No.	Ion mode	Rt(min)	Name	Formula	m/z	HMBD	Trend in model group	Regulated by syringin
1	M-H	0.46	Ornithine	C5H12N2O2	131.08	HMDB00214	up	$\checkmark$
2	M+H	0.75	L-Lysine	C6H14N2O2	147.11	HMDB00182	up	$\checkmark$
3	M+H	0.96	Citric acid	C6H8O7	215.02	HMDB00094	up	$\checkmark$
4	M+H	1.39	Uric acid	C5H4N4O3	169.04	HMDB00289	low	$\checkmark$
5	M-H	1.71	L-glutamine	C5H10N2O3	145.06	HMDB00641	low	$\checkmark$
6	M+H	2.11	Glucose	C6H12O6	181.07	HMDB00122	low	
7	M+H	2.22	Dimethyl succinate	C6H10O4	147.06	HMDB33837	up	$\checkmark$
8	M+H	2.43	N,N-dimethylglyine	C8H11N	144.08	HMDB01020	up	$\checkmark$
9	M+H	2.82	<b>D-Proline</b>	C5H9NO2	116.07	HMDB03411	up	$\checkmark$
10	M-H	3.49	Glutamyl-Serine	C8H13N2O6	232.07	HMDB28828	up	
11	M-H	3.59	Metyrosine	C10H13NO3	196.10	HMDB14903	low	
12	M+H	3.88	Carnitine	C7H15NO3	184.10	HMDB00062	up	$\checkmark$
13	M-H	4.03	Heptylmalonic acid	C10H18O4	201.11	HMDB59719	up	$\checkmark$
14	M-H	4.29	Taurocholic acid	C26H45NO7S	514.28	HMDB00036	up	$\checkmark$
15	M+H	4.56	3-Oxodecanoic acid	C14H26O3	265.17	HMDB10730	low	
16	M+H	5.35	Glycocholic acid	C26H43NO6	488.30	HMDB00138	up	
17	M-H	5.48	Pyruvic acid	C3H4O3	87.01	HMDB00243	low	$\checkmark$
18	M+H	6.05	L-Phenylalanine	C9H11NO2	166.08	HMDB00159	up	$\checkmark$
19	M-H	6.35	LysoPC(15:0)	C23H48NO7P	480.31	HMDB10381	low	$\checkmark$
20	M+H	6.68	Biliverdin	C33H34N4O6	583.26	HMDB01008	up	$\checkmark$
21	M+H	6.87	LysoPE(16:0/0:0)	C21H44NO7P	476.28	HMDB11503	low	
22	M-H	7.23	Arachidonic acid	C20H32O2	303.23	HMDB01043	up	$\checkmark$
23	M+H	7.41	LysoPE(0:0/20:0)	C25H52NO7P	510.36	HMDB11481	low	$\checkmark$
24	M-H	7.51	LysoPC(18:1(9Z))	C26H52NO7P	520.34	HMDB02815	low	$\checkmark$
25	M-H	7.83	LysoPC(P-16:0)	C24H50NO6P	478.33	HMDB10407	low	
26	M-H	8.05	LysoPC(17:0)	C25H52NO7P	508.34	HMDB12108	low	$\checkmark$
27	M+H	8.19	Oleamide	C18H35NO	282.28	HMDB02117	up	$\checkmark$
28	M-H	8.41	LysoPC(16:0)	C24H50NO7P	494.33	HMDB10382	low	$\checkmark$
29	M-H	8.55	LysoPC(18:0)	C26H54NO7P	522.36	HMDB10384	low	$\checkmark$
30	M-H	8.80	Palmitoleic acid	C16H30O2	253.22	HMDB03229	up	$\checkmark$
31	M+H	9.02	8-HETE	C20H32O3	321.24	HMDB04679	low	$\checkmark$
32	M+H	9.46	Triethyl citrate	C12H20O7	299.11	HMDB34263	up	·
33	M-H	9.81	LysoPE(18:0/0:0)	C23H48NO7P	480.31	HMDB11130	low	~

**Table S1** The differentially expressed metabolites in hepatitis B animal model using metabolomics strategy.









IL-10

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HA





PCIII



HBsAg



IFN-y









Figure S1 Biochemistry analysis of liver lesion and the anti-HBV effect of syringing. Model group vs control group: #p < 0.05, ##p < 0.01; Model group and syringin-treated group: \*p < 0.05; \*\*p < 0.01.



**Figure S2** Compared with control group, liver tissue from the model group exhibits unordered hepatocyte cords and hepatic lobule structure, cell morphology swelling, abnormal cell nuclear atrophy, severe fatty degeneration and fibrosis, spotty or focal necrosis and obvious inflammatory cells infiltration. Hemorrhagic necrosis with foci of lymphomonocytic infiltration around fibrosis tissue can be easily seen. Syringin gradually improved the pathological changes of liver tissue, and by the eighth week, there were mainly mild fibrotic lesions.