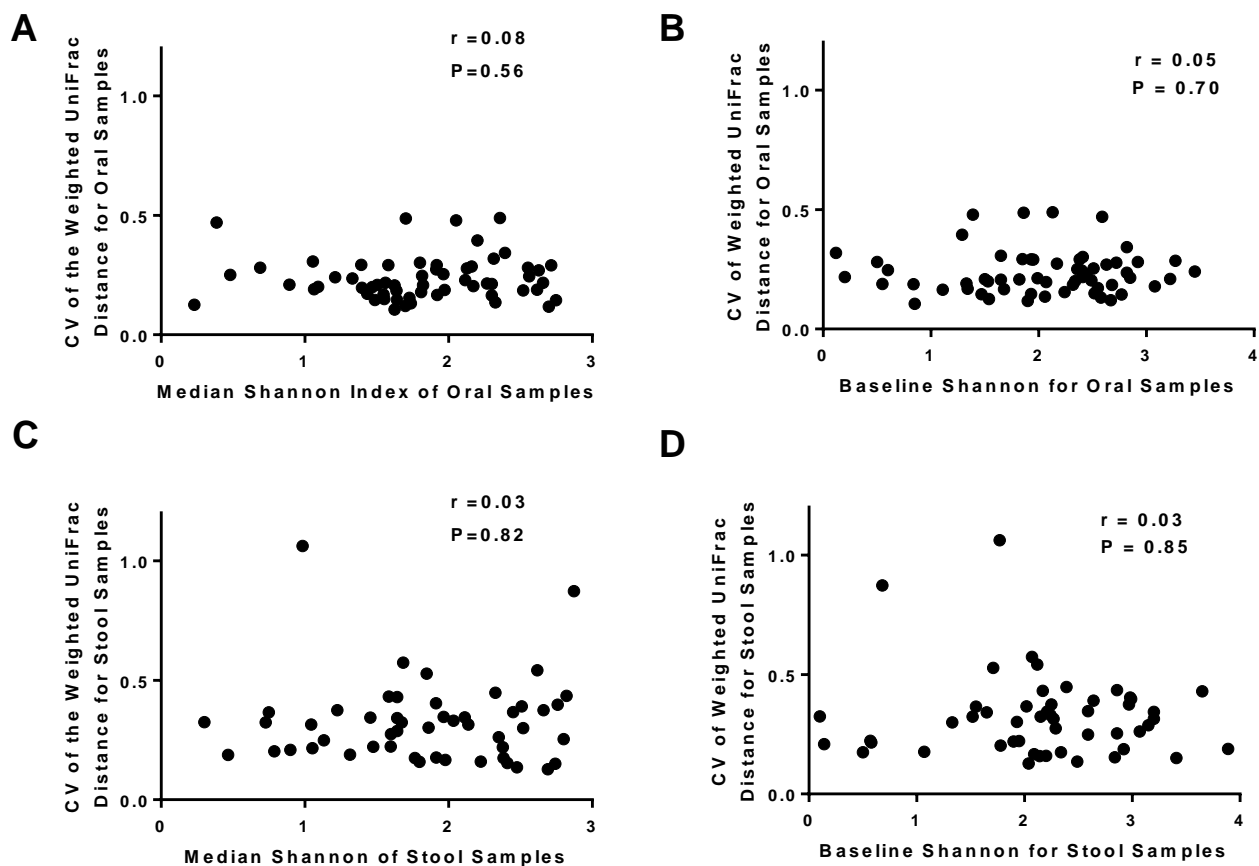


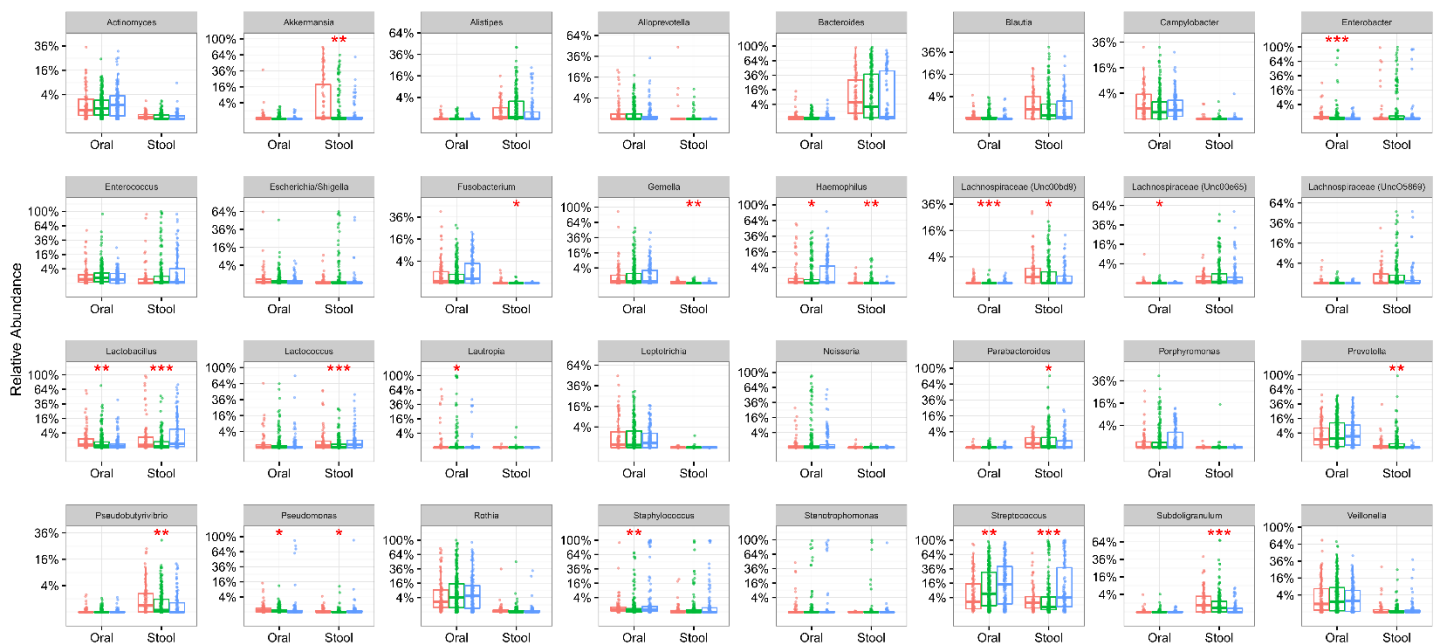
**Fig. S1. The temporal variability of the oral and stool microbial  $\alpha$ -diversity of the AML and HMP cohorts.** (A) Depicted is the oral and stool microbial  $\alpha$ -diversity intra-patient temporal variability. Each point represents the coefficient of variation (CV) of the Chao-1 diversity index for samples derived from each patient with  $\geq 3$  samples. (B) Same as (A) except using the Simpson index. (C) Depicted is the oral and stool microbial  $\alpha$ -diversity intra-patient temporal variability. Each point represents the CV of the Shannon diversity index for samples derived from the HMP cohort. In panels A -C, the bars represent mean  $\pm$  standard deviation values. (D) Portrayed is the correlation between the CV of the Weighted UniFrac Distances originating from oral and stool samples from the same patient. The Pearson's correlation ( $r$ ) value and  $P$  value for correlation analysis also are indicated.



**Fig. S2. No correlation between temporal stability of community structure and bacterial diversity found among treated acute myeloid leukemia patients.** Portrayed is the correlation between the coefficient of variation of the weighted UniFrac distances from oral (A and B) and stool (C and D) samples from each patient and either the median Shannon index of those samples (A and C), or a patient's baseline Shannon diversity (B and D). The Pearson's correlation ( $r$ ) value and  $P$  value for correlation analysis also are indicated.

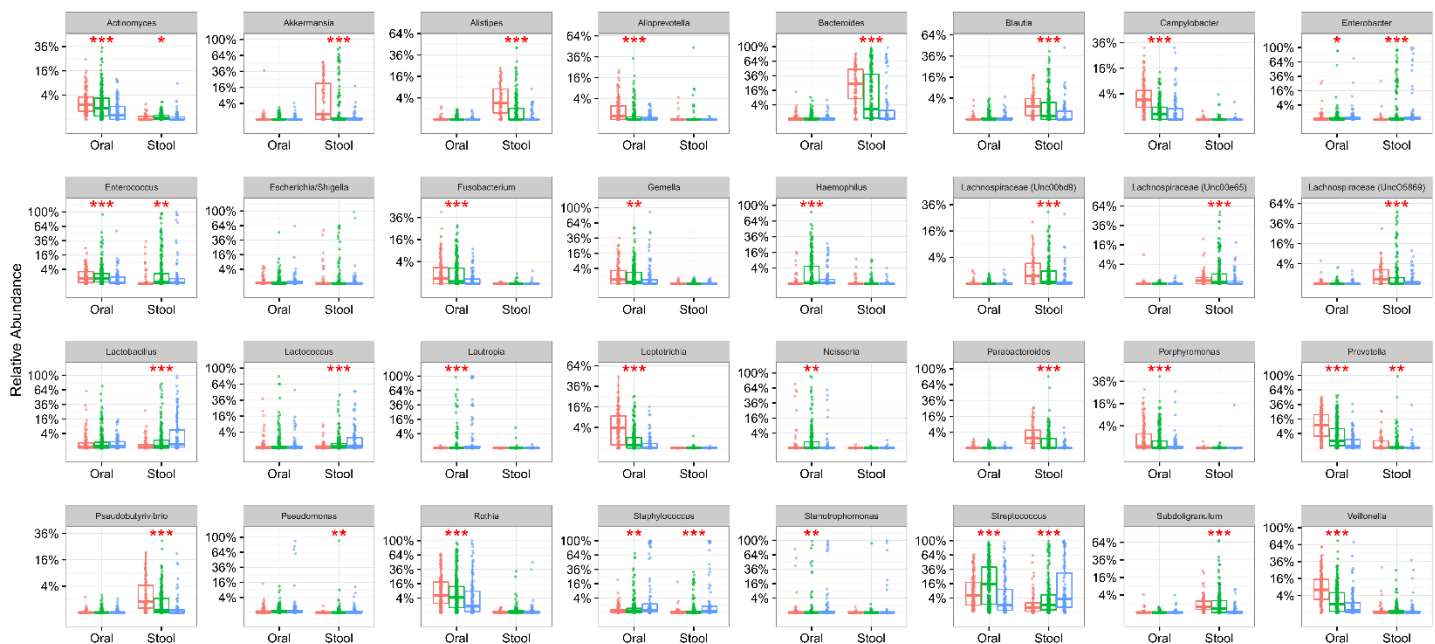
### Taxa by Patient Microbiome Stability

Patient Microbiome Stability (Coefficient of Variation of Weighted UniFrac Distance): ■ Stable ■ Average ■ Variable



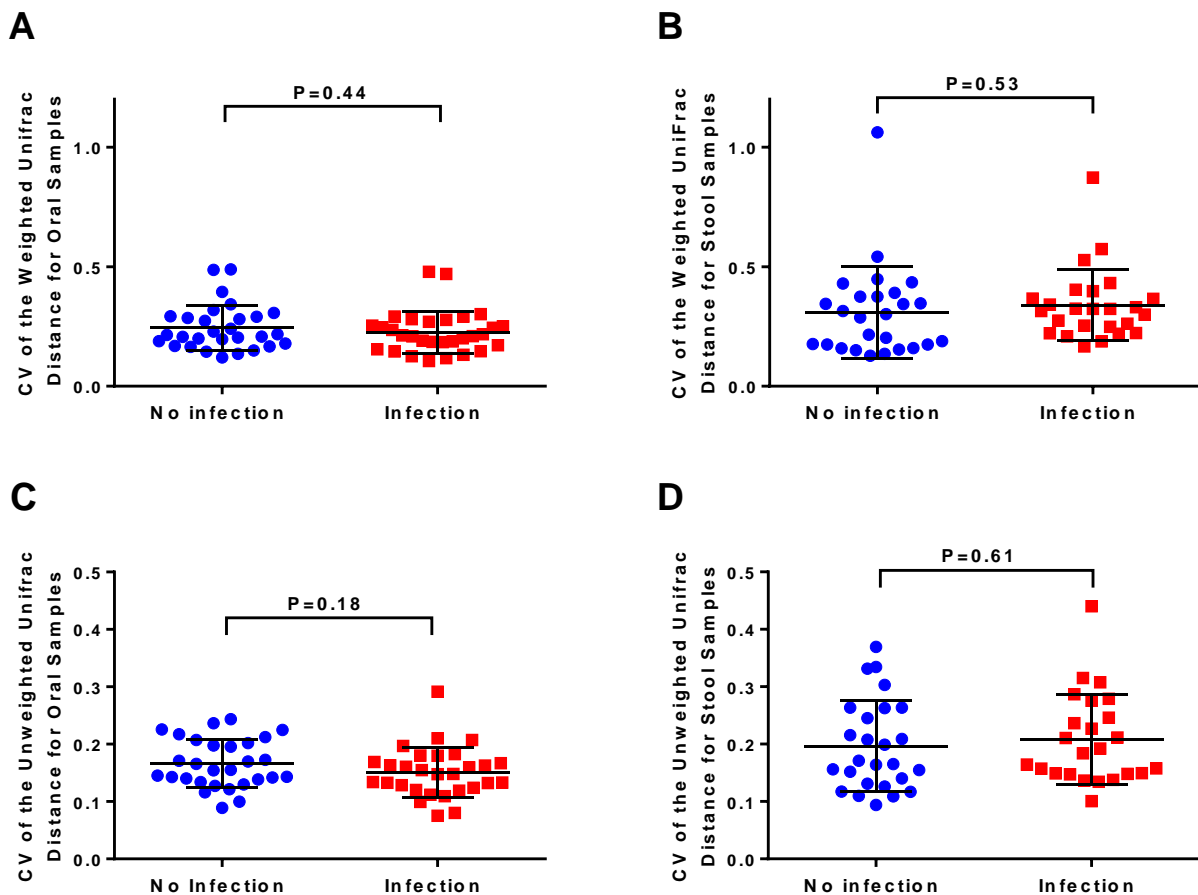
### Taxa by Patient Microbiome Diversity

Patient Microbiome Diversity (Coefficient of Variation of Shannon Diversity Index):



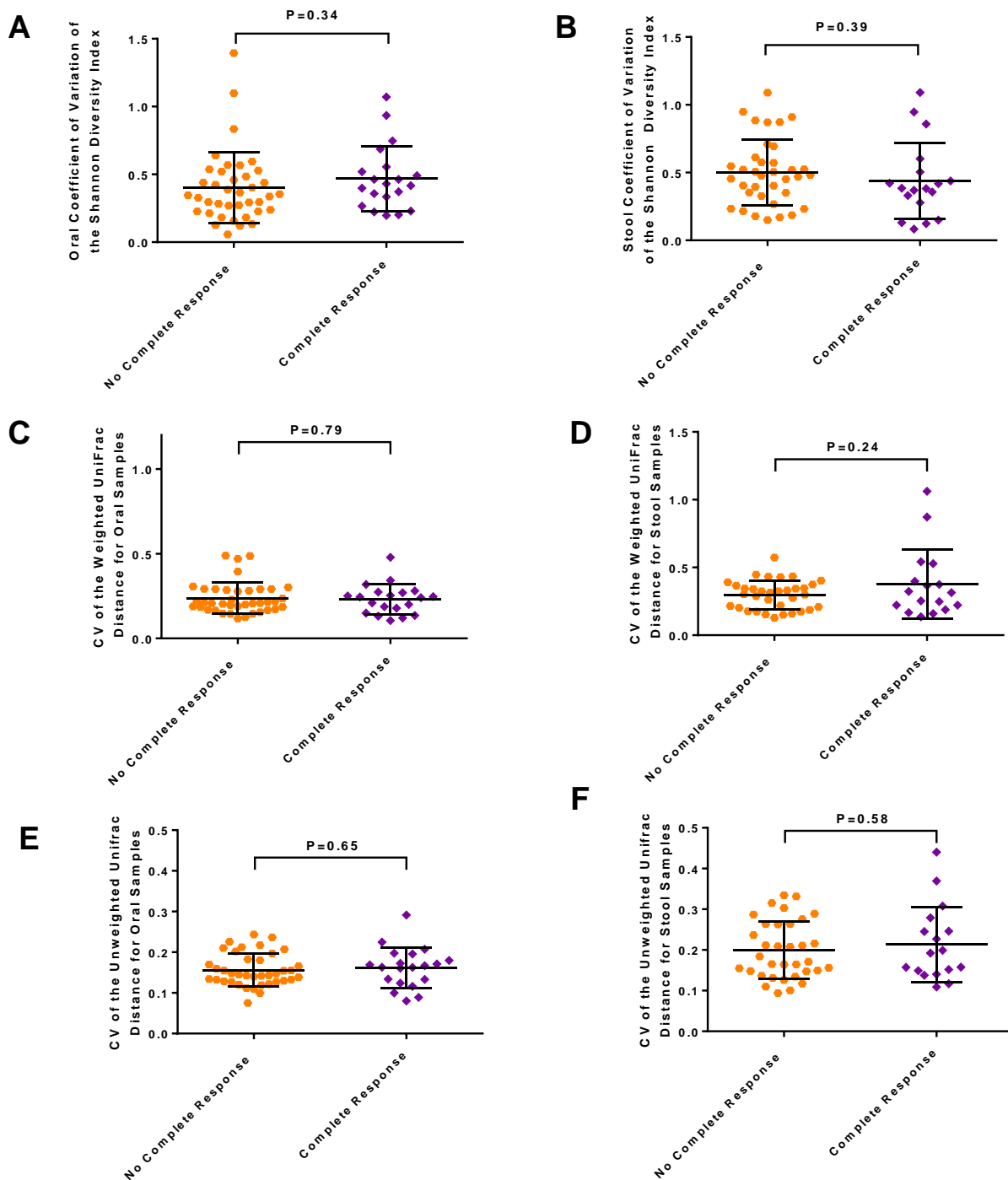
**Fig. S3. Taxonomic composition differences among different stability categories.** Shown are the significant differences in relative abundances of genera between different patient stability categories based on either coefficient of variation (CV) of weighted UniFrac distances (top panels) or the CV of the Shannon diversity index (SDI) (bottom panels). Differences in genera abundance across categories were determined using non-parametric Kruskal-Wallis analysis of variance, then corrected for the false discovery rate using the Benjamini & Hochberg method. Asterisk notations \*, \*\*, and \*\*\* indicate adjusted  $P$  values  $\leq 0.05$ ,  $0.01$ , and  $0.001$ , respectively.

Fig. S4



**Fig. S4. Temporal instability of microbiome community structure is not associated with infection during chemotherapy.** Shown are the CVs of the weighted (A and B) and unweighted (C and D) unifracs distances for oral (A and C) and stool (B and D) samples stratified by patients who did or did not contract an infection during the induction phase of chemotherapy before neutrophil recovery. In all panels the bars represent mean  $\pm$  standard deviation values, and  $P$  values comparing CV values among infectious outcomes were calculated using a 2-sample t-test with Welch's correction.

Fig. S5



**Fig. S5. Temporal variability of the microbiome is not associated with chemotherapeutic response.** Shown are the CVs of the Shannon diversity index (A and B), weighted (C and D), and unweighted (E and F) unifracs distances for oral (A, C, E) and stool (B, D, F) samples stratified by patients who did or did not have a complete response to induction chemotherapy. In all panels the bars represent mean  $\pm$  standard deviation values, and  $P$  values comparing CV values among infectious outcomes were calculated using a 2-sample t-test with Welch's correction.

**Table S1. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of the stool as measured by the Coefficient of Variation of the Shannon Diversity Index<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	-0.0030511	0.0037892	-0.805	0.425
Received Piperacillin/ Tazobactam >72hr	-0.0692387	0.1034310	-0.669	0.507
Received Cefepime >72hrs	-0.0642391	0.0961470	-0.668	0.508
Received Carbapenem >72hrs	0.0910213	0.0984667	0.924	0.360
Days on All Antibiotics <sup>d</sup>	0.0028492	0.0032929	0.865	0.392
Days on Treatment Antibiotics	-0.0004718	0.0045104	-0.105	0.917
Number of Antibiotics Received	0.0213972	0.0229633	0.932	0.357
Non-Fludarabine High Intensity Chemotherapy	0.0771821	0.0937103	0.824	0.415
Hypomethylator based Chemotherapy	0.1747187	0.1478379	1.182	0.244
ChemoOthers	0.0870571	0.1618069	0.538	0.593

<sup>1</sup> Residual standard error: 0.2507 on 43 degrees of freedom, adjusted R-squared: 0.0397 and p-value: 0.3064.

**Table S2. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of the stool as measured by the Coefficient of Variation of the Unweighted Unifrac Distance<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	-0.0003602	0.0021741	-0.283	0.7789
Received Piperacillin/ Tazobactam >72hr	0.0436241	0.0346360	1.260	0.2151
Received Cefepime >72hrs	0.0139268	0.0324944	0.430	0.6697
Received Carbapenem >72hrs	-0.0231621	0.0326590	-0.709	0.4823
Days on All Antibiotics <sup>d</sup>	0.0001982	0.0010993	0.180	0.8579
Days on Treatment Antibiotics	-0.0007410	0.0015409	-0.481	0.6332
Number of Antibiotics Received	0.0027624	0.0077760	0.355	0.7243
Non-Fludarabine High Intensity Chemotherapy	0.0165690	0.0317881	0.521	0.6051
Hypomethylator based Chemotherapy	-0.0062450	0.0493546	-0.127	0.8999
ChemoOthers	0.0390815	0.0533204	0.733	0.4679

<sup>1</sup> Residual standard error: 0.08237 on 40 degrees of freedom, adjusted R-squared: -0.1106, and p-value: 0.8784.

**Table S3. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of the stool as measured by the Coefficient of Variation of the Weighted Unifrac Distance<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	0.0004108	0.0028111	0.149	0.885
Received Piperacillin/ Tazobactam >72hr	-0.0025742	0.0764188	-0.034	0.973
Received Cefepime >72hrs	-0.0338694	0.0716937	-0.472	0.639
Received Carbapenem >72hrs	-0.0298034	0.0720568	-0.414	0.681
Days on All Antibiotics <sup>d</sup>	0.0013529	0.0024254	0.558	0.580
Days on Treatment Antibiotics	-0.0025904	0.0033998	-0.762	0.451
Number of Antibiotics Received	0.0241723	0.0171565	1.409	0.167
Non-Fludarabine High Intensity Chemotherapy	0.0527559	0.0701354	0.752	0.456
Hypomethylator based Chemotherapy	-0.0556192	0.1088930	-0.511	0.612
ChemoOthers	0.0886147	0.1176429	0.753	0.456

<sup>1</sup> Residual standard error: 0.1817 on 40 degrees of freedom, adjusted R-squared: -0.1163, and p-value: 0.8937



**Table S4. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of oral samples as measured by the Coefficient of Variation of the Shannon Diversity Index<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	0.003785	0.003515	1.077	0.2870
Received Piperacillin/ Tazobactam >72hr	-0.069843	0.097154	-0.719	0.4757
Received Cefepime >72hrs	-0.142693	0.087163	-1.637	0.1082
Received Carbapenem >72hrs	0.029032	0.089903	0.323	0.7482
Days on All Antibiotics <sup>d</sup>	-0.006798	0.003052	-2.227	0.0306
Days on Treatment Antibiotics	0.005466	0.004250	1.286	0.2046
Number of Antibiotics Received	0.038113	0.021953	1.736	0.0890
Non-Fludarabine High Intensity Chemotherapy	0.031210	0.087383	0.357	0.7225
Hypomethylator based Chemotherapy	-0.048031	0.140382	-0.342	0.7337
ChemoOthers	0.048609	0.140919	0.345	0.7316

<sup>1</sup> Residual standard error: 0.2429 on 48 degrees of freedom, adjusted R-squared: 0.08042, and p-value: 0.1661.

**Table S5. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of oral samples as measured by the Coefficient of Variation of the Unweighted Unifrac Distance<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	9.131e-01	5.338e-04	0.171	0.864908
Received Piperacillin/ Tazobactam >72hr	-1.883e-02	1.475e-02	-1.276	0.207937
Received Cefepime >72hrs	7.827e-03	1.324e-02	0.591	0.557092
Received Carbapenem >72hrs	-4.519e-03	1.365e-02	-0.305	0.761985
Days on All Antibiotics <sup>d</sup>	1.912e-03	4.635e-04	4.125	0.000147
Days on Treatment Antibiotics	-1.239e-03	6.445e-04	-1.920	0.060774
Number of Antibiotics Received	-2.951e-03	3.334e-03	-0.885	0.380505
Non-Fludarabine High Intensity Chemotherapy	-7.372e-03	1.327e-02	-0.556	0.581112
Hypomethylator based Chemotherapy	7.928e-03	2.132e-02	0.372	0.711619
ChemoOthers	-2.641e-02	2.140e-02	-1.234	0.223152

<sup>1</sup> Residual standard error: 0.03689 on 48 degrees of freedom, adjusted R-squared: 0.2682, and p-value: 0.003797

**Table S6. Multivariable regression analyses of potential clinical factors associated with the intra-patient temporal instability of oral samples as measured by the Coefficient of Variation of the Weighted Unifrac Distance<sup>1</sup>**

	Estimate	Std. Error	t value	P value
Age	-0.0007342	0.0011572	-0.634	0.52877
Received Piperacillin/ Tazobactam >72hr	-0.0439871	0.0319822	-1.375	0.17540
Received Cefepime >72hrs	-0.0020750	0.0286934	-0.072	0.94265
Received Carbapenem >72hrs	-0.0063201	0.0295955	-0.214	0.83180
Days on All Antibiotics <sup>d</sup>	0.0032774	0.0010047	3.262	0.00204
Days on Treatment Antibiotics	-0.0019812	0.0013992	-1.416	0.16326
Number of Antibiotics Received	-0.0011496	0.0072269	-0.159	0.87428
Non-Fludarabine High Intensity Chemotherapy	-0.0023525	0.0287658	-0.082	0.93516
Hypomethylator based Chemotherapy	0.0627605	0.0462125	1.358	0.18079
ChemoOthers	0.0362807	0.0463895	0.782	0.43800

<sup>1</sup> Residual standard error: 0.07998 on 48 degrees of freedom, adjusted R-squared: 0.2218, and p-value: 0.01157

Table S7. Biosample accession numbers

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