Supplementary Information

Figures



Supplementary Figure 1: Microglia activation in limbic brain regions during the CUS protocol.

Microglia activation was assessed as the percentage of surface area covered by the activation marker CD68 compared to the total area stained with Iba1 (brown columns, left axis). Microglia count in each frame is also displayed (blue line, right axis). Measures were taken at all timepoints including US, CUS, CUS+Rest and CUS+US. Significance (n = 3, +/- SEM) was determined with a one-way ANOVA and Tukey's post-hoc analyses and marked as *p < 0.05, **p < 0.01, ***p < 0.0001.



Supplementary Figure 2: Gut microbiota altering substances modulate gene and cytokine expression of immunological factors during chronic and recurrent stress.

Gene and cytokine expression in the prefrontal cortex (PFC), hippocampus, cortex and cerebellum is outlined as the z-score of relative expression normalized across all timepoints for a



particular gene or cytokine. Note that each instance is a ratio of stressed vs. non-stressed for each of the respective treatment group. For each group, n = 6 individuals are shown.

Supplementary Figure 3: Gut microbiome variations in response the CUS protocol. Alpha diversity (Fisher's alpha) and beta diversity (unweighted UniFrac PCoA plots) show the variations in the gut microbiome with respect to the CUS protocol within a treatment group for (a) CUS control vs. stressed, (b) Control, (c) BDPP, (d) Probiotic and (e) Synbiotic. (f) Enumeration of the main phyla across treatment at the respective timepoints is also shown. Each group contains n = 6 individuals +/- SEM with significance determined with one-way ANOVA *p<0.05 and **p<0.01.



Supplementary Figure 4: Gut microbiome genera variations with respect to time point during chronic and recurrent stress. 16S V4 sequencing of gut microbiome genera of stressed mice only during the CUS protocol are shown with variations with respect to timepoint during the CUS protocol for the respective treatments. Each group represents n = 6 individuals +/- SEM with significance determined with a one-way ANOVA and Tukey's post-hoc analysis where *p<0.05, **p<0.01 and ***p<0.001.



Supplementary Figure 5: Gut microbiome genera variations with respect to treatment during chronic and recurrent stress. 16S V4 sequencing of gut microbiome genera of stressed mice only during the CUS protocol are shown with variations with respect to treatment. Each group represents n = 6 individuals +/- SEM with significance determined with a one-way ANOVA and Tukey's post-hoc analysis where *p<0.05, **p<0.01 and ***p<0.001.



Supplementary Figure 6: Stress and synbiotic induced variations on the ratio of Th17 to Treg and their activated states. The ratios of Th17 to Treg cells was determined with (a) the ratio of transcription factors *Roryt* to *Foxp3*, (b) ratio of cytokines IL-17A to IL-10, (c) the ratio of unactivated Th17/Treg cells and (d) ratio of "activated" Th17 (IL-17A⁺ICOS⁺) and Treg (CTLA4⁺). For the gene and cytokine expression, n = 6 independent samples +/- SEM were used, while n = 3 individuals for the cellular data, while statistics were calculated with one-way ANOVAs in all cases with Tukey's post-hoc analysis *p<0.05, **p<0.01.



Supplementary Figure 7: Ileum additional cell population quantification. Relative cellular expression (as indicated on y-axis) or activation for various cell populations and their activation states are shown for (a) monocytes relative to total cell populations, (b) macrophages relative to total cell populations, (c) ratio of MHCII⁺ to MHCII⁻ cells, (d) dendritic cells (DC) relative to total cell populations, (e) DC1 and (f) DC2 relative to total DC cell population, (g) innate

lymphoid cells (ILCs) relative to total cell populations, (h) ILC1, (i) ILC2, (j) ILC3 IL-22⁺ and (k) ILC3 IL-17A⁺ relative to ILC populations, (l) NK cells relative to total cell populations, (m) B cells and (n) memory B cells relative total cell populations, (o) CD8 IFNya cells and (p) CD8 TNFa cells relative to CD8 cell populations and (q) Th1 and (r) Th2 cells relative to total cell populations in their naïve and activated states (solid superimposed bars). In all cases, n = 3 individuals +/- SEM are shown and significance indicated with a one-way ANOVA with Tukey's post-hoc analysis with *p<0.05, **p<0.01.



Supplementary Figure 8: Immune cell distribution in spleen and PBMCs. Major cell populations determined with CyTOF immunophenotyping in the (a) spleen and quantification of (b) genes and (c) cytokines in the spleen are shown as heatmaps with each instance being the ratio of stressed vs. non-stressed group within each treatment, with the z-score taken across time for each gene. Major cell populations from (d) PBMCs are represented in viSNE plots while

proinflammatory cytokine expression in the serum is shown as a heatmap in (e). For the cell frequencies, n = 3 independent samples +/- SEM are shown while for the gene and cytokine determination, n = 6 +/- SEM is shown.



Supplementary Figure 9: Spleen additional cell population quantification. Relative cellular expression (as indicated on y-axis) or activation for various cell populations and their activation states are shown for (a) monocytes relative to total cell populations, (b) macrophages relative to total cell populations, (c) dendritic cells (DC) relative to total cell populations, (d) innate lymphoid cells (ILCs) relative to total cell populations, (e) ILC1 and (f) ILC3 relative to total

cell populations, (g) regulatory T cells (Tregs) and (h) T helper (Th)17 cells, both with an inset of activated cell populations (Treg, CTLA4⁺ and Th17, IL17A⁺ICOS⁺) relative to total CD4+ cells, (i) NK cells relative to total cell populations, (j) B cells and (k) memory B cells relative total cell populations, (l) CD8 IFNy cells and (m) CD8 TNF α cells relative to total CD8 cell populations and (n) Th1 and (o) Th2 cells relative to total CD4+ cell populations in their naïve and activated states (solid superimposed bars). In all cases, n = 3 individuals +/- SEM are shown and significance indicated with a one-way ANOVA with Tukey's post-hoc analysis with *p<0.05, **p<0.01.



Supplementary Figure 10: PBMC additional cell population quantification. Relative cellular expression (as indicated on y-axis) or activation for various cell populations and their activation states are shown for (a) monocytes relative to total cell populations, (b) macrophages relative to total cell populations, (c) dendritic cells (DC) relative to total cell populations, (d) DC2 relative to total DC cell populations, (e) regulatory T cells (Tregs) and (f) T helper (Th)17 cells, both with an inset of activated cell populations (Treg, CTLA4⁺ and Th17, IL17A⁺ICOS⁺) relative to total CD4⁺ cells, (g) B cells and (h) memory B cells relative total cell populations, (i) NK cells relative to total cell populations, (j) CD8 TNF α cells and (k) CD8 IFN γ cells relative to total

CD8 cell populations and (I) Th1 and (m) Th2 cells relative to total CD4+ cell populations in their naïve and activated states (solid superimposed bars). In all cases, n = 3 individuals +/- SEM are shown and significance indicated with a one-way ANOVA with Tukey's post-hoc analysis with *p<0.05, **p<0.01.

Supplementary Table 1: Polyphenol-free Diet Composition The polyphenol-free diet was custom modified from Research Diets Ain-93G diet (New Brunswick, NJ, USA)

Class	Ingredient	Grams
Description		
Protein	Casein, Lactic, 30	200 g
	Mesh	
Protein	Cystine, L	3 g
Carbohydrate	Starch, Corn	397 g
Carbohydrate	vdrate Maltodextrin 10 132 g	
Carbohydrate	Sucrose, Fine	100 g
	Granulated	
Fiber	Cellulose, BW200	50 g
Fat	Corn Oil	70 g
Mineral S10022G		35 g
Vitamin	V10037	10 g
Vitamin	Choline Bitartrate	2.5 g
		Total: 1000 g

Caloric	
Information	
Protein	20.3 % kcal
Fat	15.8 % kcal
Carbohydrate	63.9 % kcal
Energy Density	4.00 kcal/g

Supplementary Table 2: Chronic Unpredictable Stress Protocol

Day	Stress 1	Stress 2						
Week 1	Week 1							
Monday	Crowding, 60 min	Cage Tilt, 45°, 12h (7PM - 7AM)						
Tuesday	Cold Swim (4°C, 5min)	Lights ON 12h, (6PM - 6AM)						
Wednesday	Cage Shaking, 20 min	Food deprivation, 12h (7PM - 7AM)						
Thursday	Wet Bedding, 6hr (10am- 4pm)	Forced swim, 5min, 4°C						
Friday	Restraint Stress, 1hr	Cage Tilt, 45°, 12h (7PM – 7AM)						
Saturday	Fox odor, 10h (8AM – 6PM)	Lights ON 12h, (6PM - 6AM)						
Sunday	Water deprivation, 10h (10AM-6PM)	Cold exposure (4C, 1hr)						
Week 2								
Monday	Forced swim, 5min, 4°C	No Bedding (12h, 6PM - 6AM)						
Tuesday	Cage Shaking, 20 min	Fox Odor (12h, 8PM - 8AM)						
Wednesday	Restraint Stress, 1hr	Water Deprivation (8h, 12AM - 8AM)						
Thursday	Forced swim, 5min, 4°C	Cold Exposure (20min, 4C)						
Friday	Wet Bedding, 10h (10AM – 8PM)	No Bedding, 12 h (8PM- 8AM)						
Saturday	Cage Tilt, 45°, 12h (7AM-7PM)	Fox odor, 10h (10PM – 8AM)						
Sunday	Cage Shaking, 20 min	Cage Tilt, 45°, 12h (7PM – 7AM)						
Week 3								
Monday	Cold Swim (4°C, 5min)	Food&Water Dep., 12h, (8PM-8AM)						
Tuesday	Cage Shaking, 20 min	Lights ON 12h, (6PM - 6AM)						
Wednesday	Crowding, 60 min	Cage Tilt, 45°, 12h (7PM – 7AM)						
Thursday	Wet Bedding, 8h (10AM- 6PM)	Restraint Stress, 1hr						
Friday	Food Deprivation, 12h (8AM-8PM)	No Bedding, 12 h (8PM- 8AM)						

Saturday	Fox odor, 8h (10AM –	Lights ON 12h, (6PM -	
	6PM)	6AM)	
Sunday	Restraint Stress, 1hr	Wet Bedding, 8h (10AM-	
		6PM)	
Week 4			
Monday	Fox odor, 10h (10AM –	Cage Tilt, 45°, 8h (12PM –	
	8PM)	8AM)	
Tuesday	Cold Swim (4°C, 5min)	Food&Water Dep., 12h,	
		(8PM-8AM)	
Wednesday	Cage Shaking, 20 min	Wet Bedding, 10h (10PM-	
		8AM)	
Thursday	Crowding, 60 min	Restraint Stress, 1hr	
Friday	Cold exposure (4C, 1hr)	Wet Bedding, 10h (10PM-	
		8AM)	
Saturday	No Bedding, 10h (8AM-	Cage Tilt, 45°, 8h (12PM –	
	6PM)	8AM)	
Sunday	Crowding, 60 min	Fox odor, 10h (10AM –	
		8PM)	

Subthreshold US						
Monday	Fox odor, 10h (10AM –	Cage Tilt, 45°, 8h (12PM –				
	8PM)	8AM)				
Tuesday	Cold Swim (4°C, 5min)	Food&Water Dep., 12h,				
		(8PM-8AM)				
Wednesday	Cage Shaking, 20 min	Wet Bedding, 10h (10PM-				
		8AM)				
Thursday	Crowding, 60 min	Restraint Stress, 1hr				
Friday	Cold exposure (4C, 1hr)	Wet Bedding, 10h (10PM-				
		8AM)				
Saturday	No Bedding, 10h (8AM-	Cage Tilt, 45°, 8h (12PM –				
	6PM)	8AM)				
Sunday	Crowding, 60 min	Fox odor, 10h (10AM –				
	_	8PM)				

Gene	Accession Number	Sequence (5' - 3')	Product Length (bp)
דו די	NIM 011005 2	F: CCGAAACCTCAGACAAAGCGTCA	144
ILK2	INM_011905.5	R: TCACACACCCCAGAAGCATCACAT	144
	NIM 021207.2	F: GCAAAGTCCCTGATGACATTCCTT	267
ILK4	<u>INIM_021297.5</u>	R: CCACAGCCACCAGATTCTCTAAA	207
II 6	NIM 001214054 1	F: CACAAGTCCGGAGACCAGAC	142
1L-0	INIM_001314034.1	R: CAGAATTGCCCATTGCACAAC	142
11 10	NIM 009261 4	F: TTCAGGCAGGCAGTATCACTC	60
IL-Ip	<u>INM_008301.4</u>	R: CCACGGGAAAGACACAGGTAG	09
Minn 2	NIM 1459274	F: AGAAGAGACCACGGCAGAA	102
Nirps	<u>INIM_143827.4</u>	R: CCTTGGACCAGGTTCAGTGT	102
Cara 1	NIM 000907.2	F: CACATTTCCAGGACTGACTGG	120
Casp1	<u>INM_009807.2</u>	R: AGACGTGTACGAGTGGTTGT	120
	NDA 021151252.2	F: CTGGTGGACACATACAGGAAGAC	105
ΝΓκΒ	<u>NM_021151352.2</u>	R: ATAGGCACTGTCTTCTTTCACCTC	195
		F: GTTTAAAACCAGACCCTGGAACT	150
ICAM	<u>NM_010493.3</u>	R: CGTCTGCAGGTCATCTTAGGAG	152
VCDA		F: CCGGCATATACGAGTGTGAAT	1.7.1
VCAM	<u>NM_011693.3</u>	R: ATGGCAGGTATTACCAAGGAAGAT	151
11 10	NDA 010549.2	F: GCTCTTACTGACTGGCATGAG	105
IL-10	<u>INM_010548.2</u>	R: CGCAGCTCTAGGAGCATGTG	105
	<u>NM 013653.3</u>	F: AGATCTCTGCAGCTGCCCTCA	170
CCLS		R: GGAGCACTTGCTGCTGGTGTAG	
	NM 011333.3	F: GCATCCACGTGTTGGCTCA	0.5
CCL2		R: CTCCAGCCTACTCATTGGGATCA	95
	ND (001014007.1	F: ACCAGAACTGTGAGGGTTGG	1.7.7
AHK	<u>NM_001314027.1</u>	R: TCTGAGGTGCCTGAACTCCT	155
IDO	ND 4 145040 2	F: CCTCATCCCTCCTTCCTTTC	010
IDO	<u>NM_145949.2</u>	R: GGAGCAATTGCCTGGTATGT	219
	ND 4 001100240 1	F: TTGGCCAGCGCCATCTT	100
Foxp3	<u>NM_001199348.1</u>	R: TGCCTCCTCCAGAGAGAAGTG	100
Th at	VM 021177762 1	F: AGCAAGGACGGCGAATGTT	107
Ibel	<u>AM_0211///02.1</u>	R: GGGTGGACATATAAGCGGTTC	18/
Cata?	XM 020547606 1	F: CGGCCATTCGTACATGGAA	135
Gaias	<u>AIVI_02934/090.1</u>	R: GGATACCTCTGCACCGTAGC	
RORC	NM 00120373/1	F: GTGGAGTTTGCCAAGCGGCTTT	108
NORC	<u>111VI_001273734.1</u>	R: CCTGCACATTCTGACTAGGACG	100
11_174	NM 010552.3	F: AAGGCAGCAGCGATCATCC	150
	<u>11111_010332.3</u>	R: GGAACGGTTGAGGTAGTCTGAG	150
Cyplal	<u>NM_001136059.2</u>	F: GGGTTTGACACAGTCACAACT	197

Supplementary Table 3: Mouse Primer Sequences

		R: GGGACGAAGGATGAATGCCG	
II 72D	NIM 144549 1	GTCCACCAAACTTCCCAGACAG	110
$\frac{1123R}{1123R} = \frac{11123R}{1112}$	CCTGAAGCAGGATGTCCTCTGA	110	
	NIM 001200726 1	F: ATGTGTCCGTCGTGGATCTGAC	122
GAPDH	<u>INIM_001289720.1</u>	R: AGACAACCTGGTCCTCAGTGTAG	152

Clone Target Source Metal Conjugate CD45 30-F11 Biolegend 112Cd, 114Cd, 116Cd MHCI(H-2)* M1/42Biolegend 112Cd, 114Cd, 116Cd Ly-6G* 1A8 Fluidigm 141 Pr CD11c* N418 Biolegend 142 Nd TCRb* H57-597 Fluidigm 143 Nd CD115* AFS98 Fluidigm 144 Nd F4/80* Fluidigm 146 Nd BM8 CD11b* M1/70 Biolegend 148 Nd CD19* 6D5 Biolegend 149 Sm CD25* 3C7 Biolegend 151 Eu CD3e* 145-2C11 Fluidigm 152 Sm CD16/32 93 Fluidigm 153 Eu CTLA4 UC10-4B9 Fluidigm 154 Sm ST2* DIH9 Biolegend 155 Gd Foxp3 FJK-16s Fluidigm 158 Gd RORgt B2D Fluidigm 159 Tb B220* RA3-6B2 Fluidigm 160 Gd 2E7' CD103* Biolegend 161 Dy TNFα MP6-T22 Fluidigm 162 Dy IL-23R* Novus Bio polyclonal 163 Dy Sca-1 D7 Fluidigm 164 Dy IFNγ XMG1.2 Fluidigm 165 Ho CD117* 2B8 Biolegend 166 Er NKp46* 167 Er 29A1.4 Fluidigm CD8* 53-6.7 Biolegend 168 Er IL22* NBP2-11699 Novus Bio 169 Tm NK1.1* Fluidigm PK136 170 Er CD4* RM4-5 Biolegend 172 Yb IL-17A TC11-18H10.1 Fluidigm 174 Yb CD127* A7R34 Biolegend 175 Lu ICOS* 7E.17G9 Fluidigm 176 Yb MHCII (I-A/I-E) M5.114.15.2 Fluidigm 209 Bi

Supplementary Table 4: Target Antibodies, clone and conjugate for CyTOF Experiments Tags marked with an asterisk (*) were used in calculating the viSNE analysis Supplementary Table 4: Fold change of relative gene expression of inflammatory markers in the prefrontal cortex. Detailed gene expression analysis of data presented in figure 2b. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
TLR4	Control	0 79 + 0 22	1 66 + 0 20	1 22 + 0 21	1 50 + 0 18
	BDPP	1.32 ± 0.25	1.00 ± 0.29	1.16 ± 0.15	0.95 ± 0.18
	Probiotic	0.72 ± 0.44	1.81 + 0.13	0.96 ± 0.21	1.27 ± 0.26
	Synbiotic	0.89 ± 0.20	1.11 ± 0.21	1.13 ± 0.19	0.98 ± 0.24
Nfkb	Control	0.97 ± 0.21	1.01 ± 0.14	0.91 ± 0.23	0.94 ± 0.24
	BDPP	1.25 ± 0.21	0.73 ± 0.23	1.03 ± 0.13	0.78 ± 0.11
	Probiotic	1.15 ± 0.24	1.10 ± 0.19	0.95 ± 0.16	1.05 ± 0.25
	Synbiotic	1.24 ± 0.17	0.93 ± 0.15	1.06 ± 0.17	0.88 ± 0.21
Casp1	Control	1.30 ± 0.20	1.45 ± 0.30	0.77 ± 0.22	1.62 ± 0.30
-	BDPP	0.99 ± 0.31	1.16 ± 0.24	1.26 ± 0.30	0.73 ± 0.34
	Probiotic	1.14 ± 0.26	1.69 ± 0.23	0.88 ± 0.30	1.04 ± 0.32
	Synbiotic	1.21 ± 0.30	1.13 ± 0.21	0.97 ± 0.36	1.25 ± 0.27
ll1b	Control	0.99 ± 0.37	1.91 ± 0.41	1.00 ± 0.24	1.60 ± 0.28
	BDPP	1.41 ± 0.39	1.57 ± 0.25	0.89 ± 0.23	0.80 ± 0.26
	Probiotic	1.05 ± 0.44	1.81 ± 0.39	1.01 ± 0.22	1.19 ± 0.09
	Synbiotic	1.11 ± 0.30	1.57 ± 0.30	1.10 ± 0.23	0.75 ± 0.42
CCL2	Control	0.97 ± 0.32	$\textbf{2.13} \pm \textbf{0.23}$	1.54 ± 0.15	3.50 ± 0.16
	BDPP	1.48 ± 0.09	1.68 ± 0.23	1.17 ± 0.19	1.95 ± 0.18
	Probiotic	1.33 ± 0.44	3.54 ± 0.32	1.58 ± 0.22	0.78 ± 0.40
	Synbiotic	1.32 ± 0.26	1.87 ± 0.26	1.57 ± 0.17	1.00 ± 0.13
CCL5	Control	0.83 ± 0.06	2.21 ± 0.16	1.09 ± 0.14	1.68 ± 0.15
	BDPP	0.65 ± 0.05	1.25 ± 0.17	1.28 ± 0.16	1.59 ± 0.06
	Probiotic	0.92 ± 0.12	1.23 ± 0.40	0.80 ± 0.05	2.09 ± 0.14
	Synbiotic	0.95 ± 0.13	1.27 ± 0.20	1.16 ± 0.02	1.20 ± 0.24
ICAM	Control	1.24 ± 0.21	2.67 ± 0.18	1.18 ± 0.14	2.35 ± 0.15
	BDPP	2.08 ± 0.20	1.99 ± 0.28	1.04 ± 0.17	0.87 ± 0.06
	Probiotic	1.30 ± 0.20	1.48 ± 0.19	1.06 ± 0.04	1.02 ± 0.47
	Synbiotic	1.40 ± 0.19	1.04 ± 0.15	1.24 ± 0.18	0.47 ± 0.13
VCAM	Control	1.16 ± 0.11	1.98 ± 0.18	0.85 ± 0.15	2.08 ± 0.23
	BDPP	1.02 ± 0.19	1.78 ± 0.19	1.02 ± 0.17	2.01 ± 0.16
	Probiotic	0.60 ± 0.15	2.00 ± 0.17	1.13 ± 0.15	1.47 ± 0.12
	Synbiotic	0.81 ± 0.25	1.74 ± 0.10	1.21 ± 0.11	1.17 ± 0.16

Supplementary Table 5: Ileal Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in figure 4c. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
Π _17Δ	Control	1 35 + 0 25	2 15 + 0 12	0 78 + 0 10	1.40 ± 0.21
	BDPP	1.03 ± 0.23 1 08 + 0 14	1.13 ± 0.12 1.87 ± 0.21	1.03 ± 0.10	0.62 ± 0.21
	Probiotic	1.19 ± 0.20	1.29 ± 0.09	1.07 ± 0.24	1.00 ± 0.13
	Synbiotic	1.08 ± 0.19	1.07 ± 0.06	0.85 ± 0.20	1.33 ± 0.20
IL-1b	Control	0.95 ± 0.11	1.83 ± 0.22	0.64 ± 0.09	1.26 ± 0.31
	BDPP	0.86 ± 0.12	0.88 ± 0.20	0.60 ± 0.10	0.51 ± 0.07
	Probiotic	0.73 ± 0.21	0.76 ± 0.19	0.52 ± 0.15	1.19 ± 0.21
	Synbiotic	0.72 ± 0.11	0.65 ± 0.12	0.94 ± 0.15	0.82 ± 0.31
IL-6	Control	0.86 ± 0.31	1.99 ± 0.09	0.77 ± 0.13	1.61 ± 0.15
	BDPP	1.75 ± 0.21	0.93 ± 0.07	0.97 ± 0.20	0.76 ± 0.10
	Probiotic	1.21 ± 0.15	1.39 ± 0.22	$\textbf{0.68} \pm \textbf{0.12}$	1.03 ± 0.25
	Synbiotic	1.43 ± 0.22	0.85 ± 0.24	$\textbf{0.88} \pm \textbf{0.12}$	0.90 ± 0.05
IL-10	Control	1.03 ± 0.11	1.34 ± 0.21	0.76 ± 0.14	0.92 ± 0.14
	BDPP	0.96 ± 0.23	1.62 ± 0.10	0.91 ± 0.10	1.14 ± 0.20
	Probiotic	0.77 ± 0.21	1.83 ± 0.31	0.58 ± 0.08	0.62 ± 0.08
	Synbiotic	1.82 ± 0.13	1.32 ± 0.21	1.06 ± 0.21	1.16 ± 0.09
Kynurenine	Control	0.83 ± 0.12	2.53 ± 0.30	1.04 ± 0.10	$\textbf{2.11} \pm \textbf{0.21}$
	BDPP	1.25 ± 0.20	0.87 ± 0.16	0.84 ± 0.17	$\textbf{0.85}\pm\textbf{0.15}$
	Probiotic	1.34 ± 0.18	1.36 ± 0.22	$\overline{0.93\pm0.20}$	0.48 ± 0.41
	Synbiotic	1.64 ± 0.21	0.62 ± 0.11	1.05 ± 0.17	0.70 ± 0.20

Supplementary Table 6: Ileal Immune Cell Marker Gene Expression. Detailed gene expression analysis of data presented in figure 4d. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
	Control	1.00 + 0.04	1.07 \ 0.04	4 47 + 0.00	4 50 1 0 40
гохр3	Control	1.20 ± 0.34	1.07 ± 0.21	1.17 ± 0.09	1.58 ± 0.18
	BDPP	1.15 ± 0.17	1.09 ± 0.06	1.57 ± 0.14	1.89 ± 0.27
	Probiotic	1.24 ± 0.18	0.77 ± 0.10	1.23 ± 0.12	1.50 ± 0.21
	Synbiotic	1.69 ± 0.10	1.75 ± 0.19	1.65 ± 0.15	1.95 ± 0.11
Rorc	Control	1.20 ± 0.10	1.76 ± 0.14	0.98 ± 0.20	$\textbf{2.49} \pm \textbf{0.24}$
	BDPP	1.69 ± 0.12	1.42 ± 0.18	0.84 ± 0.19	$\textbf{2.98} \pm \textbf{0.10}$
	Probiotic	1.15 ± 0.09	$\textbf{1.63} \pm \textbf{0.19}$	$\textbf{1.13}\pm\textbf{0.13}$	1.92 ± 0.16
	Synbiotic	1.70 ± 0.11	1.69 ± 0.20	$\textbf{1.18} \pm \textbf{0.13}$	2.37 ± 0.18
Gata3	Control	$\textbf{0.84} \pm \textbf{0.42}$	1.02 ± 0.21	1.46 ± 0.15	1.08 ± 0.11
	BDPP	$\textbf{0.99} \pm \textbf{0.11}$	1.02 ± 0.16	1.92 ± 0.17	$\textbf{0.89} \pm \textbf{0.25}$
	Probiotic	$\textbf{0.98} \pm \textbf{0.31}$	0.90 ± 0.17	1.29 ± 0.17	1.50 ± 0.20
	Synbiotic	1.32 ± 0.17	1.12 ± 0.19	2.03 ± 0.17	1.68 ± 0.21
Tbet	Control	1.33 ± 0.13	1.13 ± 0.14	1.02 ± 0.26	$\textbf{0.76} \pm \textbf{0.17}$
	BDPP	0.82 ± 0.42	1.33 ± 0.15	1.48 ± 0.42	1.73 ± 0.31
	Probiotic	1.07 ± 0.19	0.75 ± 0.27	0.72 ± 0.18	1.12 ± 0.11
	Synbiotic	1.13 ± 0.14	1.18 ± 0.29	1.05 ± 0.25	1.08 ± 0.13
Ahr	Control	1.22 ± 0.13	1.00 ± 0.13	1.30 ± 0.06	1.14 ± 0.13
	BDPP	1.26 ± 0.13	1.19 ± 0.13	1.36 ± 0.16	1.72 ± 0.10
	Probiotic	1.12 ± 0.14	1.00 ± 0.14	1.27 ± 0.12	1.43 ± 0.14
	Synbiotic	1.42 ± 0.14	1.31 ± 0.25	1.26 ± 0.21	1.31 ± 0.10
Cyp1a1	Control	0.72 ± 0.08	$\textbf{2.24} \pm \textbf{0.16}$	0.86 ± 0.12	2.59 ± 0.21
	BDPP	0.40 ± 0.28	2.27 ± 0.31	1.11 ± 0.18	2.46 ± 0.25
	Probiotic	0.64 ± 0.31	$\overline{2.55\pm0.18}$	$\overline{0.80\pm0.03}$	$\textbf{2.48} \pm \textbf{0.18}$
	Synbiotic	1.02 ± 0.17	1.27 ± 0.15	0.74 ± 0.11	1.64 ± 0.15

Supplementary Table 7: Prefrontal Cortex Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 2. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	0.99 ± 0.17	1.71 ± 0.13	1.10 ± 0.10	0.77 ± 0.06
	BDPP	1.41 ± 0.22	1.22 ± 0.20	1.02 ± 0.19	0.96 ± 0.11
	Probiotic	1.44 ± 0.20	1.44 ± 0.11	1.05 ± 0.22	1.00 ± 0.31
	Synbiotic	1.32 ± 0.09	$\textbf{0.83} \pm \textbf{0.08}$	0.91 ± 0.07	1.06 ± 0.22
IL-6	Control	1.05 ± 0.13	$\textbf{0.87} \pm \textbf{0.11}$	0.97 ± 0.21	1.00 ± 0.21
	BDPP	1.12 ± 0.22	0.98 ± 0.21	1.04 ± 0.25	0.93 ± 0.13
	Probiotic	0.98 ± 0.06	1.05 ± 0.30	0.99 ± 0.29	0.89 ± 0.13
	Synbiotic	0.99 ± 0.15	1.09 ± 0.18	1.00 ± 0.11	0.85 ± 0.14
IL-10	Control	0.90 ± 0.10	1.17 ± 0.13	1.18 ± 0.29	1.10 ± 0.07
	BDPP	0.83 ± 0.03	1.33 ± 0.20	1.04 ± 0.20	1.32 ± 0.14
	Probiotic	1.29 ± 0.10	1.78 ± 0.21	1.17 ± 0.10	1.14 ± 0.11
	Synbiotic	1.47 ± 0.07	1.15 ± 0.22	0.78 ± 0.13	1.00 ± 0.10

Supplementary Table 8: Hippocampus Inflammatory Cytokine Expression.

Detailed cytokine expression analysis of data presented in Supplementary Figure 2. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	1.50 ± 0.21	1.18 ± 0.09	0.80 ± 0.27	0.80 ± 0.22
	BDPP	1.35 ± 0.11	0.63 ± 0.22	0.97 ± 0.18	1.06 ± 0.14
	Probiotic	0.88 ± 0.10	0.92 ± 0.21	0.53 ± 0.05	1.04 ± 0.35
	Synbiotic	0.93 ± 0.22	0.72 ± 0.10	1.03 ± 0.21	1.05 ± 0.29
IL-6	Control	1.30 ± 0.24	1.03 ± 0.71	1.13 ± 0.14	$\textbf{0.99} \pm \textbf{0.28}$
	BDPP	1.07 ± 0.14	$\textbf{0.80} \pm \textbf{0.19}$	$\textbf{1.18} \pm \textbf{0.14}$	1.18 ± 0.16
	Probiotic	1.12 ± 0.21	0.80 ± 0.23	1.07 ± 0.12	1.12 ± 0.18
	Synbiotic	0.96 ± 0.22	0.72 ± 0.13	0.89 ± 0.11	0.86 ± 0.11
IL-10	Control	1.08 ± 0.34	0.99 ± 0.34	0.78 ± 0.17	0.96 ± 0.14
	BDPP	0.88 ± 0.12	$\textbf{0.78} \pm \textbf{0.21}$	1.23 ± 0.72	1.62 ± 0.32
	Probiotic	1.25 ± 0.14	1.10 ± 0.22	0.95 ± 0.17	0.75 ± 0.18
	Synbiotic	1.22 ± 0.21	0.93 ± 0.19	0.70 ± 0.12	0.97 ± 0.21

Supplementary Table 9: Hippocampal Immune Cell Marker Gene Expression.

Detailed gene expression analysis of data presented in Supplementary Figure 2. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
TIr4	Control	0.76 ± 0.12	1.93 ± 0.13	1.14 ± 0.15	$\textbf{2.10} \pm \textbf{0.14}$
	BDPP	0.77 ± 0.15	$\textbf{2.37} \pm \textbf{0.33}$	0.96 ± 0.24	2.34 ± 0.12
	Probiotic	0.80 ± 0.21	$\textbf{2.76} \pm \textbf{0.27}$	1.05 ± 0.16	$\textbf{2.22}\pm\textbf{0.22}$
	Synbiotic	0.90 ± 0.12	1.99 ± 0.16	1.79 ± 0.13	2.55 ± 0.12
Nfkb	Control	0.82 ± 0.13	1.46 ± 0.22	1.23 ± 0.23	1.46 ± 0.19
	BDPP	0.95 ± 0.13	1.15 ± 0.29	1.52 ± 0.17	1.99 ± 0.17
	Probiotic	0.83 ± 0.17	2.06 ± 0.25	1.13 ± 0.26	1.51 ± 0.25
	Synbiotic	1.03 ± 0.11	1.29 ± 0.31	1.39 ± 0.21	1.70 ± 0.14
Casp1	Control	1.03 ± 0.16	1.00 ± 0.35	1.43 ± 0.28	0.75 ± 0.15
	BDPP	0.63 ± 0.41	1.58 ± 0.42	1.06 ± 0.21	1.57 ± 0.30
	Probiotic	1.11 ± 0.19	2.69 ± 0.25	0.95 ± 0.18	1.08 ± 0.30
	Synbiotic	0.64 ± 0.12	1.42 ± 0.14	$\textbf{1.28} \pm \textbf{0.16}$	0.92 ± 0.17
ll1b	Control	1.10 ± 0.31	1.24 ± 0.13	1.09 ± 0.26	1.75 ± 0.18
	BDPP	0.95 ± 0.16	1.33 ± 0.19	0.81 ± 0.63	1.05 ± 0.30
	Probiotic	1.41 ± 0.21	2.84 ± 0.15	0.60 ± 0.18	1.38 ± 0.13
	Synbiotic	1.59 ± 0.28	$\textbf{3.13}\pm\textbf{0.30}$	1.53 ± 0.41	1.02 ± 0.17
Ccl2	Control	0.70 ± 0.12	0.99 ± 0.17	0.74 ± 0.23	0.52 ± 0.16
	BDPP	0.52 ± 0.15	1.00 ± 0.25	0.41 ± 0.29	0.54 ± 0.11
	Probiotic	0.44 ± 0.12	1.52 ± 0.27	0.81 ± 0.16	0.77 ± 0.12
	Synbiotic	1.47 ± 0.42	0.73 ± 0.25	1.34 ± 0.14	1.32 ± 0.29
Ccl5	Control	1.90 ± 0.52	1.46 ± 0.17	1.00 ± 0.13	1.71 ± 0.26
	BDPP	0.71 ± 0.12	1.33 ± 0.15	1.02 ± 0.15	1.18 ± 0.31
	Probiotic	0.84 ± 0.13	1.70 ± 0.25	1.26 ± 0.27	0.90 ± 0.31
	Synbiotic	$1.0\ \pm 0.19$	0.87 ± 0.17	1.37 ± 0.22	0.84 ± 0.21
Icam	Control	0.78 ± 0.14	1.01 ± 0.81	1.04 ± 0.41	0.98 ± 0.22
	BDPP	0.66 ± 0.26	0.90 ± 0.16	1.23 ± 0.29	0.29 ± 0.45
	Probiotic	0.95 ± 0.31	1.80 ± 0.13	0.72 ± 0.14	0.41 ± 0.23
	Synbiotic	1.63 ± 0.17	1.80 ± 0.23	$\textbf{1.18} \pm \textbf{0.31}$	0.31 ± 0.31
Vcam	Control	0.86 ± 0.18	0.85 ± 0.31	$0.89\pm0.2\overline{5}$	$1.09\pm0.1\overline{6}$
	BDPP	0.70 ± 0.18	0.73 ± 0.27	0.93 ± 0.28	0.53 ± 0.42
	Probiotic	0.72 ± 0.22	1.18 ± 0.14	0.79 ± 0.30	1.10 ± 0.43
	Synbiotic	$1.09\pm0.3\overline{2}$	0.80 ± 0.20	1.07 ± 0.28	$0.98\pm0.2\overline{6}$

Supplementary Table 10: Cortex Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 2. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/-SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	1.34 ± 0.14	1.54 ± 0.18	0.57 ± 0.34	0.94 ± 0.32
	BDPP	1.22 ± 0.19	1.70 ± 0,24	0.80 ± 0.26	1.04 ± 0.31
	Probiotic	0.85 ± 0.17	0.95 ± 0.22	0.78 ± 0.21	0.77 ± 0.21
	Synbiotic	1.05 ± 0.15	1.10 ± 0.10	$\textbf{0.93} \pm \textbf{0.32}$	1.56 ± 0.31
IL-6	Control	1.08 ± 0.08	$\textbf{0.83} \pm \textbf{0.09}$	0.86 ± 0.16	1.21 ± 0.16
	BDPP	1.05 ± 0.21	$\textbf{0.43}\pm\textbf{0.12}$	1.88 ± 0.23	$\textbf{0.92}\pm\textbf{0.21}$
	Probiotic	1.01 ± 0.11	1.04 ± 0.25	$\textbf{1.48} \pm \textbf{0.31}$	$\textbf{0.38} \pm \textbf{0.30}$
	Synbiotic	1.01 ± 0.21	$\textbf{0.68} \pm \textbf{0.18}$	1.50 ± 0.21	0.94 ± 0.32
IL-10	Control	1.21 ± 0.21	1.63 ± 0.13	$\textbf{0.63} \pm \textbf{0.12}$	0.97 ± 0.17
	BDPP	1.25 ± 0.31	1.40 ± 0.21	$\textbf{0.85}\pm\textbf{0.21}$	$\textbf{0.88} \pm \textbf{0.21}$
	Probiotic	1.10 ± 0.17	$\textbf{0.89} \pm \textbf{0.19}$	0.72 ± 0.18	0.97 ± 0.16
	Synbiotic	1.37 ± 0.21	0.90 ± 0.17	1.00 ± 0.13	1.07 ± 0.22

Supplementary Table 11: Cortex Immune Cell Marker Gene Expression. Detailed gene expression analysis of data presented in Supplementary Figure 2. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
Tlr4	Control	0.91 ± 0.16	1.41 ± 0.19	1.48 ± 0.15	1.26 ± 0.18
	BDPP	0.97 ± 0.06	1.47 ± 0.27	1.32 ± 0.13	1.26 ± 0.17
	Probiotic	0.85 ± 0.20	1.41 ± 0.17	1.07 ± 0.08	1.04 ± 0.19
	Synbiotic	1.02 ± 0.16	1.42 ± 0.17	1.33 ± 0.07	0.84 ± 0.18
Nfkb	Control	0.85 ± 0.09	0.92 ± 0.14	$\textbf{0.88} \pm \textbf{0.17}$	1.23 ± 0.11
	BDPP	0.84 ± 0.11	0.82 ± 0.27	0.86 ± 0.26	1.29 ± 0.20
	Probiotic	0.95 ± 0.20	1.30 ± 0.31	0.58 ± 0.45	$\textbf{0.96} \pm \textbf{0.11}$
	Synbiotic	0.97 ± 0.20	0.80 ± 0.15	1.02 ± 0.15	0.99 ± 0.16
Casp1	Control	1.58 ± 0.24	1.40 ± 0.23	0.50 ± 0.09	1.05 ± 0.15
	BDPP	1.71 ± 0.25	1.74 ± 0.24	0.66 ± 0.05	1.20 ± 0.25
	Probiotic	1.38 ± 0.36	1.24 ± 0.22	0.60 ± 0.16	1.07 ± 0.20
	Synbiotic	1.06 ± 0.28	1.03 ± 0.22	0.76 ± 0.19	1.63 ± 0.24
ll1b	Control	1.06 ± 0.13	2.62 ± 0.30	1.50 ± 0.24	1.71 ± 0.25
	BDPP	0.73 ± 0.16	1.76 ± 0.16	1.08 ± 0.34	0.90 ± 0.17
	Probiotic	1.12 ± 0.16	$\textbf{2.66} \pm \textbf{0.31}$	1.50 ± 0.19	0.84 ± 0.14
	Synbiotic	1.06 ± 0.18	$\textbf{2.68} \pm \textbf{0.30}$	$\textbf{0.93} \pm \textbf{0.17}$	1.43 ± 0.30
Ccl2	Control	1.13 ± 0.34	1.04 ± 0.19	0.61 ± 0.19	0.95 ± 0.23
	BDPP	0.76 ± 0.29	1.30 ± 0.53	$\textbf{0.79} \pm \textbf{0.20}$	0.68 ± 0.14
	Probiotic	1.95 ± 0.26	$\textbf{2.11} \pm \textbf{0.22}$	0.41 ± 0.32	1.14 ± 0.11
	Synbiotic	1.56 ± 0.22	1.98 ± 0.16	0.30 ± 0.12	1.46 ± 0.16
Ccl5	Control	1.16 ± 0.19	1.72 ± 0.22	0.70 ± 0.30	1.00 ± 0.14
	BDPP	0.72 ± 0.30	1.32 ± 0.29	0.61 ± 0.13	1.31 ± 0.16
	Probiotic	0.77 ± 0.18	1.50 ± 0.26	1.01 ± 0.17	0.90 ± 0.30
	Synbiotic	1.11 ± 0.16	1.43 ± 0.28	0.51 ± 0.14	1.05 ± 0.18
Icam	Control	0.99 ± 0.06	$\textbf{2.33} \pm \textbf{0.18}$	2.47 ± 0.40	1.89 ± 0.17
	BDPP	0.60 ± 0.22	1.49 ± 0.15	1.98 ± 0.38	2.62 ± 0.25
	Probiotic	0.87 ± 0.15	$\textbf{3.60} \pm \textbf{0.17}$	1.29 ± 0.34	$\textbf{2.18} \pm \textbf{0.31}$
	Synbiotic	1.68 ± 0.21	1.48 ± 0.22	0.54 ± 0.14	1.58 ± 0.44
Vcam	Control	$\textbf{0.92} \pm \textbf{0.22}$	0.98 ± 0.25	$\textbf{1.19}\pm\textbf{0.14}$	1.17 ± 0.15
	BDPP	0.83 ± 0.10	1.03 ± 0.28	$\overline{1.16\pm0.10}$	1.22 ± 0.25
	Probiotic	0.83 ± 0.02	1.55 ± 0.13	$\overline{0.69\pm0.19}$	$\overline{1.50\pm0.31}$
	Synbiotic	0.97 ± 0.09	$1.83\pm0.2\overline{4}$	$0.67\pm0.2\overline{5}$	$1.06\pm0.2\overline{1}$

Supplementary Table 12: Cerebellum Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 2. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/-SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	0.92 ± 0.08	0.95 ± 0.06	0.93 ± 0.13	0.76 ± 0.14
	BDPP	1.19 ± 0.09	1.02 ± 0.21	1.23 ± 0.15	0.69 ± 0.21
	Probiotic	1.07 ± 0.22	0.76 ± 0.15	0.64 ± 0.21	0.82 ± 0.12
	Synbiotic	1.39 ± 0.21	$\textbf{0.78} \pm \textbf{0.21}$	0.89 ± 0.22	$\textbf{0.88} \pm \textbf{0.21}$
IL-6	Control	$\textbf{0.68} \pm \textbf{0.21}$	0.86 ± 0.14	$\textbf{0.83} \pm \textbf{0.13}$	0.63 ± 0.04
	BDPP	0.78 ± 0.15	0.86 ± 0.21	1.36 ± 0.41	0.72 ± 0.09
	Probiotic	1.30 ± 0.22	0.96 ± 0.22	0.88 ± 0.15	0.97 ± 0.20
	Synbiotic	1.45 ± 0.21	0.84 ± 0.29	1.00 ± 0.25	1.11 ± 0.25
IL-10	Control	0.80 ± 0.28	1.58 ± 0.21	0.73 ± 0.14	1.01 ± 0.21
	BDPP	$\textbf{0.83} \pm \textbf{0.19}$	1.30 ± 0.19	1.03 ± 0.25	$\textbf{0.93} \pm \textbf{0.14}$
	Probiotic	0.95 ± 0.21	0.90 ± 0.17	0.70 ± 0.21	0.87 ± 0.21
	Synbiotic	0.86 ± 0.15	1.25 ± 0.21	0.93 ± 0.22	2.41 ± 0.31

Supplementary Table 13: Cortex Immune Cell Marker Gene Expression. Detailed gene expression analysis of data presented in Supplementary Figure 2. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM.

		US	CUS	CUS+Rest	CUS+US
TIr4	Control	0.99 ± 0.17	0.85 ± 0.16	1.04 ± 0.19	0.95 ± 0.15
	BDPP	0.79 ± 0.26	0.82 ± 0.17	1.11 ± 0.23	1.07 ± 0.29
	Probiotic	1.46 ± 0.20	0.84 ± 0.13	0.98 ± 0.29	0.81 ± 0.19
	Synbiotic	1.26 ± 0.20	1.63 ± 0.17	0.84 ± 0.27	1.02 ± 0.17
Nfkb	Control	0.87 ± 0.15	1.31 ± 0.24	1.37 ± 0.17	1.64 ± 0.18
	BDPP	1.02 ± 0.61	1.02 ± 0.15	$\textbf{0.83}\pm\textbf{0.19}$	1.63 ± 0.14
	Probiotic	1.05 ± 0.21	0.70 ± 0.16	0.64 ± 0.32	1.34 ± 0.17
	Synbiotic	1.25 ± 0.31	0.75 ± 0.19	$\textbf{0.95}\pm\textbf{0.21}$	$\textbf{3.70} \pm \textbf{0.34}$
Casp1	Control	1.17 ± 0.21	1.30 ± 0.25	$\textbf{1.18} \pm \textbf{0.13}$	1.05 ± 0.21
	BDPP	0.64 ± 0.12	$\textbf{1.21}\pm\textbf{0.10}$	$\textbf{1.38} \pm \textbf{0.19}$	0.74 ± 0.19
	Probiotic	1.38 ± 0.10	1.32 ± 0.09	1.40 ± 0.22	1.06 ± 0.14
	Synbiotic	1.84 ± 0.16	1.05 ± 0.24	1.39 ± 0.16	0.96 ± 0.21
ll1b	Control	0.92 ± 0.22	$\textbf{2.63} \pm \textbf{0.40}$	2.32 ± 0.13	1.67 ± 0.16
	BDPP	0.28 ± 0.59	1.06 ± 0.32	0.82 ± 0.16	1.14 ± 0.17
	Probiotic	1.25 ± 0.43	1.02 ± 0.27	0.66 ± 0.12	1.29 ± 0.15
	Synbiotic	$\textbf{0.86} \pm \textbf{0.19}$	0.37 ± 0.30	0.57 ± 0.15	$\textbf{2.49} \pm \textbf{0.19}$
Ccl2	Control	0.75 ± 0.20	1.81 ± 0.12	0.91 ± 0.21	$\textbf{0.93} \pm \textbf{0.20}$
	BDPP	0.79 ± 0.27	1.16 ± 0.22	0.80 ± 0.15	1.16 ± 0.13
	Probiotic	1.26 ± 0.20	1.02 ± 0.22	1.01 ± 0.20	1.39 ± 0.23
	Synbiotic	1.03 ± 0.12	1.22 ± 0.14	1.18 ± 0.17	1.01 ± 0.17
Ccl5	Control	0.85 ± 0.29	2.07 ± 0.24	$\textbf{1.13}\pm\textbf{0.13}$	1.53 ± 0.15
	BDPP	0.98 ± 0.23	1.24 ± 0.17	$\textbf{1.10}\pm\textbf{0.21}$	1.48 ± 0.22
	Probiotic	0.72 ± 0.42	1.66 ± 0.18	$\textbf{0.91} \pm \textbf{0.11}$	$\textbf{1.18} \pm \textbf{0.21}$
	Synbiotic	0.78 ± 0.17	1.09 ± 0.12	1.29 ± 0.14	1.51 ± 0.17
Icam	Control	1.28 ± 0.15	1.37 ± 0.26	2.11 ± 0.16	2.67 ± 0.19
	BDPP	0.57 ± 0.18	0.60 ± 0.07	0.72 ± 0.09	1.39 ± 0.12
	Probiotic	$\textbf{2.16} \pm \textbf{0.18}$	0.81 ± 0.20	2.02 ± 0.16	1.45 ± 0.15
	Synbiotic	1.60 ± 0.14	0.55 ± 0.06	1.80 ± 0.19	1.14 ± 0.24
Vcam	Control	1.13 ± 0.20	0.90 ± 0.14	$\textbf{0.89} \pm \textbf{0.14}$	1.38 ± 0.14
	BDPP	0.92 ± 0.16	0.85 ± 0.15	$\overline{0.54\pm0.22}$	0.76 ± 0.12
	Probiotic	1.71 ± 0.23	0.66 ± 0.13	$\overline{0.96\pm0.20}$	1.07 ± 0.21
	Synbiotic	1.67 ± 0.27	$0.62\pm0.1\overline{8}$	$1.00\pm0.1\overline{4}$	$1.44\pm0.1\overline{8}$

Supplementary Table 14: Spleen Immune Cell Marker Gene Expression. Detailed gene expression analysis of data presented in Supplementary Figure 8. Each value is represented as the fold change of the relative gene expression (relative to *gapdh*) from stressed vs. unstressed controls at the same timepoint. Each value is the mean of n = 6 independent samples +/- SEM, where the SEM was calculated as the geometric mean of the SEM of the two groups used to build the ratio.

		US	CUS	CUS+Rest	CUS+US
Foxp3	Control	1.25 ± 0.21	0.84 ± 0.12	1.26 ± 0.19	1.21 ± 0.13
	BDPP	1.70 ± 0.30	0.91 ± 0.08	1.32 ± 0.21	1.84 ± 0.13
	Probiotic	1.01 ± 0.12	0.69 ± 0.12	0.91 ± 0.16	1.15 ± 0.13
	Synbiotic	1.22 ± 0.08	0.84 ± 0.24	0.80 ± 0.21	1.94 ± 0.23
Rorc	Control	1.58 ± 0.18	3.49 ± 0.35	1.83 ± 0.16	0.90 ± 0.13
	BDPP	2.01 ± 0.16	3.35 ± 0.53	1.55 ± 0.23	1.83 ± 0.26
	Probiotic	0.84 ± 0.17	2.75 ± 0.39	1.12 ± 0.30	0.81 ± 0.16
	Synbiotic	1.06 ± 0.17	1.96 ± 0.30	1.03 ± 0.16	0.96 ± 0.41
Gata3	Control	1.49 ± 0.10	1.19 ± 0.09	1.14 ± 0.22	0.82 ± 0.30
	BDPP	1.11 ± 0.15	1.16 ± 0.13	1.26 ± 0.21	1.21 ± 0.16
	Probiotic	0.92 ± 0.09	1.24 ± 0.08	1.34 ± 0.35	0.90 ± 0.16
	Synbiotic	1.09 ± 0.06	1.31 ± 0.10	1.16 ± 0.11	1.12 ± 0.18
Tbet	Control	0.76 ± 0.22	0.94 ± 0.17	1.15 ± 0.20	$\textbf{0.86} \pm \textbf{0.12}$
	BDPP	1.08 ± 0.20	1.05 ± 0.09	1.94 ± 0.40	1.74 ± 0.17
	Probiotic	$\textbf{0.84} \pm \textbf{0.19}$	1.07 ± 0.16	0.95 ± 0.17	$\textbf{0.73} \pm \textbf{0.18}$
	Synbiotic	0.72 ± 0.14	0.72 ± 0.13	1.21 ± 0.29	1.11 ± 0.28
Ahr	Control	0.87 ± 0.23	$\textbf{0.87} \pm \textbf{0.11}$	1.24 ± 0.22	0.74 ± 0.19
	BDPP	$\textbf{0.83}\pm\textbf{0.19}$	$\textbf{0.78} \pm \textbf{0.16}$	1.32 ± 0.18	0.81 ± 0.26
	Probiotic	$\textbf{0.85}\pm\textbf{0.17}$	$\textbf{0.73} \pm \textbf{0.15}$	$\textbf{1.28} \pm \textbf{0.12}$	$\textbf{0.83} \pm \textbf{0.15}$
	Synbiotic	0.87 ± 0.13	0.72 ± 0.12	0.87 ± 0.22	0.75 ± 0.17
Cyp1a1	Control	1.06 ± 0.23	2.00 ± 0.12	1.21 ± 0.18	$\textbf{2.14} \pm \textbf{0.21}$
	BDPP	0.94 ± 0.17	1.07 ± 0.19	0.94 ± 0.17	1.94 ± 0.41
	Probiotic	0.70 ± 0.14	0.86 ± 0.13	1.09 ± 0.19	2.29 ± 0.30
	Synbiotic	1.08 ± 0.19	0.61 ± 0.15	1.26 ± 0.13	1.27 ± 0.21
TIr4	Control	0.93 ± 0.11	1.20 ± 0.16	0.85 ± 0.20	0.96 ± 0.17
	BDPP	0.97 ± 0.21	1.02 ± 0.14	1.11 ± 0.16	1.13 ± 0.15
	Probiotic	0.86 ± 0.22	1.28 ± 0.20	0.86 ± 0.24	1.08 ± 0.13
	Synbiotic	0.94 ± 0.22	1.02 ± 0.15	0.76 ± 0.25	1.07 ± 0.14
Nfkb	Control	0.85 ± 0.11	0.95 ± 0.12	1.08 ± 0.17	$\textbf{0.88} \pm \textbf{0.11}$
	BDPP	0.80 ± 0.10	$\textbf{0.88} \pm \textbf{0.11}$	1.23 ± 0.21	1.07 ± 0.21
	Probiotic	0.78 ± 0.16	1.21 ± 0.11	0.89 ± 0.16	0.82 ± 0.18
	Synbiotic	0.78 ± 0.18	1.24 ± 0.20	0.91 ± 0.16	0.91 ± 0.14
ll1b	Control	0.72 ± 0.16	1.23 ± 0.14	1.44 ± 0.23	1.11 ± 0.15
	BDPP	0.77 ± 0.21	0.78 ± 0.19	0.91 ± 0.22	0.74 ± 0.10
	Probiotic	0.74 ± 0.18	1.55 ± 0.19	1.04 ± 0.32	0.92 ± 0.13
	Synbiotic	1.01 ± 0.26	1.47 ± 0.17	1.16 ± 0.31	0.94 ± 0.16
Ido	Control	0.45 ± 0.37	0.80 ± 0.11	0.86 ± 0.18	0.87 ± 0.18
	BDPP	0.97 ± 0.12	1.18 ± 0.21	1.63 ± 0.35	0.56 ± 0.17
	Probiotic	0.71 ± 0.20	$\textbf{0.79} \pm \textbf{0.15}$	0.83 ± 0.44	0.51 ± 0.12
	Synbiotic	0.68 ± 0.20	1.02 ± 0.17	0.82 ± 0.87	0.51 ± 0.16

Supplementary Table 15: Splenic Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 8. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/-SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	1.02 ± 0.08	1.34 ± 0.15	1.52 ± 0.21	1.00 ± 0.21
	BDPP	1.02 ± 0.11	1.03 ± 0.18	1.00 ± 0.13	0.47 ± 0.17
	Probiotic	0.93 ± 0.12	0.79 ± 0.21	1.19 ± 0.31	0.65 ± 0.13
	Synbiotic	1.20 ± 0.31	0.52 ± 0.17	1.36 ± 0.21	0.65 ± 0.14
IL-6	Control	0.87 ± 0.04	1.54 ± 0.09	1.01 ± 0.21	1.04 ± 0.30
	BDPP	0.90 ± 0.10	0.99 ± 0.12	1.19 ± 0.12	0.91 ± 0.13
	Probiotic	0.90 ± 0.12	0.61 ± 0.10	1.18 ± 0.17	0.90 ± 0.15
	Synbiotic	1.27 ± 0.31	0.70 ± 0.07	0.97 ± 0.20	0.88 ± 0.17
IL-10	Control	0.99 ± 0.12	$\textbf{0.78} \pm \textbf{0.13}$	1.22 ± 0.21	0.68 ± 0.13
	BDPP	0.90 ± 0.31	$\textbf{0.89} \pm \textbf{0.19}$	1.28 ± 0.26	1.06 ± 0.11
	Probiotic	1.40 ± 0.41	0.74 ± 0.17	0.94 ± 0.17	0.96 ± 0.12
	Synbiotic	1.36 ± 0.21	0.91 ± 0.04	0.98 ± 0.21	1.36 ± 0.40

Supplementary Table 16: Splenic Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 8. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/-SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	1.02 ± 0.12	1.34 ± 0.13	1.52 ± 0.22	1.00 ± 0.14
	BDPP	1.02 ± 0.09	1.03 ± 0.14	1.00 ± 0.16	0.47 ± 0.13
	Probiotic	0.93 ± 0.08	0.79 ± 0.19	1.19 ± 0.06	0.65 ± 0.07
	Synbiotic	1.20 ± 0.11	0.52 ± 0.21	1.36 ± 0.26	0.65 ± 0.12
IL-6	Control	0.87 ± 0.23	1.54 ± 0.13	1.01 ± 0.17	1.04 ± 0.21
	BDPP	0.90 ± 0.27	0.99 ± 0.25	$\textbf{1.19} \pm \textbf{0.22}$	0.91 ± 0.14
	Probiotic	0.90 ± 0.21	$\textbf{0.61} \pm \textbf{0.18}$	$\textbf{1.18} \pm \textbf{0.27}$	$\textbf{0.90} \pm \textbf{0.21}$
	Synbiotic	1.27 ± 0.27	0.70 ± 0.20	0.97 ± 0.31	$\textbf{0.88} \pm \textbf{0.17}$
IL-10	Control	0.99 ± 0.13	$\textbf{0.78} \pm \textbf{0.14}$	1.22 ± 0.30	$\textbf{0.68} \pm \textbf{0.13}$
	BDPP	0.90 ± 0.17	$\textbf{0.89} \pm \textbf{0.13}$	$\textbf{1.18} \pm \textbf{0.29}$	1.06 ± 0.14
	Probiotic	1.40 ± 0.21	0.74 ± 0.23	0.94 ± 0.08	0.96 ± 0.16
	Synbiotic	1.36 ± 0.28	0.91 ± 0.22	0.98 ± 0.09	1.36 ± 0.21

Supplementary Table 17: Serum Inflammatory Cytokine Expression. Detailed cytokine expression analysis of data presented in Supplementary Figure 8. Each value represents the fold change of the absolute cytokine expression between the stressed and non-stressed groups. Each value is the mean of n = 6 independent samples +/-SEM.

		US	CUS	CUS+Rest	CUS+US
IL-1b	Control	1.10 ± 0.09	1.13 ± 0.21	1.15 ± 0.21	1.69 ± 0.31
-	BDPP	1.06 ± 0.12	1.10 ± 0.08	0.88 ± 0.09	0.74 ± 0.21
	Probiotic	1.12 ± 0.22	1.11 ± 0.11	1.08 ± 1.12	2.18 ± 0.30
	Synbiotic	0.91 ± 0.21	0.91 ± 0.22	1.13 ± 0.12	0.49 ± 0.17
IL-6	Control	1.28 ± 0.21	1.25 ± 0.16	2.60 ± 0.41	$\textbf{2.15}\pm\textbf{0.21}$
	BDPP	0.65 ± 0.08	$\textbf{0.68} \pm \textbf{0.11}$	1.12 ± 0.22	$\textbf{0.86} \pm \textbf{0.21}$
	Probiotic	0.28 ± 0.34	$\textbf{2.23}\pm\textbf{0.21}$	2.61 ± 0.18	2.44 ± 31
	Synbiotic	1.07 ± 0.25	$\textbf{0.72} \pm \textbf{0.18}$	0.52 ± 0.28	0.50 ± 0.21
IL-10	Control	1.77 ± 0.21	6.04 ± 0.31	4.46 ± 0.21	$5.07 \pm \ 0.31$
	BDPP	1.88 ± 0.15	3.61 ± 0.22	2.39 ± 0.24	1.51 ± 0.21
	Probiotic	1.32 ± 0.20	$\textbf{2.19} \pm \textbf{0.28}$	0.93 ± 0.09	2.67 ± 0.16
	Synbiotic	1.62 ± 0.17	0.92 ± 0.13	0.17 ± 0.22	0.46 ± 0.21