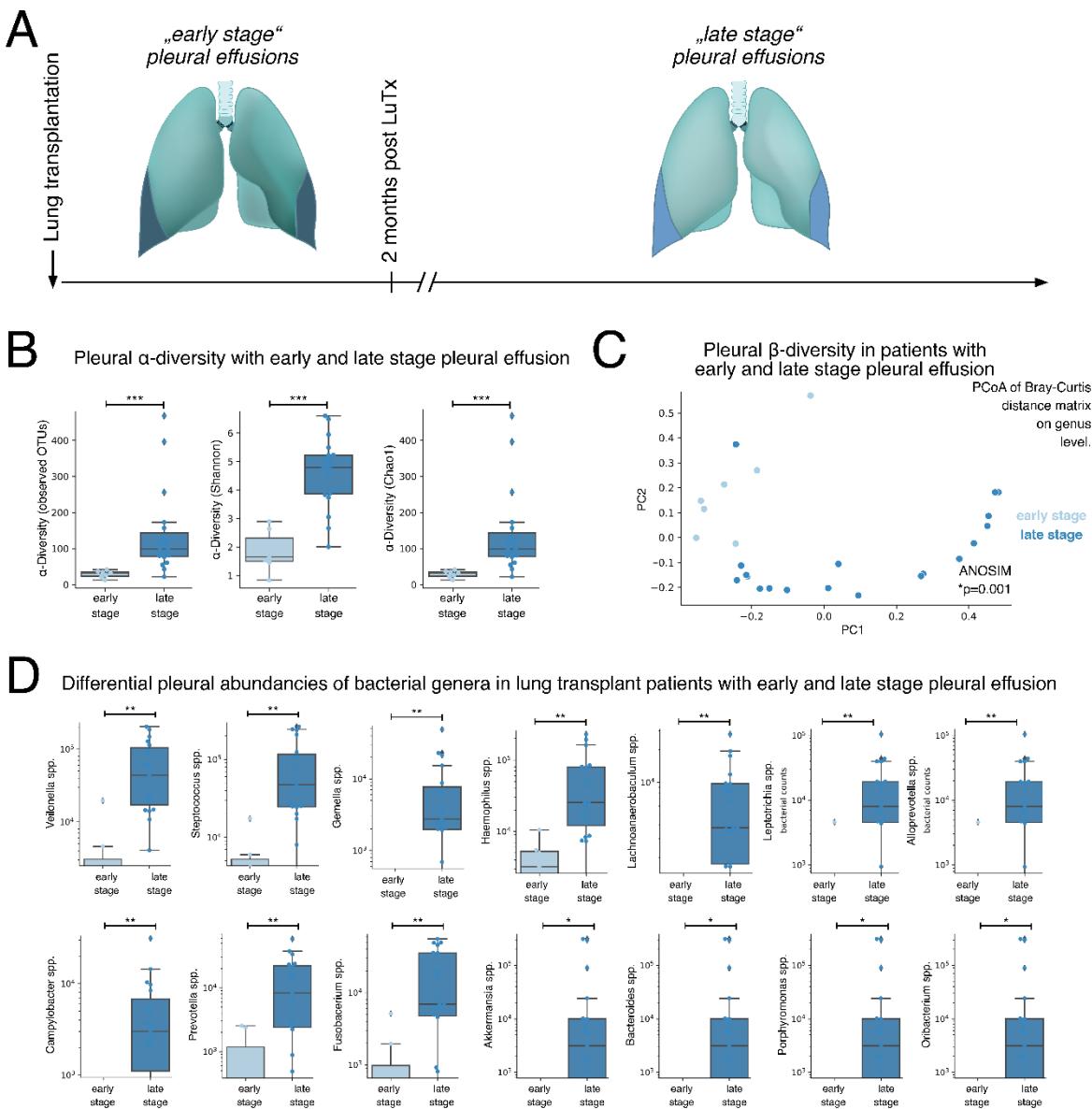
CHARACTERISTICS	EARLY STAGE	LATE STAGE	P
NUMBER OF PATIENTS	n=7	n=19	
AGE AT PE (YEARS), MEDIAN (IQR)	56.89 (53.87-61.23)	59.56 (54.94-61.64)	0.470 <sup>#</sup>
SEX, N (%) FEMALE PATIENTS	1/7 (14.3%)	6/19 (31.6%)	0.629 <sup>o</sup>
SEX, N (%) MALE PATIENTS	6/7 (85.7%)	13/19 (68.4%)	
TIME TO PE (DAYS AFTER LTX), MEDIAN (IQR)	21 (20-24)	188 (131-513)	0.0001 <sup>#</sup>
PREVIOUS THORACOCENTESIS, N (%)	1/7 (14%)	7/19 (37%)	0.269 <sup>o</sup>
<b>TRANSPLANT TYPE</b>			
SINGLE LTX, N (%)	-	1/19 (5.3%)	
DOUBLE LTX, N (%)	7/7 (100.0%)	18/19 (94.7%)	
<b>ETIOLOGY OF PLEURAL EFFUSION</b>			
POSTOPERATIVE, N (%)	7/7 (100.0%)	-	
UNCLEAR, N (%)	-	16 (84.2%)	
CHYLOTHORAX, N (%)	-	2 (10.5%)	
RENAL FAILURE, N (%)	-	1 (5.3%)	
<b>CLASSIFICATION OF PLEURAL EFFUSION</b>			
EXUDATE, N (%)	7/7 (100.0%)	15 (78.9%)	
TRANSUDATE, N (%)	-	4 (21.1%)	
<b>UNDERLYING DISEASE FOR LTX PATIENTS</b>			
ILD, N (%)	5/7 (71.4%)	13/19 (68.4%)	1.000 <sup>o</sup>
COPD, N (%)	0/7 (0%)	3/19 (15.8%)	0.540 <sup>o</sup>
CF / BRONCHIECTASIS, N (%)	1/7 (14.3%)	2/19 (10.5%)	1.000 <sup>o</sup>
RE-LTX FOR CLAD, N (%)	1/7 (14.3%)	1/19 (5.3%)	0.474 <sup>o</sup>
<b>ANTIMICROBIAL PROPHYLAXIS / THERAPY</b>			
AZITHROMYCIN, N (%)	0/7 (0%)	8/19 (42.1%)	0.062 <sup>o</sup>
PCP PROPHYLAXIS, N (%)	7/7 (100%)	17/19 (89.5%)	1.000 <sup>o</sup>
CMV PROPHYLAXIS, N (%)	3/7 (42.9%)	8/19 (42.1%)	1.000 <sup>o</sup>
FUNGAL PROPHYLAXIS, N (%)	7/7 (100%)	14/19 (73.7%)	0.278 <sup>o</sup>
IV OR PO ANTIBIOTICS AT DRAINAGE, N (%)	2/7 (28.6%)	2/19 (10.5%)	0.287 <sup>o</sup>

**Table 1:** Comparison of pleural effusions occurring early (n=7) (up to 60 days) vs. late (n=19) after lung transplantation.  
**Statistics:** <sup>#</sup> unpaired Mann-Whitney test for non-parametric testing, <sup>o</sup> Chi-squared test for categorical data (Fisher exact test for 2 cells (>25%)).

**Table 2: Comparison of late-stage pleural effusions of LTX recipients vs. pleural effusions of non-transplant patients**

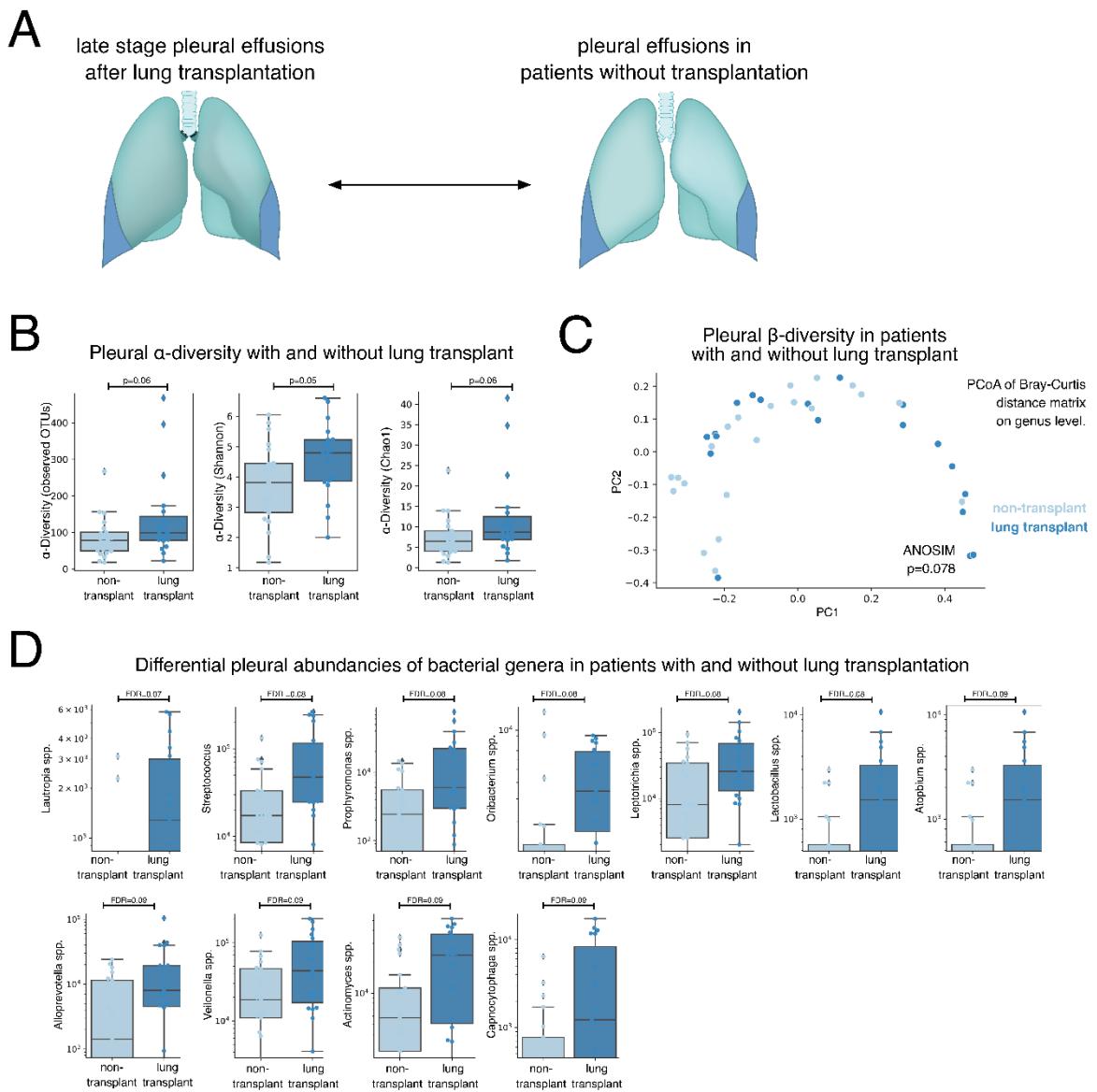
CHARACTERISTICS	LUNG TRANSPLANT	NON-TRANSPLANT	P
NUMBER OF PATIENTS	n=19	n=21	
AGE AT PE (YEARS), MEDIAN (IQR)	59.56 (54.94-61.64)	75.71 (65.79-82.43)	<b>0.0002<sup>#</sup></b>
SEX, N (%) FEMALE PATIENTS	6/19 (31.6%)	7/21 (33.3%)	0.906 <sup>o</sup>
SEX, N (%) MALE PATIENT	13/19 (68.4%)	14/21 (66.7%)	
TIME TO PE (DAYS AFTER LTX), MEDIAN (IQR)	184 (129.75-448.50)	-	
PREVIOUS THORACOCENTESIS, N (%)	7/19 (37%)	11/21 (52%)	0.324 <sup>o</sup>
TRANSPLANT TYPE			
SINGLE LTX, N (%)	1/19 (5.3%)	-	
DOUBLE LTX, N (%)	18/19 (94.7%)	-	
ETIOLOGY OF PLEURAL EFFUSION			
UNCLEAR, N (%)	16 (84.2%)	1 (4.8%)	
CHYLOTHORAX, N (%)	2 (10.5%)	-	
RENAL FAILURE, N (%)	1 (5.3%)	2 (9.5%)	
HEART FAILURE, N (%)	-	8 (38.1%)	
(PARA-)MALIGNANT, N (%)	-	8 (38.1%)	
RHEUMATOID, N (%)	-	1 (4.8%)	
(PARA-)INFECTION, N (%)	-	1 (4.8%)	
CLASSIFICATION OF PLEURAL EFFUSION			
EXUDATE, N (%)	15 (78.9%)	11 (52.4%)	
TRANSUDATE, N (%)	4 (21.1%)	10 (47.6%)	
UNDERLYING DISEASE FOR LTX PATIENTS			
ILD, N (%)	13/19 (68.4%)	-	
COPD, N (%)	3/19 (15.8%)	-	
CF / BRONCHIECTASIS, N (%)	2/19 (10.5%)	-	
RE-LTX FOR CLAD, N (%)	1/19 (5.3%)	-	
ANTIMICROBIAL PROPHYLAXIS / THERAPY			
AZITHROMYCIN, N (%)	8/19 (42.1%)	0/21 (0%)	0.001 <sup>o</sup>
PCP PROPHYLAXIS, N (%)	17/19 (89.5%)	2/21 (9.5%)	<0.0001 <sup>o</sup>
CMV PROPHYLAXIS, N (%)	8/19 (42.1%)	0/21 (0%)	0.001 <sup>o</sup>
FUNGAL PROPHYLAXIS, N (%)	14/19 (73.7%)	0/21 (0%)	<0.0001 <sup>o</sup>
IV OR PO ANTIBIOTICS AT DRAINAGE, N (%)	2/19 (10.5%)	7/21 (33.3%)	0.1328 <sup>o</sup>

**Table 2:** Comparison of late-stage pleural effusions in LTX recipients (n=19) vs. pleural effusions in non-transplant patients (n=21). Statistics: <sup>#</sup> unpaired Mann-Whitney test for non-parametric testing, <sup>o</sup> Chi-squared test for categorical data (Fisher exact test for >2 cells (25%)).



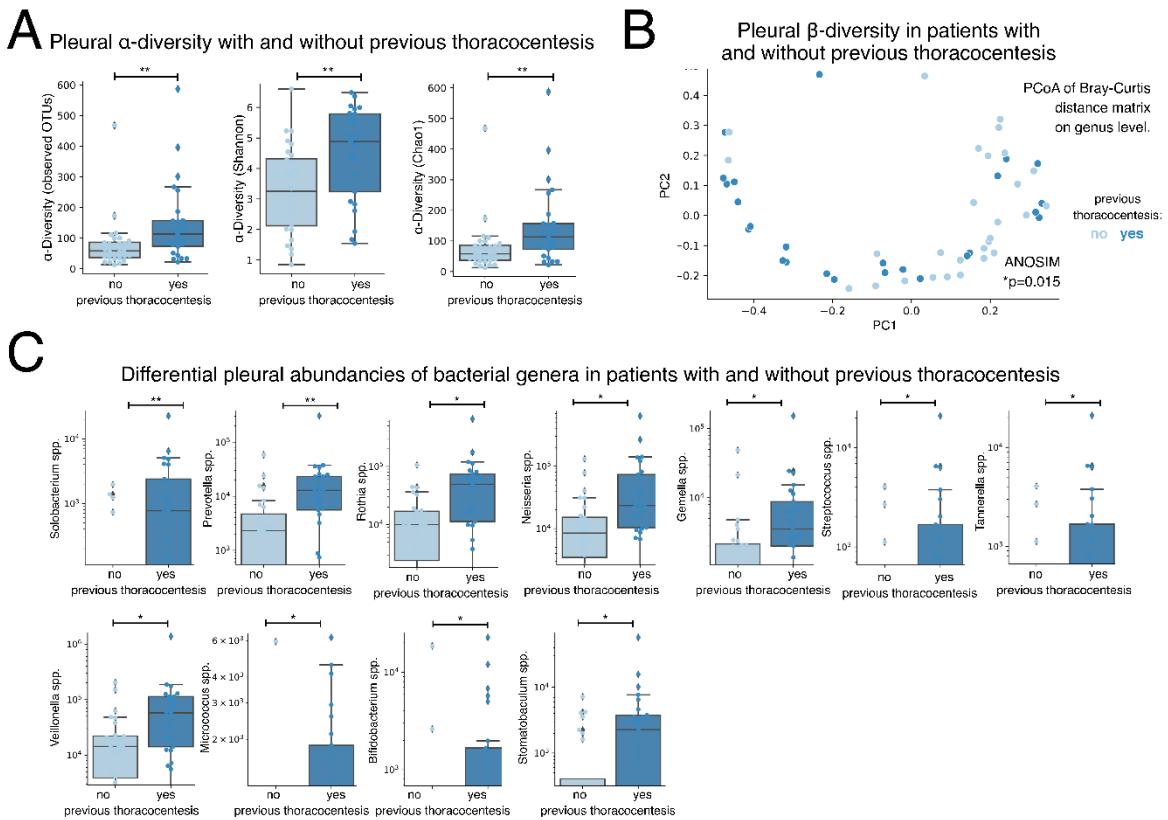
**Figure 1: Pleural microbiome differences between early- and late-stage pleural effusions after lung transplantation. (A)** Illustration of early-stage pleural effusions occurring less than 60 days after lung transplantation and late-stage pleural effusion seen later-on. **(B)** Differences between early-stage ( $n=7$ ) and late-stage ( $n=19$ ) pleural effusions regarding their  $\alpha$ -diversity quantified by three different non-phylogenetic  $\alpha$  diversity indices on OTU level. **(C)** Differences between early- and late-stage pleural effusions regarding their  $\beta$  diversity using a non-phylogenetic Bray-Curtis distance matrix and Principal Coordinate Analysis (PCoA) on genus level. **(D)** Differential abundances of bacterial genera between early- and late-stage pleural effusions post lung transplantation.

Statistics: For comparison of  $\alpha$  diversity Mann-Whitney-U test was used. For comparison of  $\beta$  diversity Analysis of Similarities (ANOSIM) was used. For comparison of differential abundances repeated Mann-Whitney-U test with multiple testing adjustment using the Benjamini-Hochberg procedure was used. \* $p<0.05$ , \*\* $p<0.01$ , \*\*\* $p<0.001$ .



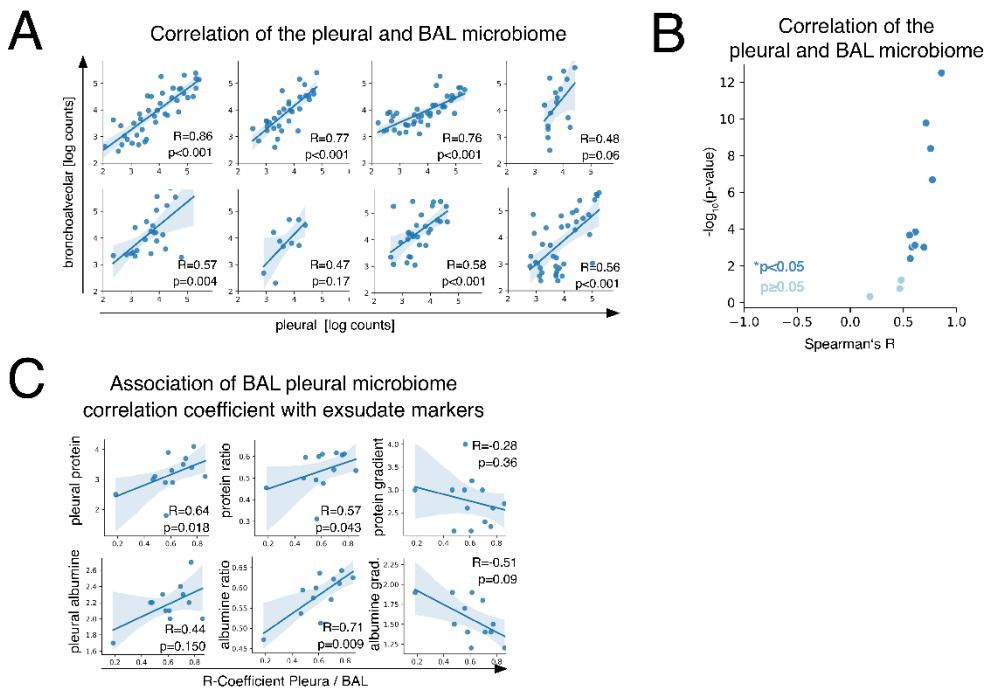
**Figure 2: Pleural microbiome differences between LTX recipients and non-transplant patients. (A)** Comparison of the pleural effusion microbiome of lung transplant recipients with late-stage pleural effusion microbiome ( $n=19$ ) with non-transplant patients ( $n=21$ ). **(B)** Differences between lung transplant pleural effusions with pleural effusions from non-transplant patients regarding their  $\alpha$  diversity quantified by three different non-phylogenetic  $\alpha$  diversity indices on OTU level. **(C)** Differences between lung transplant and non-transplant pleural effusions regarding their  $\beta$  diversity using a non-phylogenetic Bray-Curtis distance matrix and Principal Coordinate Analysis (PCoA) on genus level. **(D)** Differential abundances of bacterial genera between pleural effusions of lung transplant patients with pleural effusions on non-transplant patients.

Statistics: For comparison of  $\alpha$  diversity Mann-Whitney-U test was used. For comparison of  $\beta$ -diversity Analysis of Similarities (ANOSIM) was used. For comparison of differential abundances repeated Mann-Whitney-U test with multiple testing adjustment using the Benjamini-Hochberg procedure was used. \* $p<0.05$ , \*\*  $p<0.01$ , \*\*\* $p<0.001$ .



**Figure 3: Previous thoracenteses influence the pleural microbiome in pleural effusions.** **(A)** Differences between pleural effusions with (n=24) and without previous thoracentesis (n=28) regarding their  $\alpha$  diversity quantified by three different non-phylogenetic  $\alpha$ -diversity indices on OTU level. **(B)** Differences between pleural effusions with and without previous thoracentesis regarding their  $\beta$  diversity using a non-phylogenetic Bray-Curtis distance matrix and Principal Coordinate Analysis (PCoA) on genus level. **(C)** Differential abundances of bacterial genera between pleural effusions with and without previous thoracentesis.

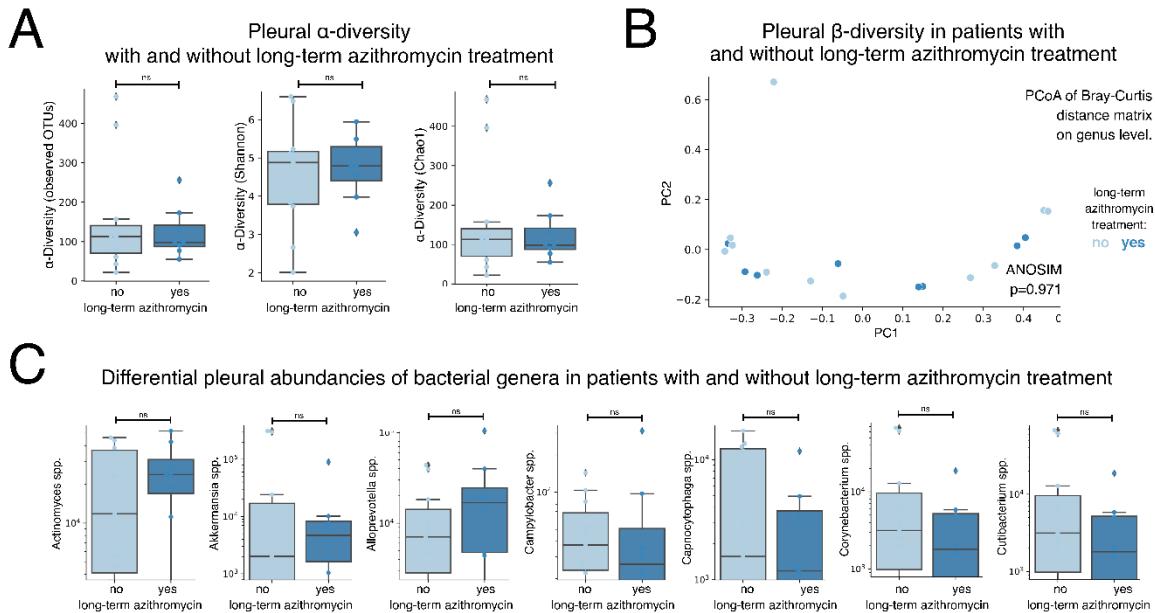
Statistics: For comparison of  $\alpha$  diversity Mann-Whitney-U test was used. For comparison of  $\beta$ -diversity Analysis of Similarities (ANOSIM) was used. For comparison of differential abundances repeated Mann-Whitney-U test with multiple testing adjustment using the Benjamini-Hochberg procedure was used. \*p<0.05, \*\* p<0.01, \*\*\*p<0.001.



**Figure 4: Correlation of the microbiome of pleural effusions and associated bronchoalveolar lavages (n=14).** (A) Representative regression plot of pleural and bronchoalveolar lavage (BAL) microbial DNA abundance on genus level. (B) Volcano plot of the Spearman's correlation coefficient and type II error. (C) Correlation of the Spearman's rank correlation coefficient between pleural and BAL microbiome and different exudate markers.

Statistics: For all correlations the Spearman's rank correlation coefficient was used.

## Supplementary Tables and Figures



**Figure S1: Long term azithromycin therapy does not influence the pleural microbiome in pleural effusions of LTX recipients.** **(A)** Differences between pleural effusions with (n=8) and without long-term azithromycin treatment (n=11) regarding their  $\alpha$  diversity quantified by three different non-phylogenetic  $\alpha$ -diversity indices on OTU level. **(B)** Differences between pleural effusions with and without long-term azithromycin treatment regarding their  $\beta$  diversity using a non-phylogenetic Bray-Curtis distance matrix and Principal Coordinate Analysis (PCoA) on genus level. **(C)** Differential abundances of bacterial genera between pleural effusions with and without long-term azithromycin therapy.

**Statistics:** For comparison of  $\alpha$  diversity Mann-Whitney-U test was used. For comparison of  $\beta$  diversity Analysis of Similarities (ANOSIM) was used. For comparison of differential abundances repeated Mann-Whitney-U test with multiple testing adjustment using the Benjamini-Hochberg procedure was used.

**Table S1: Laboratory findings of early-stage and late-stage pleural effusions in LTX recipients.**

CHARACTERISTICS	EARLY STAGE	LATE STAGE	P
<b>SERUM CHEMISTRY</b>		n=7	n=19
TOTAL PROTEIN (G/DL), MEAN (SD)	5.2 (4.6-5.8)	5.8 (5.6-6.1)	<b>0.030<sup>#</sup></b>
ALBUMIN (G/DL), MEAN (SD)	3.1 (2.9-3.2)	3.7 (3.2-3.9)	<b>0.012<sup>#</sup></b>
LDH (U/L), MEDIAN (IQR)	238 (205-254)	285 (241-392)	0.053 <sup>#</sup>
CRP (MG/DL), MEDIAN (IQR)	4.3 (0.8-6.3)	0.5 (0.2-2.3)	0.060 <sup>#</sup>
KREATININ (MG/DL), MEAN (SD)	0.9 (0.7-1.2)	1.5 (1.3-1.9)	<b>0.002<sup>#</sup></b>
NT-PROBNP (PG/ML), MEDIAN (IQR)	489 (45-970)	1938 (932-4541)	<b>0.013<sup>#</sup></b>
<b>PLEURAL FLUID CHEMISTRY</b>			
PROTEIN (G/DL), MEAN (SD)	2.7 (2.2-3.0)	3.1 (2.8-3.7)	0.132 <sup>#</sup>
ALBUMIN (G/DL), MEDIAN (IQR)	1.7 (1.0-2.0)	2.2 (2.0-2.4)	<b>0.005<sup>#</sup></b>
LDH (U/L), MEDIAN (IQR)	473 (206-906)	152 (134-194)	<b>0.005<sup>#</sup></b>
LDH (U/L) > 500 U/L, N (%)	3 (42.9%)	0	
LDH (U/L) > 1000 U/L, N (%)	1 (14.3%)	0	
TRIGLYCERIDES (MG/DL), MEDIAN (IQR)	24 (13-57)	17 (14-26)	0.397 <sup>#</sup>
TRIGLYCERIDES (MG/DL) > 110 MG/DL, N (%)	0	2	
CHOLESTERIN (MG/DL), MEDIAN (IQR)	66 (51-71)	60 (47-69)	0.647 <sup>#</sup>
HEMOGLOBIN (MG/DL), MEDIAN (IQR)	0.15 (0.05-3.10)	0.04 (0.02-0.25)	0.166 <sup>#</sup>
PH, MEDIAN (IQR)	7.42 (7.30-7.45)	7.37 (7.35-7.43)	0.859 <sup>#</sup>
LACTATE (MMOL/L), MEDIAN (IQR)	2.4 (2.1-4.3)	2.6 (1.9-3.8)	0.709 <sup>#</sup>
GLUCOSE (MG/DL), MEAN (SD)	109 (51-149)	127 (93-144)	0.456 <sup>#</sup>
<b>PLEURAL FLUID CELL COUNTS</b>			
TOTAL CELL COUNT (CELLS/µL), MEDIAN (IQR)	1592 (325-3577)	503 (258-1110)	0.312 <sup>#</sup>
POLYMORPHONUCLEAR (%), MEDIAN (IQR)	63 (9-74)	4 (3-20)	<b>0.004<sup>#</sup></b>
NEUTROPHIL GRAN. (%), MEDIAN (IQR)	12 (4-74)	3 (0-14)	0.067 <sup>#</sup>
EOSINOPHIL GRAN. (%), MEDIAN (IQR)	0 (0-2)	0 (0-0)	0.117 <sup>#</sup>
MONOCYTES (%), MEDIAN (IQR)	6 (2-13)	7 (4-13)	0.214 <sup>#</sup>
MACROPHAGES (%), MEDIAN (IQR)	0 (0-0)	9 (0-41)	<b>0.048<sup>#</sup></b>
LYMPHOCYTES (%), MEDIAN (IQR)	21 (5-70)	81 (54-86)	<b>0.043<sup>#</sup></b>
<b>MICROBIOLOGY</b>			
POS. BACTERIAL CULTURE, N (AVAILABLE N)	0 (7)	1 (19)	
POS. TBC MICROSCOPY, N (AVAILABLE N)	0 (4)	0 (17)	
POS. TBC PCR, N (AVAILABLE N)	0 (4)	0 (17)	
POS. TBC CULTURE, N (AVAILABLE N)	0 (4)	0 (17)	
POS. NTM PCR, N (AVAILABLE N)	-	0 (18)	

**Table S1:** Comparison of pleural effusions occurring early (n=7) (up to 60 days) vs. late (n=19) after lung transplantation.

**Statistics:** <sup>#</sup> unpaired Mann-Whitney test for non-parametric testing, <sup>o</sup> Chi-squared test for categorical data (Fisher exact test for >2 cells (25%)).

**Table S2: Laboratory findings of pleural effusions in LTX recipients and non-transplant patients.**

CHARACTERISTICS	LUNG TRANSPLANT	NON-TRANSPLANT	P
	n=19	n=21	
<b>SERUM CHEMISTRY</b>			
TOTAL PROTEIN (G/DL), MEAN (SD)	5.8 (5.6-6.1)	5.9 (5.6-6.6)	0.497#
ALBUMIN (G/DL), MEAN (SD)	3.7 (3.2-3.9)	3.7 (3.1-4.0)	0.966#
LDH (U/L), MEDIAN (IQR)	285 (241-392)	252 (189-494)	0.542#
CRP (MG/DL), MEDIAN (IQR)	0.5 (0.2-2.3)	3.5 (0.8-12)	<b>0.008#</b>
KREATININ (MG/DL), MEAN (SD)	1.5 (1.3-1.9)	1.4 (0.8-2.3)	0.322#
NT-PROBNP (PG/ML), MEDIAN (IQR)	1938 (932-4541)	3590 (1266-5915)	0.506#
<b>PLEURAL FLUID CHEMISTRY</b>			
PROTEIN (G/DL), MEAN (SD)	3.1 (2.8-3.7)	3.1 (2.2-4.1)	0.644#
ALBUMIN (G/DL), MEAN (SD)	2.2 (2.0-2.4)	1.7 (1.5-2.4)	0.371#
LDH (U/L), MEDIAN (IQR)	152 (134-194)	168 (107-445)	0.616#
LDH (U/L) > 500 U/L, N (%)	0	4 (19%)	
LDH (U/L) > 1000 U/L, N (%)	0	2 (9.5%)	
TRIGLYCERIDES (MG/DL), MEDIAN (IQR)	17 (14-26)	19 (11-29)	1.000#
TRIGLYCERIDES (MG/DL) > 110 MG/DL, N (%)	2	0	
CHOLESTERIN (MG/DL), MEDIAN (IQR)	60 (47-69)	44 (32-66)	0.155#
HEMOGLOBIN (MG/DL), MEDIAN (IQR)	0.04 (0.02-0.25)	0.02 (0.01-0.05)	0.327#
PH, MEDIAN (IQR)	7.37 (7.35-7.43)	7.44 (7.35-7.46)	0.055#
LACTATE (MMOL/L), MEDIAN (IQR)	2.6 (1.9-3.8)	1.7 (1.2-2.6)	0.091#
GLUCOSE (MG/DL), MEAN (SD)	127 (93-144)	113 (91-123)	0.279#
<b>PLEURAL FLUID CELL COUNTS</b>			
TOTAL CELL COUNT (CELLS/ $\mu$ L), MEDIAN (IQR)	503 (258-1110)	747 (348-2267)	0.119#
POLYMORPHONUCLEAR (%), MEDIAN (IQR)	4 (3-20)	10 (3-13)	0.462#
NEUTROPHIL GRAN. (%), MEDIAN (IQR)	3 (0-14)	9 (1-17)	0.146#
EOSINOPHIL GRAN. (%), MEDIAN (IQR)	0 (0-0)	0 (0-0)	0.965#
MONOCYTES (%), MEDIAN (IQR)	7 (4-13)	10 (4-26)	0.162#
MACROPHAGES (%), MEDIAN (IQR)	9 (0-41)	4 (2-10)	<b>0.049#</b>
LYMPHOCYTES (%), MEDIAN (IQR)	81 (54-86)	52 (12-66)	<b>0.018#</b>

**Table S2:** Comparison of late-stage effusions of LTX recipients (n=19) vs pleural effusions of non-transplant patients (n=21). Statistics: # unpaired Mann-Whitney test for non-parametric testing, ° Chi-squared test for categorical data (Fisher exact test for >2 cells (25%)).

**Table S3: Most abundant bacterial genera in pleural effusions after lung transplantation (n=26).**

BACTERIA GENUS	MEDIAN READS
STREPTOCOCCUS	25015
VEILLONELLA	20673
PREVOTELLA_7	18487
ROTHIA	17156
LEPTOTRICHIA	16259
HAEMOPHILUS	15706
NEISSERIA	12198
ACTINOMYCES	10925
STAPHYLOCOCCUS	6405
FUSOBACTERIUM	5618
ALLOPREVOTELLA	4805
CORYNEBACTERIUM	4717
PORPHYROMONAS	4085
PREVOTELLA	2756
LACHNOANAEROBACULUM	2683

**Table S4: Differential pleural abundancies of bacterial genera in early- (n=7) and late-stage (n=19) pleural effusions of LTX recipients.**

BACTERIA GENUS	SAMPLES WITH DETECTED GENUS	COUNTS	COUNTS	FDR
		EARLY STAGE [IQR]	LATE STAGE [IQR]	
VEILLONELLA	19/26	0 - 3.1x10 <sup>3</sup>	1.7x10 <sup>4</sup> - 1.0x10 <sup>5</sup>	0.0043
GEMELLA	7/26	0-0	2.0x10 <sup>3</sup> - 7.7x10 <sup>3</sup>	0.0043
HAEMOPHILUS	21/26	0-5.3x10 <sup>3</sup>	1.2x10 <sup>4</sup> - 8.0x10 <sup>4</sup>	0.0043
LACHNOANAEROBACULUM	9/26	0-0	1.7x10 <sup>3</sup> - 9.7x10 <sup>3</sup>	0.0043
STREPTOCOCCUS	20/26	0-5.2x10 <sup>3</sup>	2.5x10 <sup>4</sup> - 1.2x10 <sup>5</sup>	0.0043
LEPTOTRICHIA	20/26	0-4.7x10 <sup>3</sup>	1.3x10 <sup>4</sup> - 6.8x10 <sup>4</sup>	0.0043
ALLOPREVOTELLA	12/26	0-0	4.6x10 <sup>3</sup> - 1.9x10 <sup>4</sup>	0.0073
CAMPYLOBACTER	6/26	0-0	1.1x10 <sup>3</sup> - 6.8x10 <sup>3</sup>	0.0076
PREVOTELLA	11/26	0-1.2x10 <sup>3</sup>	1.2x10 <sup>3</sup> - 2.4x10 <sup>3</sup>	0.0076
FUSOBACTERIUM	14/26	0-9.8x10 <sup>2</sup>	4.8x10 <sup>3</sup> - 3.6x10 <sup>4</sup>	0.0076
ORIBACTERIUM	8/26	0-0	1.7x10 <sup>3</sup> - 6.9x10 <sup>3</sup>	0.0093
PORPHYROMONAS	13/26	0-1.4x10 <sup>3</sup>	3.0x10 <sup>3</sup> - 2.2x10 <sup>4</sup>	0.0096
AKKERMANSIA	9/26	0-0	0 - 1.0x10 <sup>4</sup>	0.0171
FACALIBACTERIUM	6/26	0-0	0 - 6.7x10 <sup>3</sup>	0.0155
ACTINOMYCES	16/26	0-5.3x10 <sup>3</sup>	5.1x10 <sup>3</sup> - 3.7 x10 <sup>4</sup>	0.0171
STAPHYLOCCUS	14/26	0-3.2x10 <sup>3</sup>	4.1x10 <sup>3</sup> - 1.7x10 <sup>4</sup>	0.0182
NEISSERIA	20/26	0-9.9x10 <sup>3</sup>	9.0x10 <sup>3</sup> - 7.6x10 <sup>4</sup>	0.0182
SELENOMONAS	9/26	0-0	0 - 9.7x10 <sup>3</sup>	0.0199
LACTOBACILLUS	6/26	0-0	0 - 8.0x10 <sup>3</sup>	0.0223
PREVOTELLA_7	22/26	2.7x10 <sup>3</sup> - 1.4x10 <sup>3</sup>	1.2 x10 <sup>4</sup> - 7.4 x10 <sup>4</sup>	0.0287

**Table S5: Most abundant bacterial genera in pleural effusions of non-transplant patients (n=21).**

BACTERIA GENUS	MEDIAN READS
VEILLONELLA	18793
STREPTOCOCCUS	17322
PREVOTELLA_7	14252
HAEMOPHILUS	13592
NEISSERIA	12763
ROTHIA	10268
LEPTOTRICHIA	8048
STAPHYLOCOCCUS	7809
PREVOTELLA	5887
ACTINOMYCES	5810
CORYNEBACTERIUM	4494
FUSOBACTERIUM	3186
MEGASPHAERA	2858
PORPHYROMONAS	2441
GEMELLA	2071

**Table S6: Differential pleural abundancies of bacterial genera in late-stage effusions of patients after lung transplantation (n=19) and pleural effusions of patients without transplantation (n=21).**

BACTERIA GENUS	SAMPLES WITH DETECTED GENUS	COUNTS	COUNTS	FDR
		NON-TRANSPLANT [IQR]	LUNG TRANSPLANT [IQR]	
LEPTOTRICHIA	32	2.5x10 <sup>3</sup> -3.4x10 <sup>3</sup>	1.3x10 <sup>4</sup> -6.8x10 <sup>4</sup>	0.0563
PORPHYROMONAS	18	0-5.5x10 <sup>3</sup>	3.0x10 <sup>3</sup> -2.2x10 <sup>4</sup>	0.0563
ORIBACTERIUM	10	0-1.3 x10 <sup>3</sup>	1.7x10 <sup>3</sup> -6.9x10 <sup>3</sup>	0.0563
STREPTOCOCCUS	36	8.5x10 <sup>3</sup> -3.3x10 <sup>4</sup>	2.5x10 <sup>4</sup> -1.2x10 <sup>5</sup>	0.0563
LACTOBACILLUS	8	0-0	0-8.0x10 <sup>3</sup>	0.0563
ACTINOMYCES	25	2.8x10 <sup>3</sup> -1.1x10 <sup>4</sup>	5.1x10 <sup>3</sup> -3.7x10 <sup>4</sup>	0.0700
ALLOPREVOTELLA	19	0-1.1x10 <sup>4</sup>	4.6x10 <sup>3</sup> -1.9x10 <sup>4</sup>	0.0700
VEILLONELLA	36	1.1x10 <sup>4</sup> -4.7x10 <sup>4</sup>	1.7x10 <sup>4</sup> -1.0x10 <sup>5</sup>	0.0700
HAEMOPHILUS	35	5.9x10 <sup>3</sup> -2.0x10 <sup>4</sup>	1.2x10 <sup>4</sup> -8.0x10 <sup>4</sup>	0.0744
FUSOBACTERIUM	21	0-7.8x10 <sup>3</sup>	4.8x10 <sup>3</sup> -3.6x10 <sup>4</sup>	0.0744
LACHNOANAEROBACULUM	13	0-4.5x10 <sup>3</sup>	1.7x10 <sup>3</sup> -9.7x10 <sup>3</sup>	0.0785
ROTHIA	33	6.4x10 <sup>3</sup> -2.0x10 <sup>4</sup>	1.4x10 <sup>4</sup> -6.4x10 <sup>4</sup>	0.0917
AKKERMANSIA	14	0-3.8x10 <sup>3</sup>	0-1.0x10 <sup>4</sup>	0.1337
PREVOTELLA_7	33	9.2x10 <sup>3</sup> -4.4x10 <sup>4</sup>	1.2x10 <sup>4</sup> -7.4x10 <sup>4</sup>	0.1457
GEMELLA	9	0-3.5x10 <sup>3</sup>	2.0x10 <sup>3</sup> -7.7x10 <sup>3</sup>	0.1457
CUTIBACTERIUM	11	0-3.7x10 <sup>3</sup>	0-6.1x10 <sup>3</sup>	0.1925
NEISSERIA	35	6.8x10 <sup>3</sup> -2.6x10 <sup>4</sup>	9.0x10 <sup>3</sup> -7.6x10 <sup>4</sup>	0.1925
PREVOTELLA	23	0-9.7x10 <sup>3</sup>	2.4x10 <sup>3</sup> -2.2x10 <sup>4</sup>	0.1925
SELENOMONAS	15	0-5.7x10 <sup>3</sup>	0-9.7x10 <sup>3</sup>	0.1925
CORYNEBACTERIUM	18	1.5x10 <sup>2</sup> -7.8x10 <sup>3</sup>	4.0x10 <sup>3</sup> -1.0x10 <sup>4</sup>	0.2062

**Table S7: Differential pleural abundances of bacterial genera in patients with (n=24) and without (n=28) previous thoracocentesis.**

BACTERIA GENUS	SAMPLES WITH DETECTED GENUS	COUNTS WITHOUT PREVIOUS THORACOCENTESIS [IQR]	COUNTS WITH PREVIOUS THORACOCENTESIS [IQR]	FDR
PREVOTELLA	25	0-4.7x10 <sup>3</sup>	5.3x10 <sup>3</sup> -2.3x10 <sup>4</sup>	0.0105
ROTHIA	40	2.4x10 <sup>3</sup> -1.7x10 <sup>4</sup>	1.1x10 <sup>4</sup> -7.1x10 <sup>4</sup>	0.0121
GEMELLA	11	0-2.1x10 <sup>3</sup>	1.8x10 <sup>3</sup> -7.2x10 <sup>3</sup>	0.0121
NEISSERIA	42	3.4x10 <sup>3</sup> -1.5x10 <sup>4</sup>	1.0x10 <sup>4</sup> -7.6x10 <sup>4</sup>	0.0121
STREPTOCOCCUS	43	5.6x10 <sup>3</sup> -2.5x10 <sup>4</sup>	1.7x10 <sup>4</sup> -1.1x10 <sup>5</sup>	0.0145
VEILLONELLA	42	3.9x10 <sup>3</sup> -2.2x10 <sup>4</sup>	1.4x10 <sup>4</sup> -1.0x10 <sup>5</sup>	0.0203
CORYNEBACTERIUM	23	0-6.2x10 <sup>3</sup>	3.9x10 <sup>3</sup> -1.5x10 <sup>4</sup>	0.0484
FUSOBACTERIUM	24	0-6.9x10 <sup>3</sup>	2.6x10 <sup>3</sup> -3.7x10 <sup>4</sup>	0.0484
HAEMOPHILUS	42	4.6x10 <sup>3</sup> -1.9x10 <sup>4</sup>	9.7x10 <sup>3</sup> -7.9x10 <sup>4</sup>	0.0484
PREVOTELLA_7	40	4.1x10 <sup>3</sup> -2.2x10 <sup>4</sup>	9.9x10 <sup>3</sup> -7.0x10 <sup>4</sup>	0.0558
PORPHYROMONAS	21	0-5.2x10 <sup>3</sup>	1.7x10 <sup>3</sup> -1.6x10 <sup>4</sup>	0.0679
ACTINOMYCES	30	0-1.1x10 <sup>4</sup>	3.8x10 <sup>3</sup> -3.5x10 <sup>4</sup>	0.0679
LEPTOTRICHIA	38	2.5x10 <sup>3</sup> -1.9x10 <sup>4</sup>	7.0x10 <sup>3</sup> -7.7x10 <sup>4</sup>	0.0698
ALLOPREVOTELLA	21	0-6.8x10 <sup>3</sup>	4.1x10 <sup>2</sup> -2.0x10 <sup>4</sup>	0.0708
CUTIBACTERIUM	15	0-2.9x10 <sup>3</sup>	0-1.2x10 <sup>4</sup>	0.0759
SELENOMONAS	17	0-3.4x10 <sup>3</sup>	0-9.7x10 <sup>3</sup>	0.0759
LACHNOANEROBACULUM	16	0-3.7x10 <sup>3</sup>	0-8.9x10 <sup>3</sup>	0.0861
MEGASPHAERA	15	0-3.4x10 <sup>3</sup>	0-9.0x10 <sup>3</sup>	0.1578
ORIBACTERIUM	12	0-2.6x10 <sup>3</sup>	0-5.9x10 <sup>3</sup>	0.2257
STAPHYLOCOCCUS	30	1.9x10 <sup>3</sup> -1.6x10 <sup>4</sup>	4.3x10 <sup>3</sup> -1.7x10 <sup>4</sup>	0.2363