Supplementary

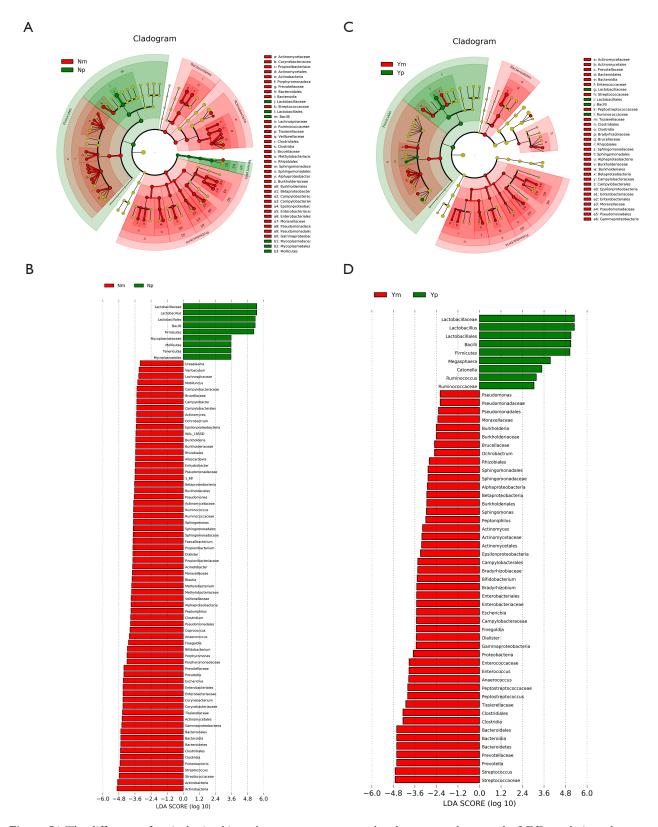


Figure S1 The difference of vaginal microbiome between pre-menopausal and menopausal women by LEfSe analysis at the genus level. (A,B) Nm: menopausal women without VAIN, Np: pre-menopausal women without VAIN; Ym: menopausal women with VAIN; Yp: pre-menopausal women with VAIN. In group Nm, observing 68 significant different genera (P<0.05), In group Np, discovering as many as 9 genera enriched (P<0.05); (C,D) in group Ym, observing 47 significant different genera (P<0.05). In group Yp, discovering as many as 9 genera enriched (P<0.05); only taxa with absolute values of LDA scores >2 were presented.

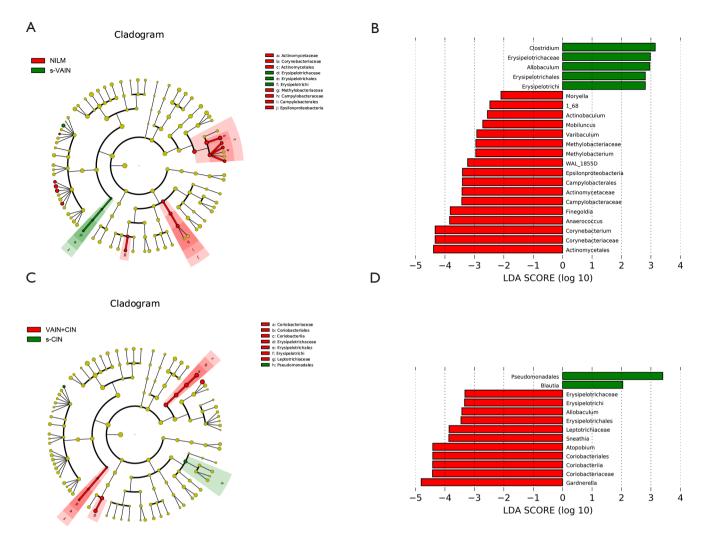


Figure S2 The difference of vaginal microbiome between VAIN and non-VAIN by LEfSe analysis at the genus level. (A,B) In group s-VAIN, observing 5 significant different genera (P<0.05), In group NILM, discovering as many as 17 genera enriched (P<0.05); (C,D) in group VAIN + CIN, observing 11 significant different genera (P<0.05). In group s-CIN, discovering as many as genera enriched (P<0.05); only taxa with absolute values of LDA scores >2 were presented.

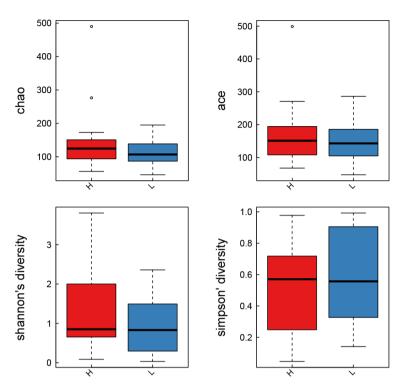


Figure S3 Bacteria species diversity and richness in VAIN1 compared with VAIN2/3. L: VAIN1; H: VAIN2/3; Shannon index, P=0.216; Simpson index, P=0.320; chao index, P=0.328; ace index, P=0.779.

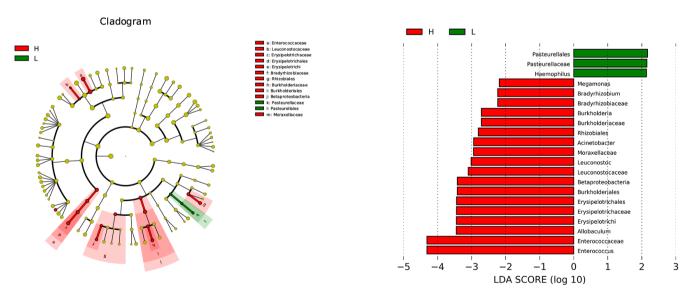


Figure S4 The difference of vaginal microbiome between VAIN1 and VAIN2/3 by LEfSe analysis at the genus level. L: VAIN1; H: VAIN2/3; In VAIN2/3, observing 18 significant different genera (P<0.05). InVAIN1, discovering as many as 3 genera enriched (P<0.05); only taxa with absolute values of LDA scores >2 were presented.

1 **Table S1.** HPV types prevalence

	VAINa	VAIN+CIN ^b	Number (%)
HPV16	9	6	15 (24.2)
HPV52	11	4	15 (24.2)
HPV53	9	1	10 (16.1)
HPV58	7	2	9 (14.5)
HPV66	7	2	9 (14.5)
HPV81	7	1	8 (12.9)
HPV56	6	2	8 (12.9)
HPV68	4	2	6 (9.7)
HPV33	4	2	6 (9.7)
HPV39	3	2	5 (8.1)
HPV59	3	1	4 (6.5)
HPV6	2	2	4 (6.5)
HPV35	3	0	3 (4.8)
HPV45	2	1	3 (4.8)
HPV31	2	0	2 (3.2)
HPV51	2	0	2 (3.2)
HPV18	2	0	2 (3.2)
HPV73	1	0	1 (1.6)

² aVAIN, vaginal intraepithelial neoplasia

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^{3 &}lt;sup>b</sup>CIN, cervical intraepithelial neoplasia

Table S2. Type-specific HPV viral load^a (CIN versus VAIN+CIN)

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Log10-transformed HPV viral load				
	CIN ^d	VAIN ^e +CIN	P	
	Mean (SD)	Mean (SD)	<u>, </u>	
all	5.37±1.26	4.79±1.57	0.244	
$\alpha 9^b$ quantity	3.95 ± 1.24	4.08±1.36	0.813	
HPV16	4.90 ± 0.25	4.00 ± 1.37	0.415	
HPV52	4.43 ± 0.89	4.20 ± 1.58	0.810	
HPV58	2.54 ± 0.76	4.36±2.15	0.375	
$\alpha7^{c}$ quantity	5.99 ± 0.82	4.66±2.64	0.393	

^aThe mean and SD of the log10-transformed virus copy numbers per10,000 human cells

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⁷ were considered the type-specific HPV viral load. Values are presented as mean (SD,

⁸ standard deviation) and compared using t test

⁹ $^{b}\alpha 9$ species including HPV-16, -31, -33, -35, -52, -58

¹⁰ $^{c}\alpha7$ species including HPV-18, -39, -45, -59, -68

¹¹ dCIN, cervical intraepithelial neoplasia

¹² eVAIN, vaginal intraepithelial neoplasia