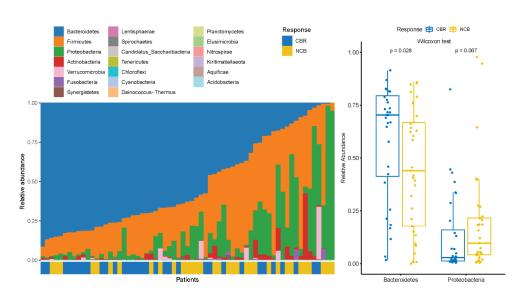
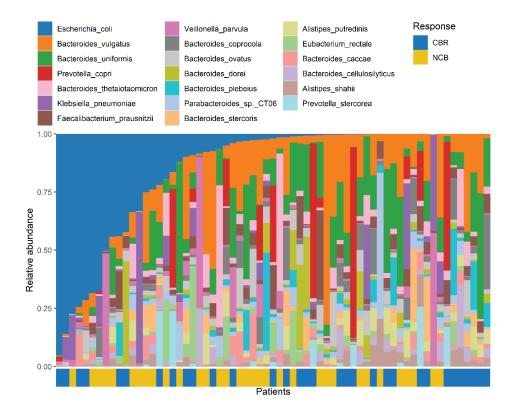
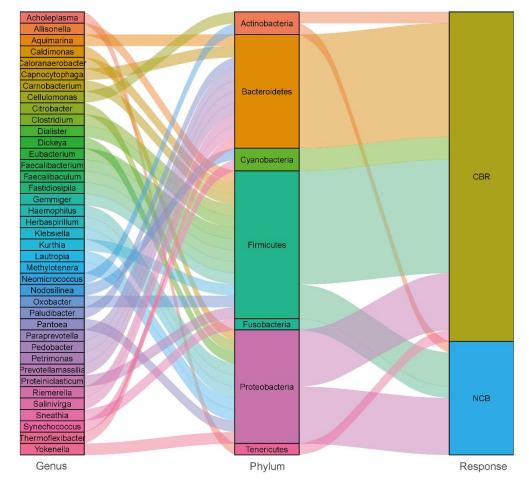
## **Supplementary Figures**



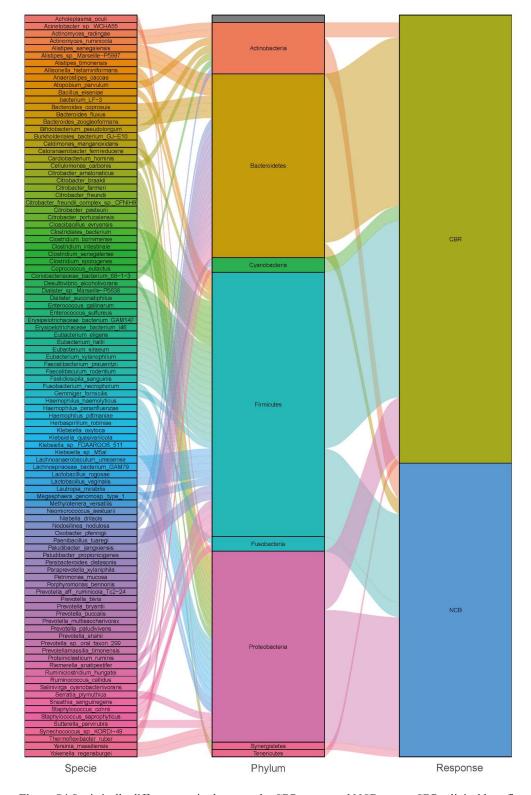
**Figure S1** Gut microbiome composition at the phylum level in 65 patients with hepatobiliary cancers (ordered by the most abundant taxa, Bacteroidetes phylum). CBR, clinical benefit response; NCB, non-clinical benefit.



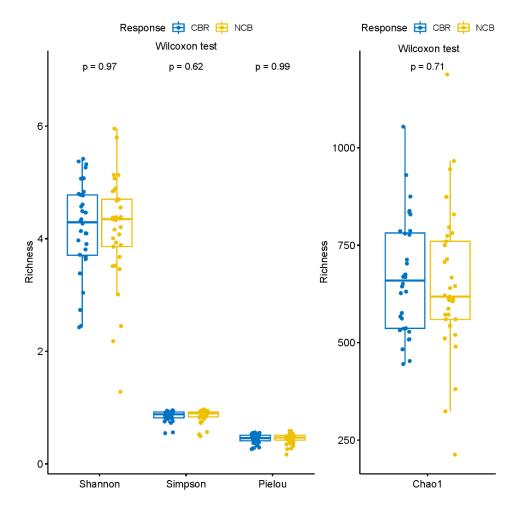
**Figure S2** Gut microbiome composition at the specie level in 65 patients with hepatobiliary cancers. CBR, clinical benefit response; NCB, non-clinical benefit.



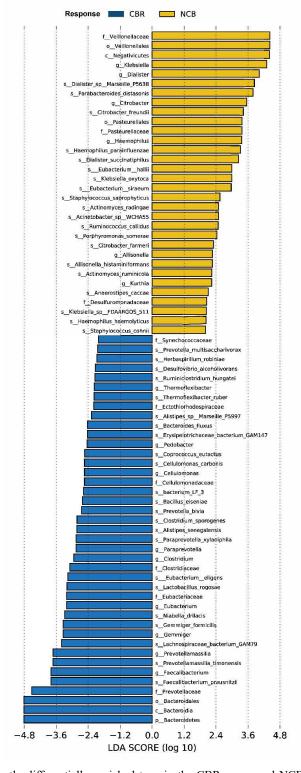
**Figure S3** Statistically different genera between the CBR group and NCB group. CBR, clinical benefit response; NCB, non-clinical benefit.



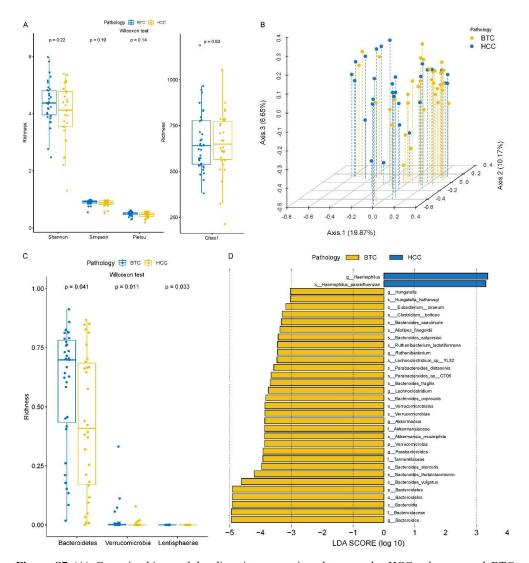
**Figure S4** Statistically different species between the CBR group and NCB group. CBR, clinical benefit response; NCB, non-clinical benefit.



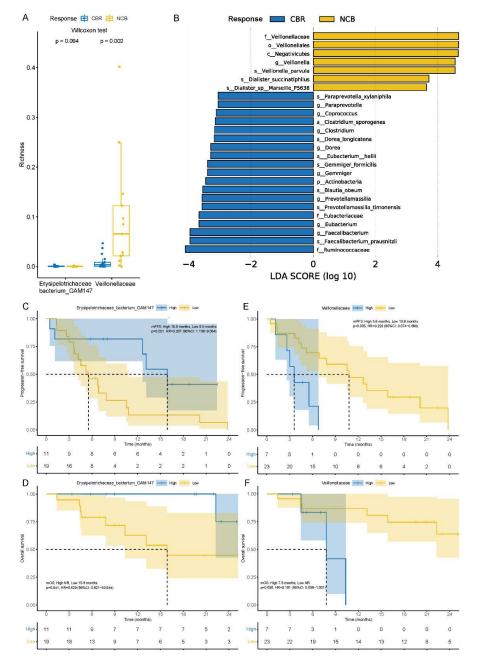
**Figure S5** Gut microbiome alpha diversity comparison between the CBR group and NCB group (Wilcoxon test). CBR, clinical benefit response; NCB, non-clinical benefit.



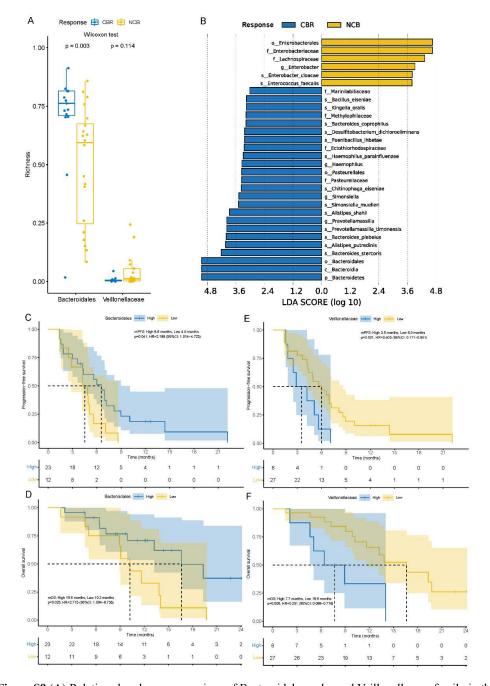
**Figure S6** Significantly differentially enriched taxa in the CBR group and NCB group identified by LEfSe (LDA>2, p<0.05). CBR, clinical benefit response; NCB, non-clinical benefit; LDA, linear discriminant analysis; LEfSe, linear discriminant analysis effect size.



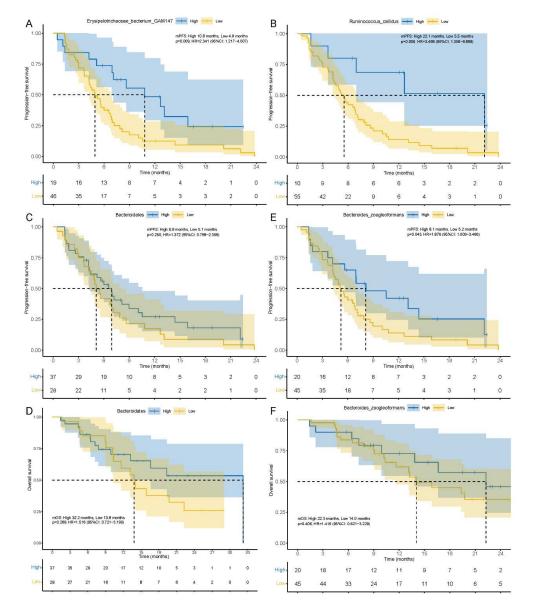
**Figure S7** (**A**) Gut microbiome alpha diversity comparison between the HCC subgroup and BTC subgroup (Wilcoxon test). (**B**) PCoA showed the beta diversity evaluated by Bray-Curtis distance between the HCC subgroup and BTC subgroup. (**C**) Significantly different phylum between the HCC subgroup and BTC subgroup (Wilcoxon test). (**D**) Significantly differentially enriched taxa in patients with HCC and BTC identified by LEfSe of the 30 patients with HCC (LDA>3, p<0.05). HCC, hepatocellular carcinoma; BTC, biliary tract cancer; PCoA, principal coordinate analysis; LEfSe, linear discriminant analysis effect size; LDA, linear discriminant analysis.



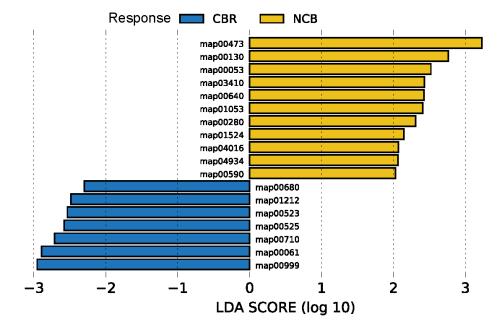
**Figure S8** (**A**) Relative abundance comparison of Erysipelotrichaceae bacterium-GAM147 and Veillonellaceae family in the CBR group and NCB group among the 30 HCC patients (Wilcoxon test). (**B**) Significantly differentially enriched taxa in the CBR group and NCB group among the 30 HCC patients identified by LEfSe (LDA>3, p<0.05). The Kaplan-Meier method with log-rank test estimates the median progression-free survival and median overall survival for the 30 HCC patients with higher or lower abundance of (**C-D**) Erysipelotrichaceae bacterium-GAM147 and (**E-F**) Veillonellaceae family. CBR, clinical benefit response; NCB, non-clinical benefit; HCC, hepatocellular carcinoma; LEfSe, linear discriminant analysis effect size; LDA, linear discriminant analysis; mPFS, median progression-free survival; mOS, median overall survival.



**Figure S9** (**A**) Relative abundance comparison of Bacteroidales order and Veillonellaceae family in the CBR group and NCB group among the 35 BTC patients (Wilcoxon test). (**B**) Significantly differentially enriched taxa in the CBR group and NCB group among the 35 BTC patients identified by LEfSe (LDA>3, p<0.05). The Kaplan-Meier method with log-rank test estimates the median progression-free survival and median overall survival for the 35 BTC patients with higher or lower abundance of (**C-D**) Bacteroidales order and (**E-F**) Veillonellaceae family. CBR, clinical benefit response; NCB, non-clinical benefit; BTC, biliary tract cancer; LEfSe, linear discriminant analysis effect size; LDA, linear discriminant analysis; mPFS, median progression-free survival; mOS, median overall survival.



**Figure S10** The Kaplan-Meier method with log-rank test estimates the median progression-free survival and overall survival for all 65 patients with higher or lower abundance of **(A)** Erysipelotrichaceae bacterium-GAM147, **(B)** Ruminococcus callidus, **(C-D)** Bacteroidales order and **(E-F)** Bacteroides zoogleoformans. mPFS, median progression-free survival; mOS, median overall survival.



**Figure S11** Significantly different KEGG pathways enriched in the CBR group and NCB group identified by LEfSe (LDA>2, p<0.05). KEGG, Kyoto Encyclopedia of Genes and Genomes; CBR, clinical benefit response; NCB, non-clinical benefit; LEfSe, linear discriminant analysis effect size; LDA, linear discriminant analysis.

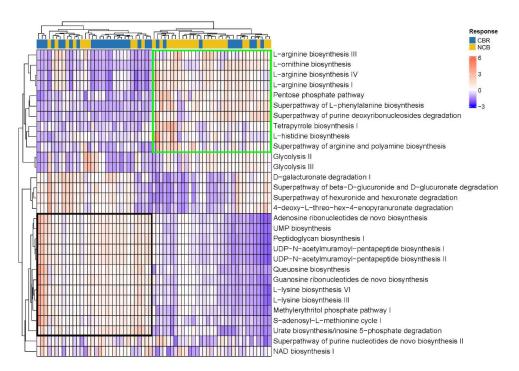


Figure S12 MetaCyc hierarchy of pathway classifications showed different metabolic pathways enriched in the CBR group and NCB group. CBR, clinical benefit response; NCB, non-clinical benefit.