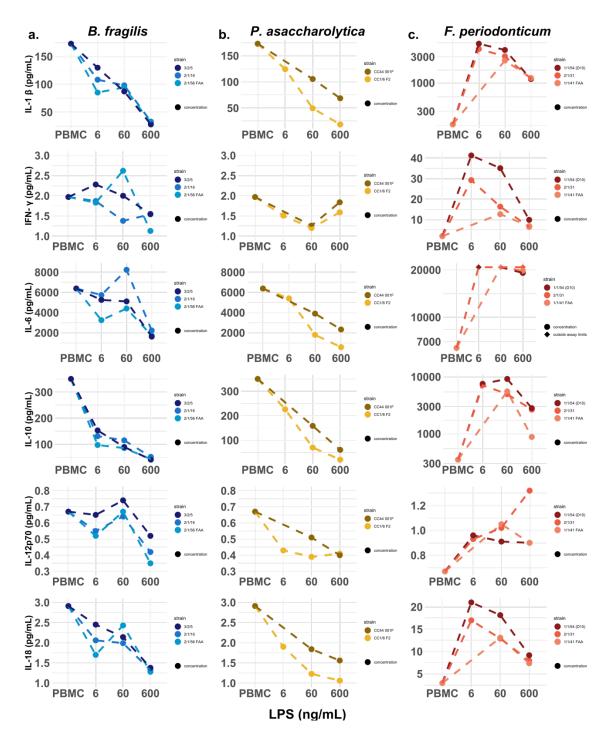
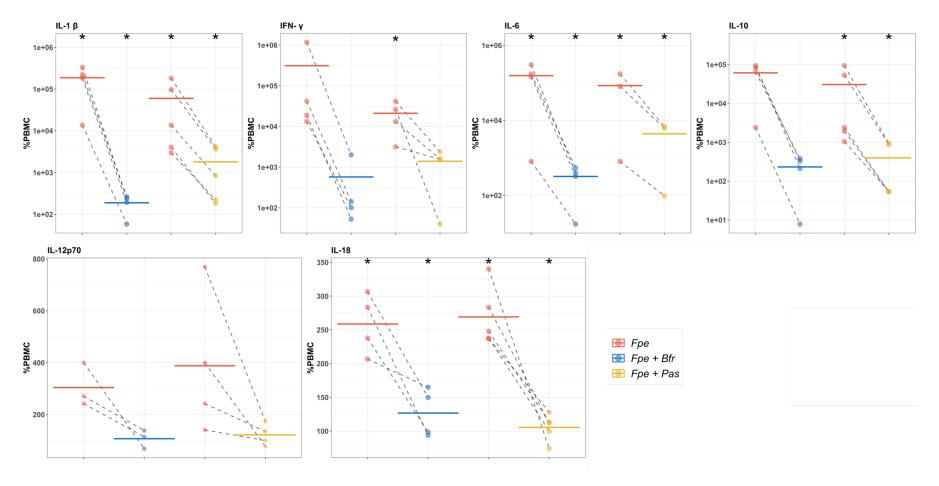
## **SUPPLEMENTARY MATERIAL**

## **SUPPLEMENTARY FIGURES**



Supplementary Figure 1. Secreted cytokine concentrations (pg/mL) after overnight incubation of PBMCs with different LPS concentrations (6 ng/mL, 60 ng/mL, 600 ng/mL) from different strains of a. *B. fragilis* b. *F. periodonticum*, and c. *P. asaccharolytica* compared to PBMC baseline (no treatment). Dashed lines connect data points from the same strains. Colors differentiate species (*B. fragilis:* blue; *P. asaccharolytica*: yellow; *F. periodonticum:* red) and different shades of a color differentiate the strains of each species. Where values are outside of assay limits, we show the assay limit, and denote the point with ' $\spadesuit$ '. For *F. periodonticum* plots, IL-1 $\beta$ , IL-6, and IL-10 y-axes are at scale log<sub>10</sub>.



Supplementary Figure 2. Changes in cytokine expression in peripheral blood mononuclear cells (PBMCs) following treatment with *F. periodonticum* alone (red) or in combination with *B. fragilis* (blue) or *P. asaccharolytica* (yellow). We show results for cytokines of interest IL-1 $\beta$ , IFN- $\gamma$ , IL-6, IL-10, IL-12p70, and IL-18. *F. periodonticum* 2/1/31 LPS used was at a concentration of 6 ng/mL. *B. fragilis* 2/1/16 LPS and *P. asaccharolytica* CC1/6 F2 LPS used were at a concentration of 600ng/mL. Values are shown as percentages of PBMC baseline secretion, which is set at 100%. Dashed lines indicate a single experimental run. Colored, solid horizontal lines represent the means of repeat experiments. Y-axes of IL-1 $\beta$ , IFN- $\gamma$ , IL-6, and IL-10 are in log<sub>10</sub> scale, while Y-axes of IL-12p70 and IL-18 are in linear scale. *Fpe* = *F. periodonticum* (6ng/mL), *Fpe* + *Bfr* = *F. periodonticum* (6ng/mL) + *P. asaccharolytica* (600 ng/mL); (\*) = *Paired student's t-tests p-value* < 0.05 (*F. periodonticum vs PBMC*, *F. periodonticum* + *B. fragilis vs F. periodonticum* alone, or *F. periodonticum* + *P. asaccharolytica vs F.* 

periodonticum alone).

## **SUPPLEMENTARY TABLES**

	Minimum	Maximum	Median	
Raw Reads	3518434	18689703	12231622.5	
Human Analysis (STAR	alignment) <sup>1</sup> - Pa	nired End	•	
Input	3479791	18670112	12152988	
Aligned Uniquely (%)	17.04	93.82	85.84	
Human Analysis (STAR	alignment) <sup>1</sup> - Si	ngle End	<u>.</u>	
Input	35	5402	2875	
Aligned Uniquely (%)	3.7	92.49	84.15	
Microbial Analysis <sup>1</sup>	•		•	
Input (to Kaiju)	53909	4711734	641559	
Identified Species <sup>2</sup> (%)	12.2	65.97	19.02	

<sup>&</sup>lt;sup>1</sup> The trimming and quality control steps provide paired and single end reads which were used to align to the human genome using STAR, separately. It is only after featureCounts that their respective gene counts were combined for subsequent analysis. For microbial analysis, the same was carried out, however Kaiju recommends combining results of paired-end (PE) and single-end (SE) reads immediately, allowing for combined read count information of the PE and SE reads.

<sup>&</sup>lt;sup>2</sup> Identified species that passed the relative abundance (in percent) cutoff of 0.01%.

Supplementary Table 2. Test of Differences between Cytokine Concentrations Released after PBMC Treatment using <i>F. periodonticum and B. fragilis</i> LPS									
Cytokine	Reference Group	Group 2	N1	N2	р	Cohen's d	Magnitude		
IL-12p70	PBMC	Fp6	3	3	0.16	-0.72	moderate		
	Fp6	Fp6 + Bf600	3	3	0.20	0.62	moderate		
	•	•							
IL-10	PBMC	Fp6	4	4	0.057	-1.10	large		
	Fp6	Fp6+Bf600	4	4	0.055	1.11	large		
IFN-γ	PBMC	Fp6	4	4	0.09	-0.90	large		
	Fp6	Fp6+Bf600	4	4	0.09	0.90	large		
IL-1β	PBMC	Fp6	4	4	0.041	-1.26	large		
	Fp6	Fp6+Bf600	4	4	0.041	1.26	large		
					<b>.</b>	<del>,</del>	,		
IL-18	PBMC	Fp6	4	4	0.017	-1.77	large		
	Fp6	Fp6+Bf600	4	4	0.05	1.163	large		
	1			r		<u> </u>			
IL-6	PBMC	Fp6	4	4	0.025	-1.52	large		
	Fp6	Fp6+Bf600	4	4	0.023	1.57	large		

Legend: Fp6=F. periodonticum 2/1/31 (6 ng/mL) alone; Fp6+Bf600 =F. periodonticum 2/1/31 (6 ng/mL) + B. fragilis 2/1/16 (600 ng/mL) co-incubation; N=number of repeats used in statistical calculations; p=paired t-test p-value; d=cohen's d; Magnitude = interpretation of cohen's d

Supplementary Table 3. Test of Differences between Cytokine Concentrations Released after PBMC Treatment using <i>F. periodonticum</i> and <i>P. asaccharolytica</i> LPS								
Reference Group	Group 2	N1	N2	р	Cohen's d	Magnitude		
PBMC	Fp6	4	4	0.06	-1.07	large		
Fp6	Fp6+Pa600	4	4	0.08	0.95	large		
PBMC	Fp6	5	5	0.017	-1.41	large		
Fp6	Fp6+Pa600	5	5	0.016	1.45	large		
PBMC	Fp6	4	4	0.035	-1.34	large		
Fp6	Fp6+Pa600	4	4	0.06	1.07	large		
PBMC	Fp6	5	5	0.010	-1.63	large		
Fp6	Fp6+Pa600	5	5	0.011	1.61	large		
PBMC	Fp6	5	5	0.02	-1.35	large		
Fp6	Fp6+Pa600	5	5	0.021	1.33	large		
PBMC	Fp6	3	3	0.017	-2.52	large		
Fp6	Fp6+Pa600	3	3	0.015	2.69	large		
	Reference Group PBMC Fp6	tment using F. periodonticum and P.           Reference Group         Group 2           PBMC         Fp6           Fp6         Fp6+Pa600           PBMC         Fp6           Fp6         Fp6+Pa600	tment using F. periodonticum and P. asac           Reference Group         Group 2         N1           PBMC         Fp6         4           Fp6         Fp6+Pa600         4           PBMC         Fp6         5           Fp6         Fp6+Pa600         5           PBMC         Fp6         4           Fp6         Fp6+Pa600         4           PBMC         Fp6         5           Fp6         Fp6+Pa600         5           PBMC         Fp6         5           Fp6         Fp6+Pa600         5           PBMC         Fp6         5           Fp6         Fp6+Pa600         5	tment using F. periodonticum and P. asacchard           Reference Group         Group 2         N1         N2           PBMC         Fp6         4         4           Fp6         Fp6+Pa600         4         4           PBMC         Fp6         5         5           Fp6         Fp6+Pa600         5         5           PBMC         Fp6         4         4           PBMC         Fp6+Pa600         4         4           PBMC         Fp6+Pa600         5         5           PBMC         Fp6         5         5           Fp6         Fp6+Pa600         5         5           PBMC         Fp6         5         5           Fp6         Fp6+Pa600         5         5           Fp6         Fp6+Pa600         5         5           PBMC         Fp6+Pa600         5         5	tment using F. periodonticum and P. asaccharolytica LP           Reference Group         Group 2         N1         N2         p           PBMC         Fp6         4         4         0.06           Fp6         Fp6+Pa600         4         4         0.08           PBMC         Fp6         5         5         0.017           Fp6         Fp6+Pa600         5         5         0.016           PBMC         Fp6         4         4         0.035           Fp6         Fp6+Pa600         4         4         0.06           PBMC         Fp6         5         5         0.010           Fp6         Fp6+Pa600         5         5         0.011           PBMC         Fp6         5         5         0.021           PBMC         Fp6+Pa600         5         5         0.021           PBMC         Fp6         3         3         0.017	Reference Group         Group 2         N1         N2         p         Cohen's d           PBMC         Fp6         4         4         0.06         -1.07           Fp6         Fp6+Pa600         4         4         0.08         0.95           PBMC         Fp6         5         5         0.017         -1.41           Fp6         Fp6+Pa600         5         5         0.016         1.45           PBMC         Fp6         4         4         0.035         -1.34           Fp6         Fp6+Pa600         4         4         0.06         1.07           PBMC         Fp6         5         5         0.010         -1.63           Fp6         Fp6+Pa600         5         5         0.011         1.61           PBMC         Fp6         5         5         0.021         -1.35           Fp6         Fp6+Pa600         5         5         0.021         1.33           PBMC         Fp6+Pa600         5         5         0.021         1.33           PBMC         Fp6         3         3         0.017         -2.52		

Legend: Fp6=*F. periodonticum 2/1/31* (6 ng/mL) alone; Fp6+Pa600=*F. periodonticum 2/1/31* (6 ng/mL) + *P. asaccharolytica CC1/6 F2*(600 ng/mL) co-incubation; N=number of repeats used in statistical calculations; p=paired t-test p-value; d=cohen's d; Magnitude = interpretation of cohen's d