Supporting Information

Advanced Generation of Paeonol-Phenylsulfonyl Derivatives as Potential Anti-HBV Agents

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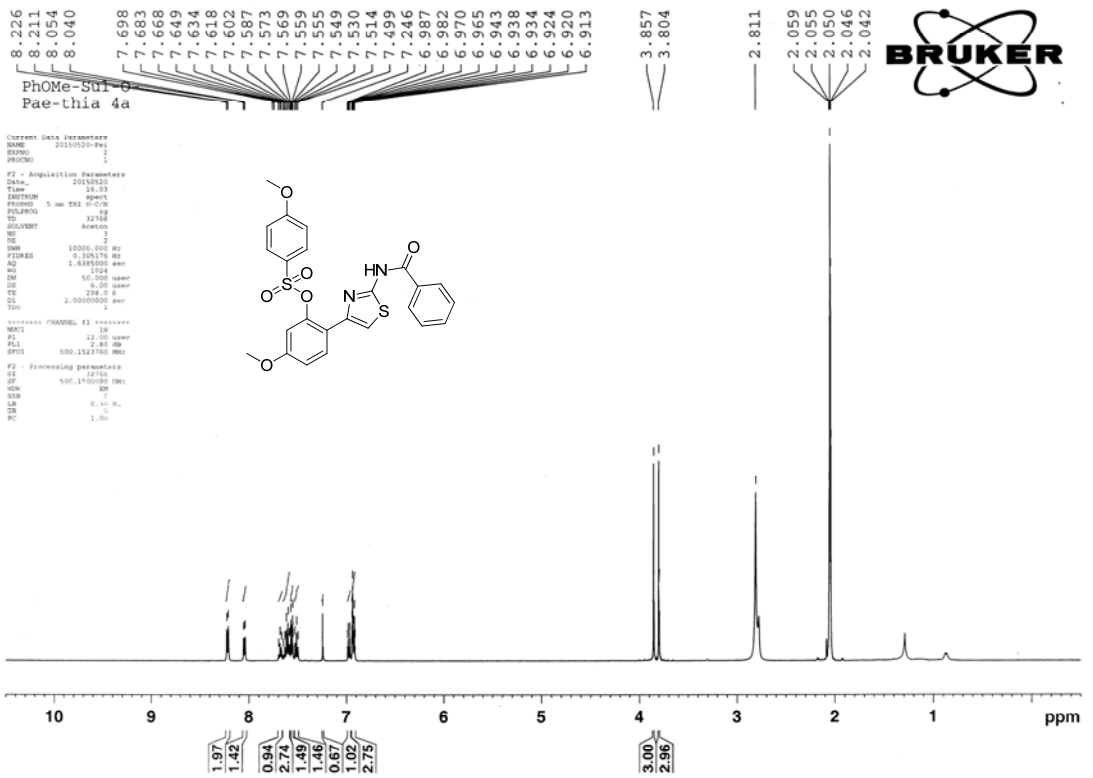
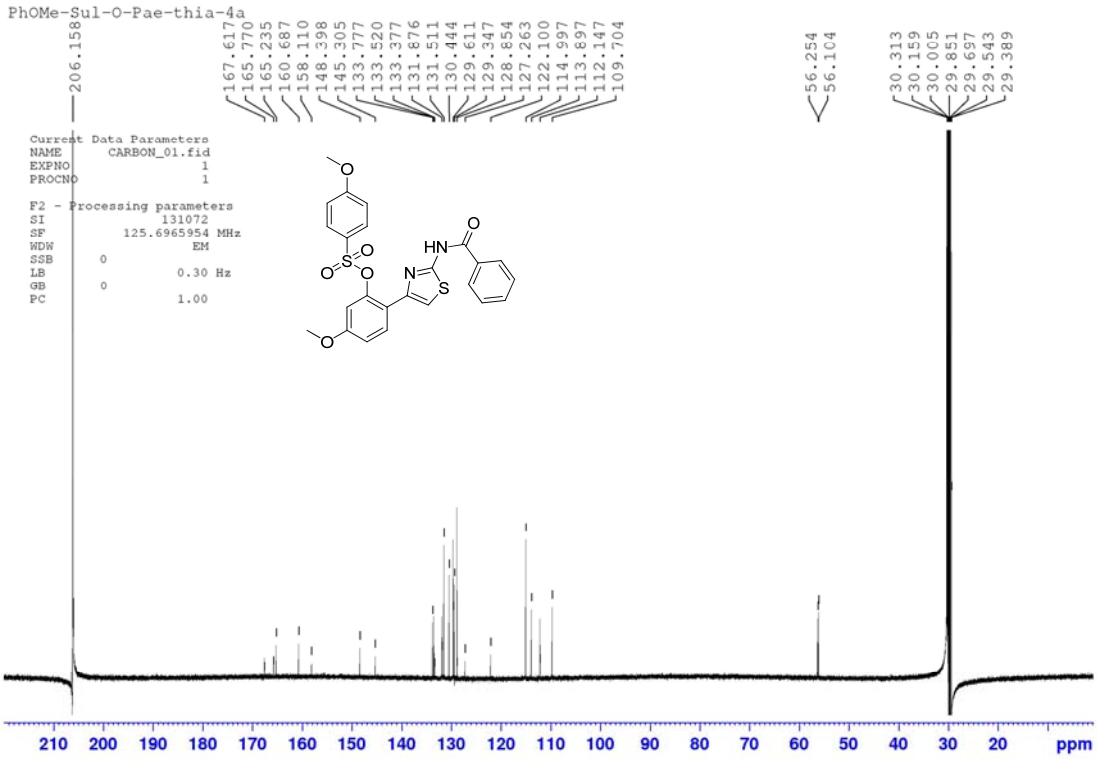
^{1}H and ^{13}C NMR of 8a





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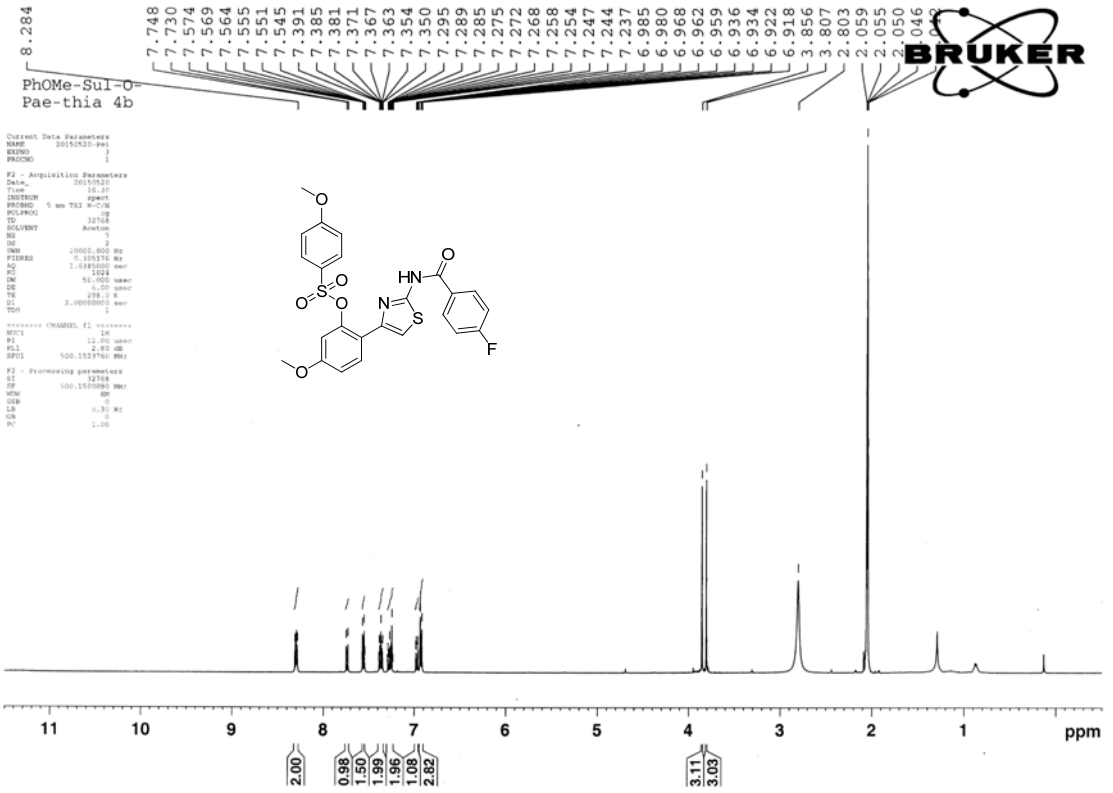

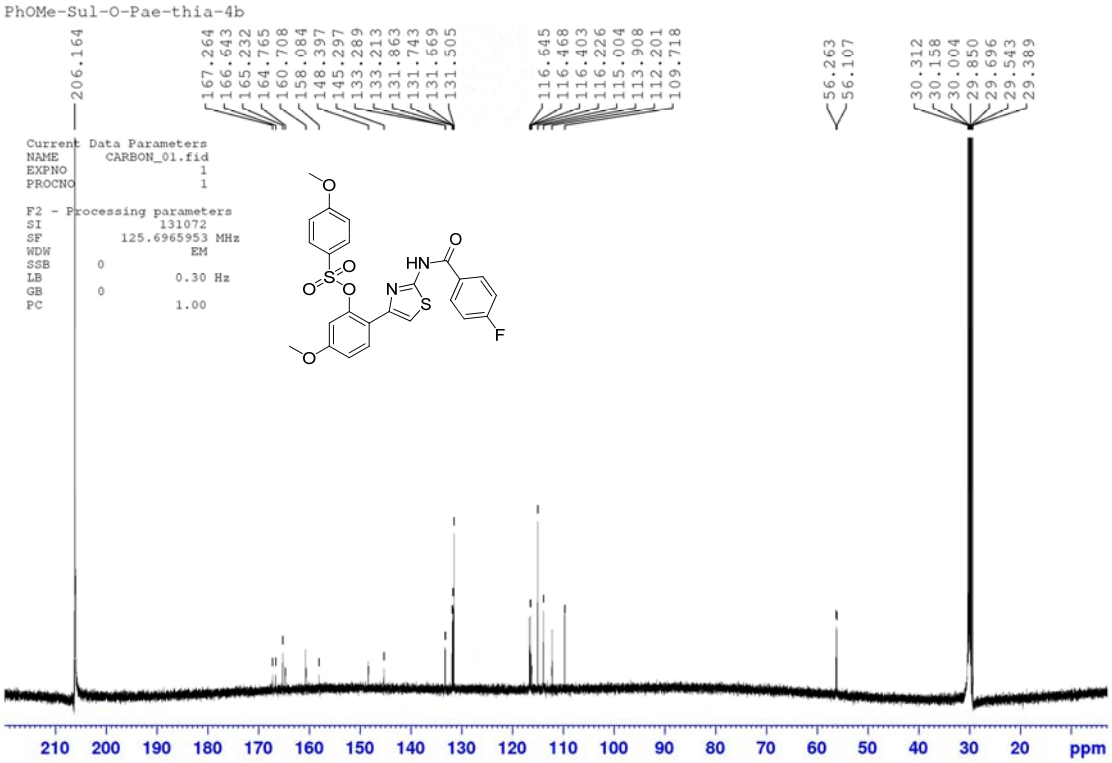

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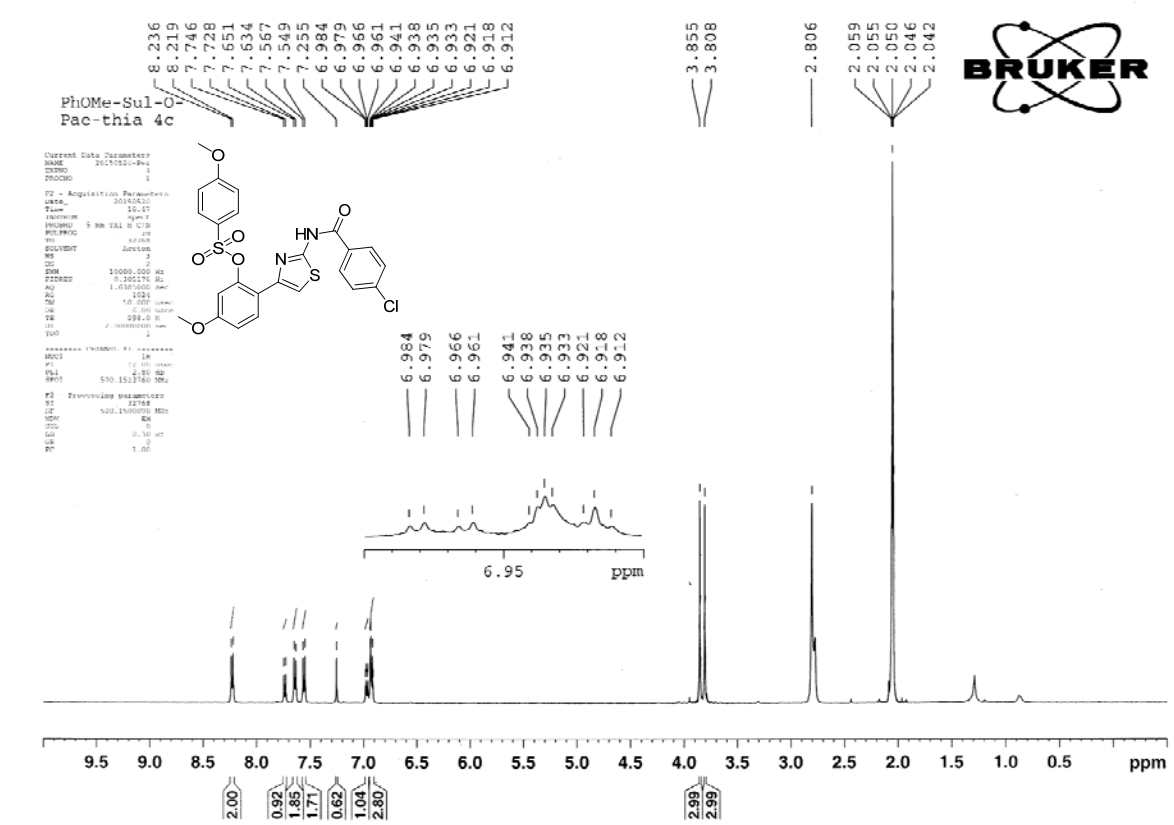

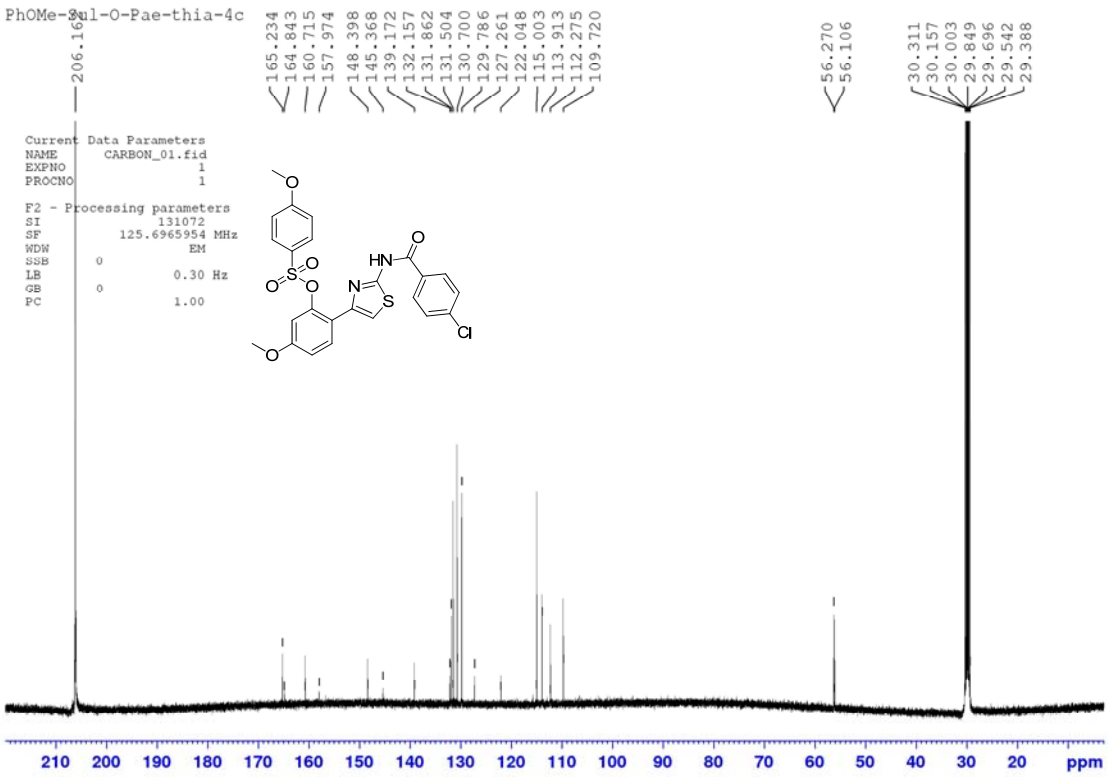

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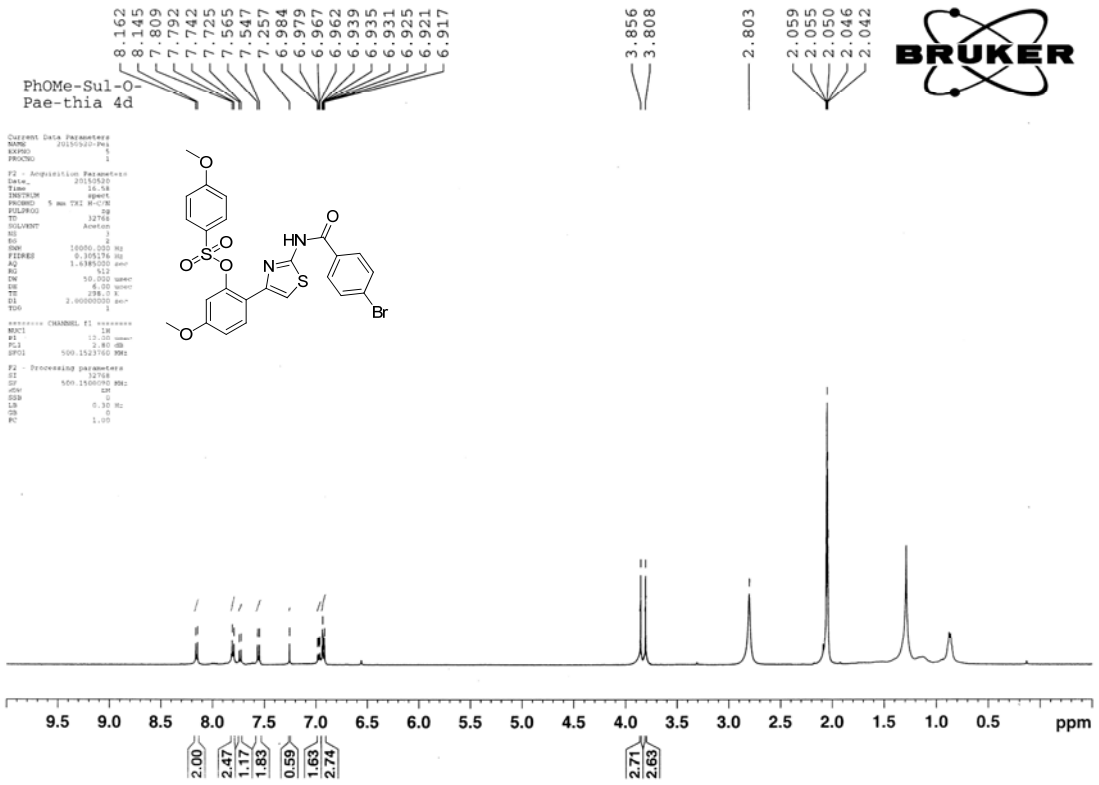

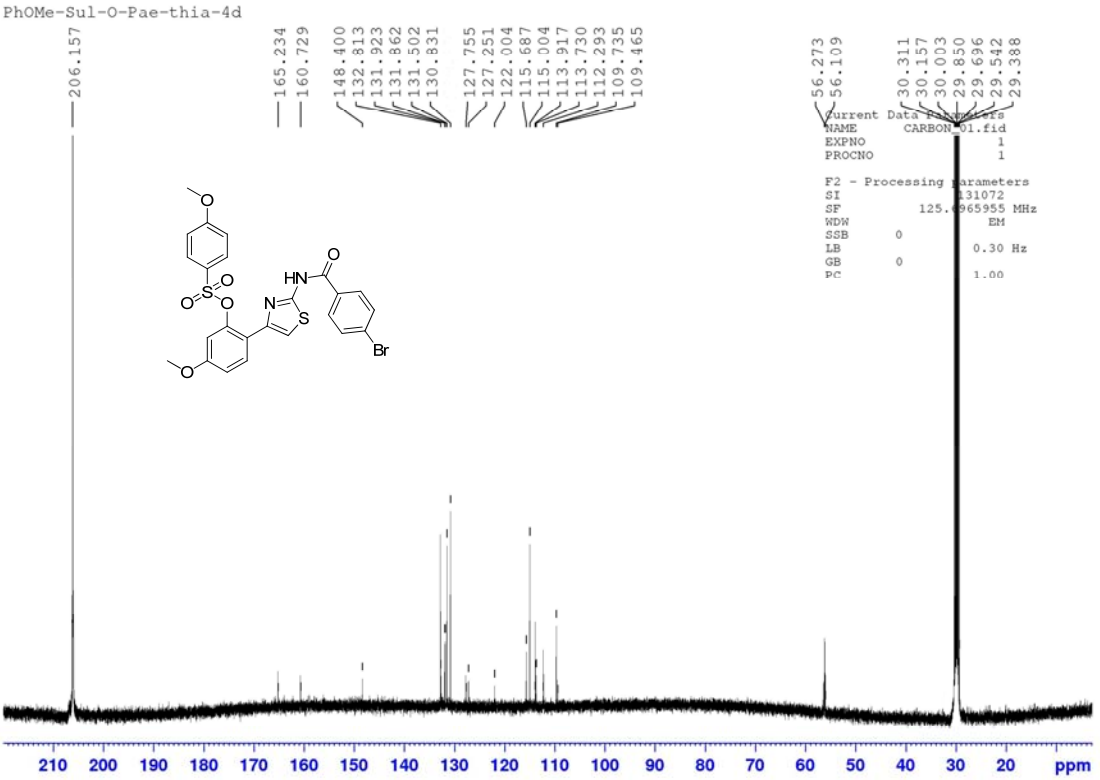

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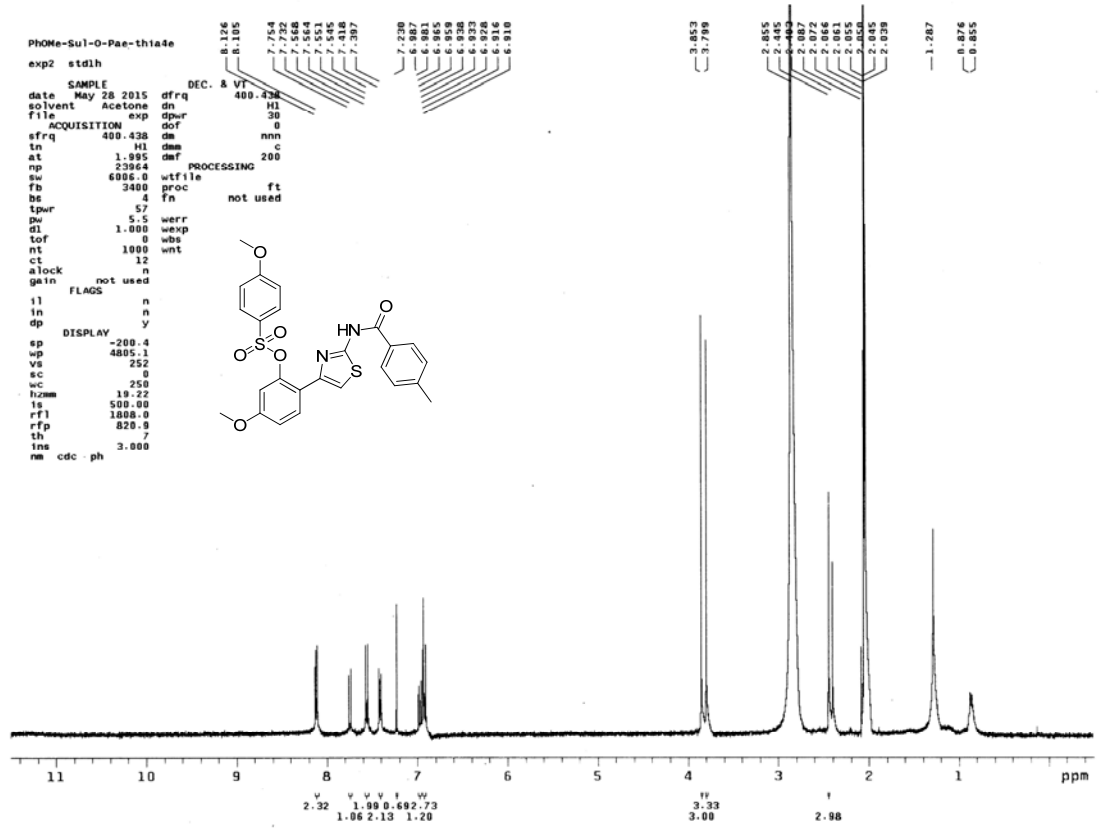

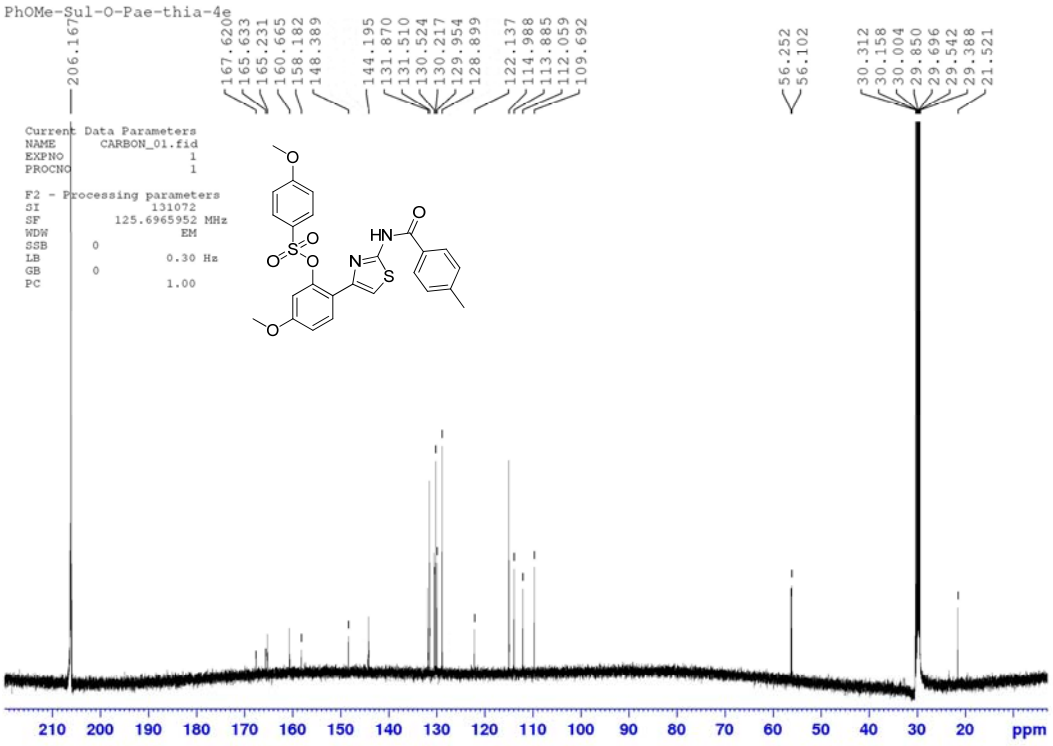

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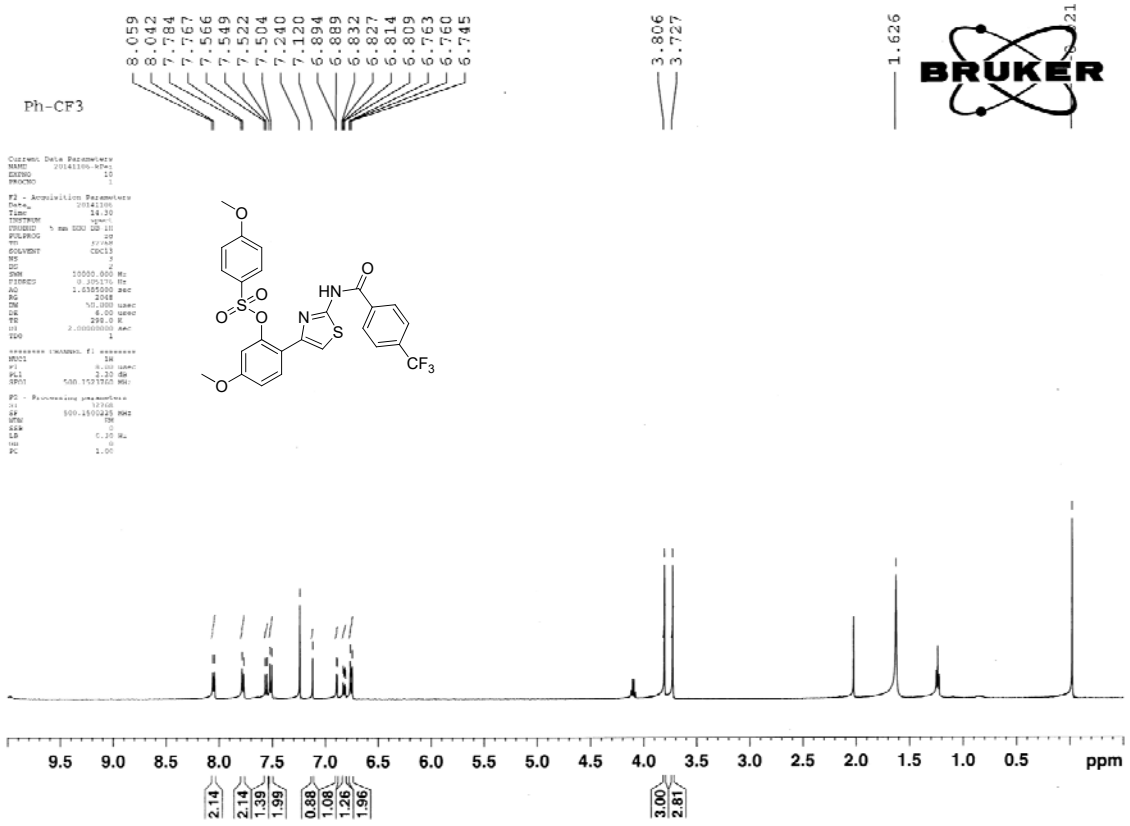
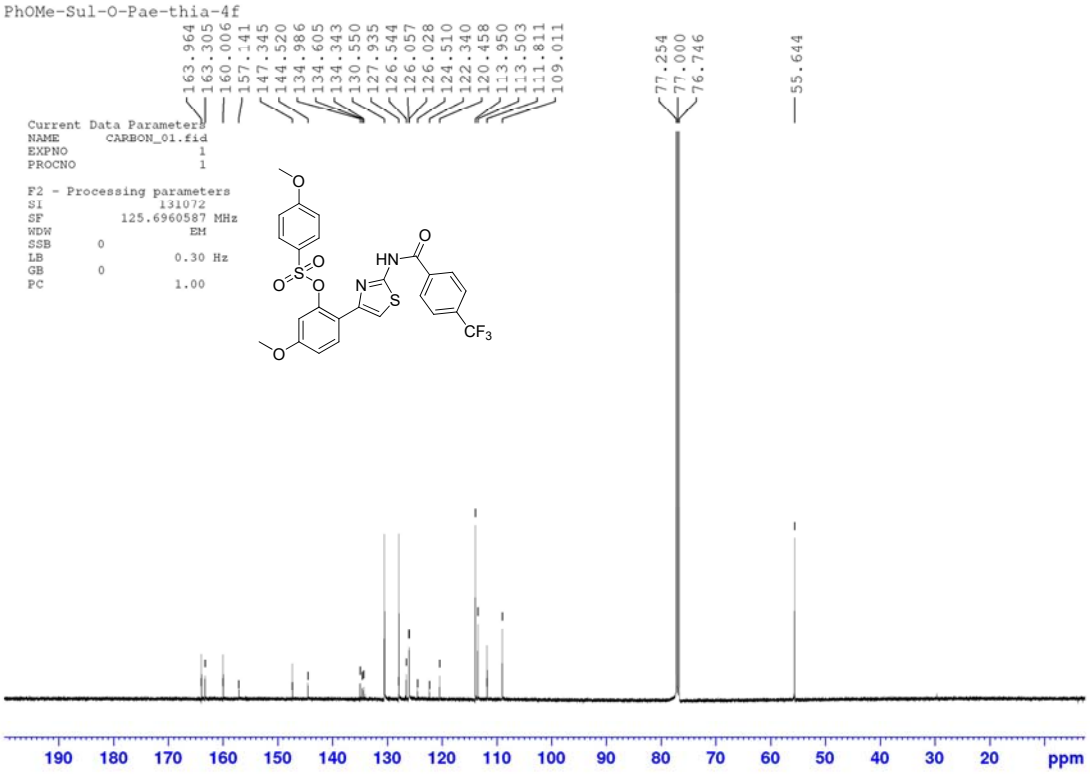
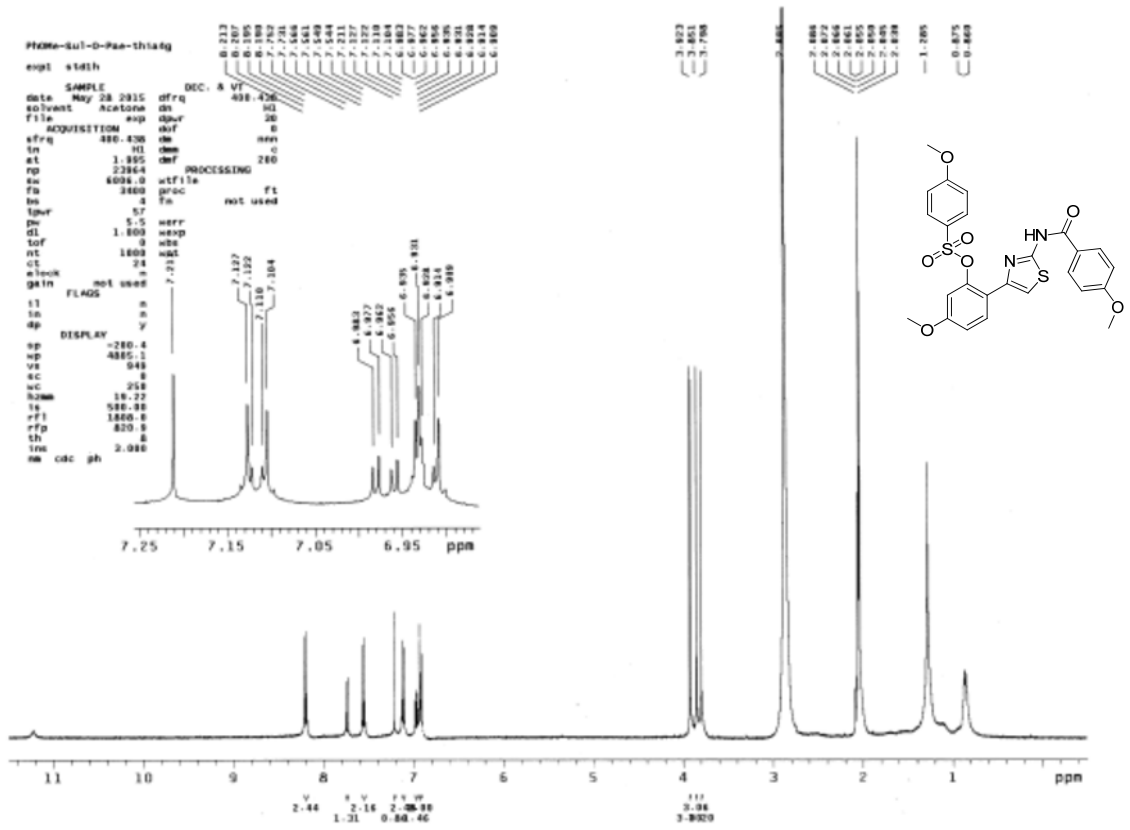
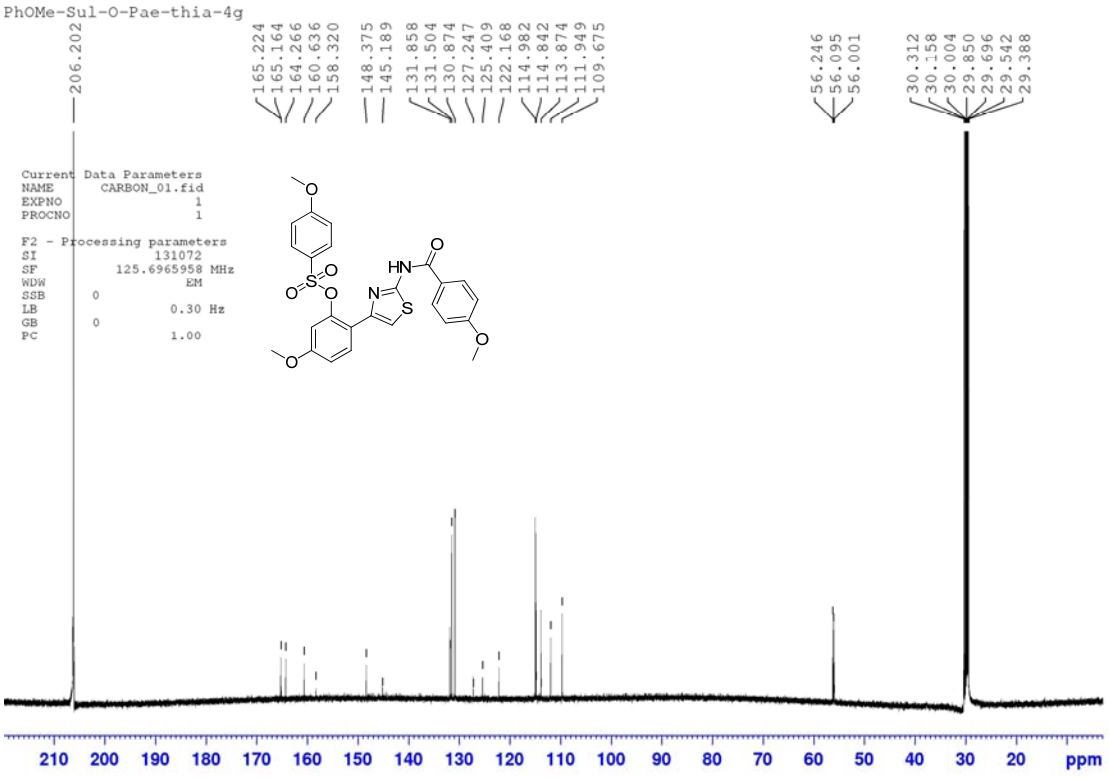


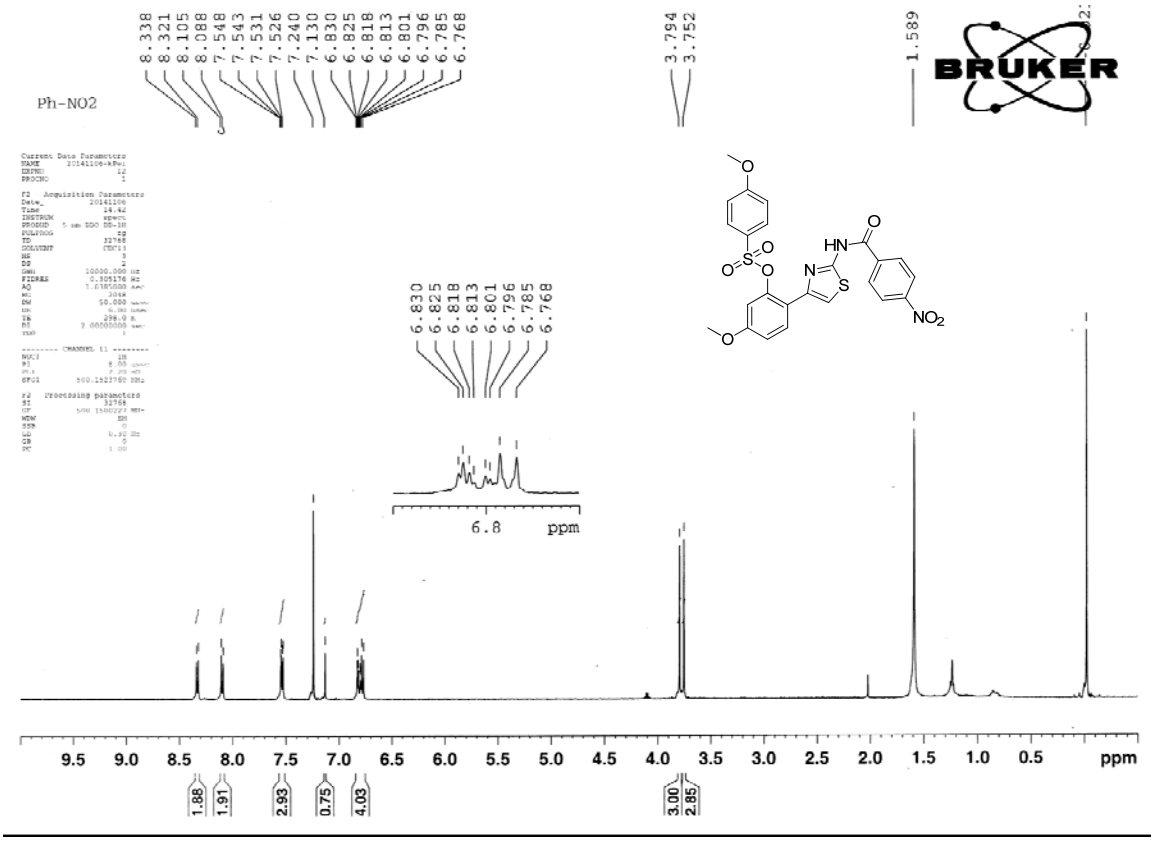


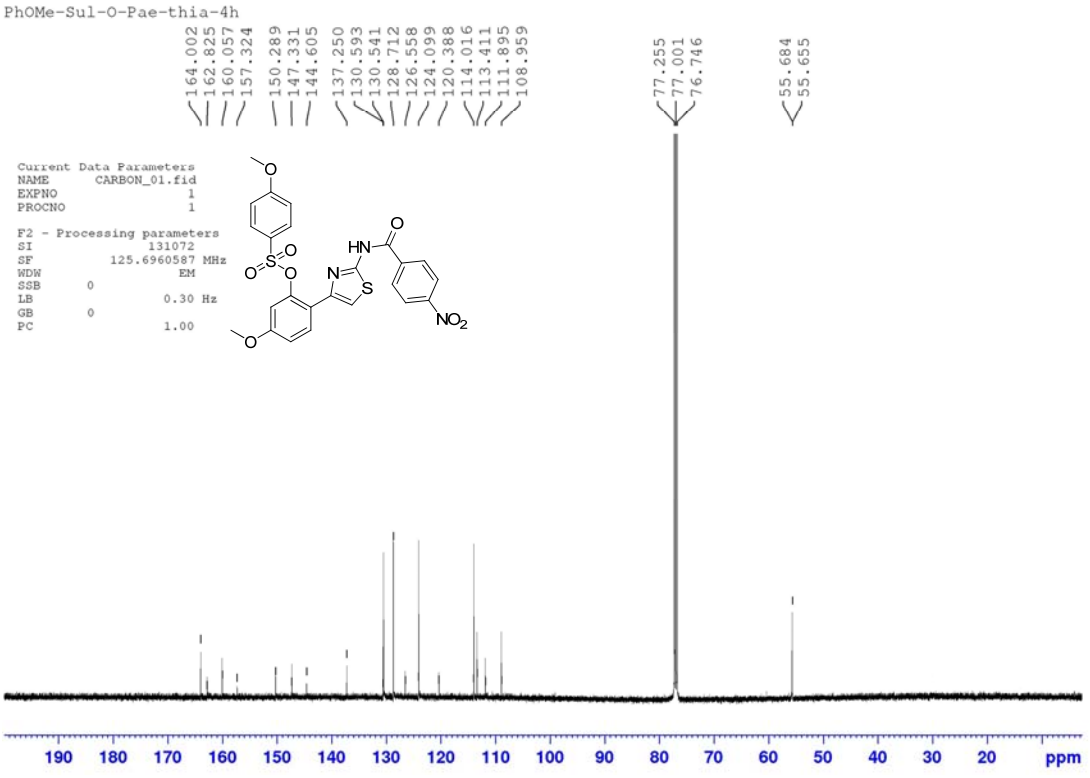
^{1}H and ^{13}C NMR of 8g



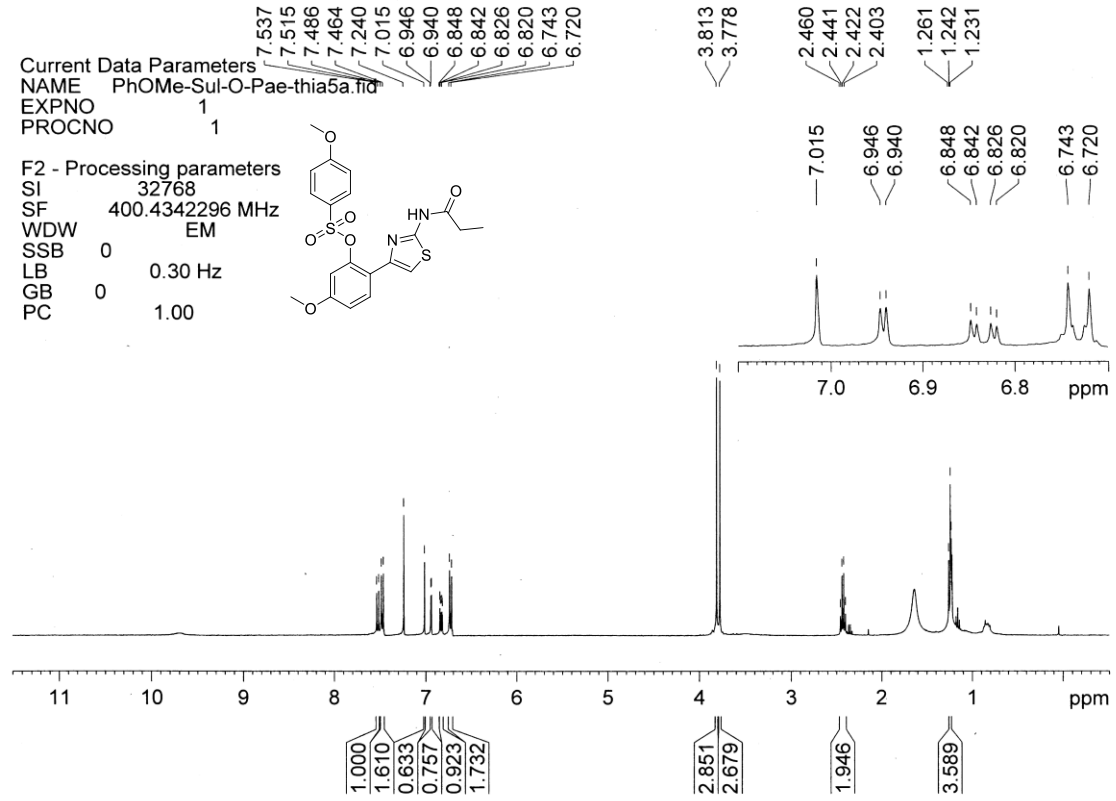


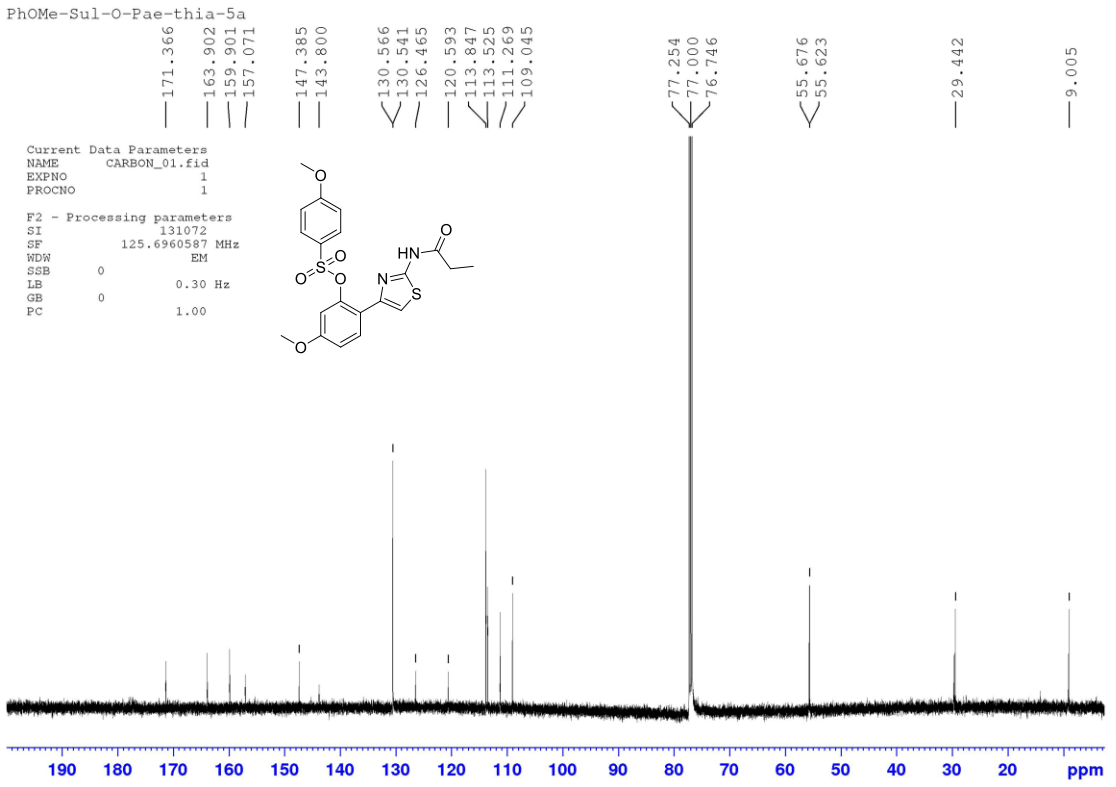
^{1}H and ^{13}C NMR of 8h





^{1}H and ^{13}C NMR of 9a





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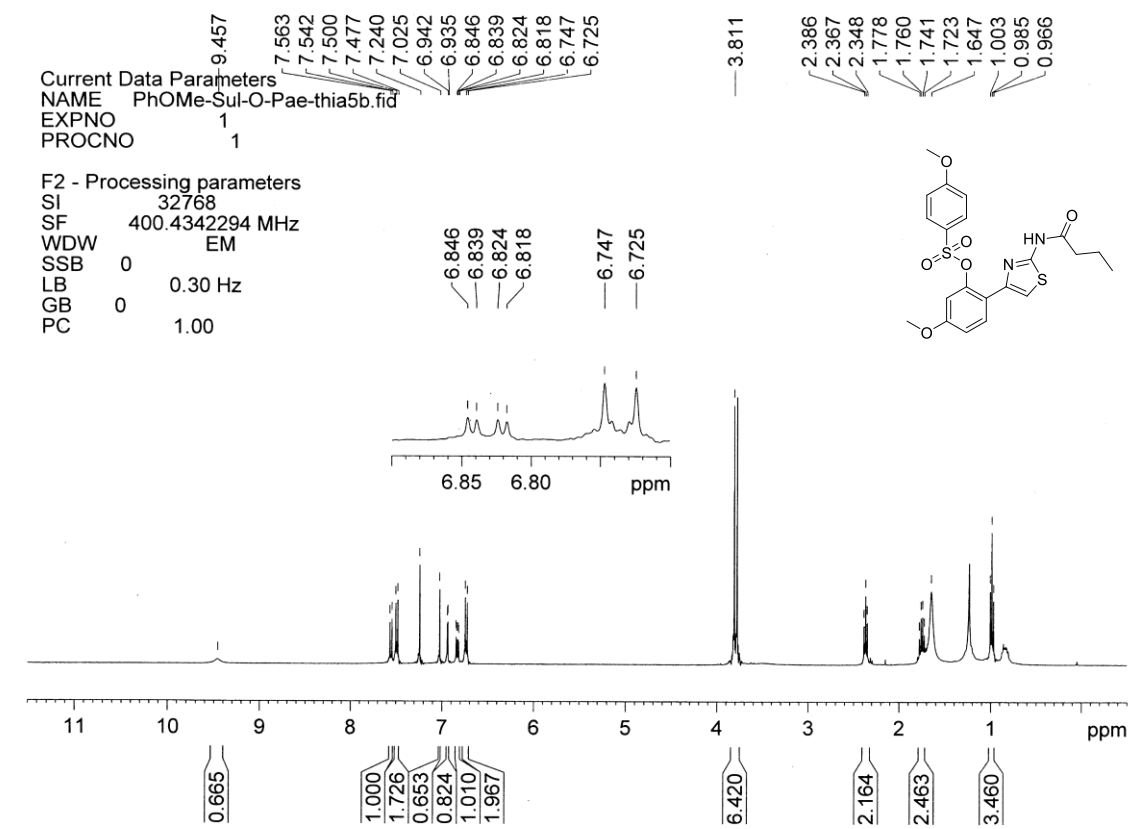

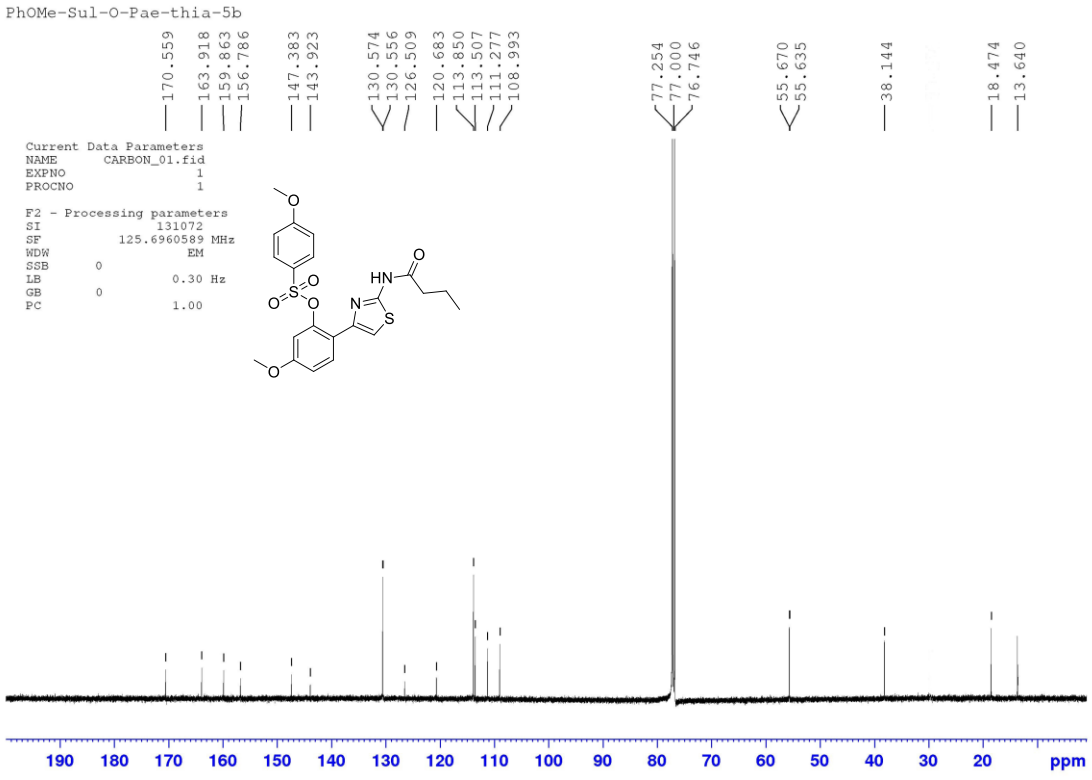

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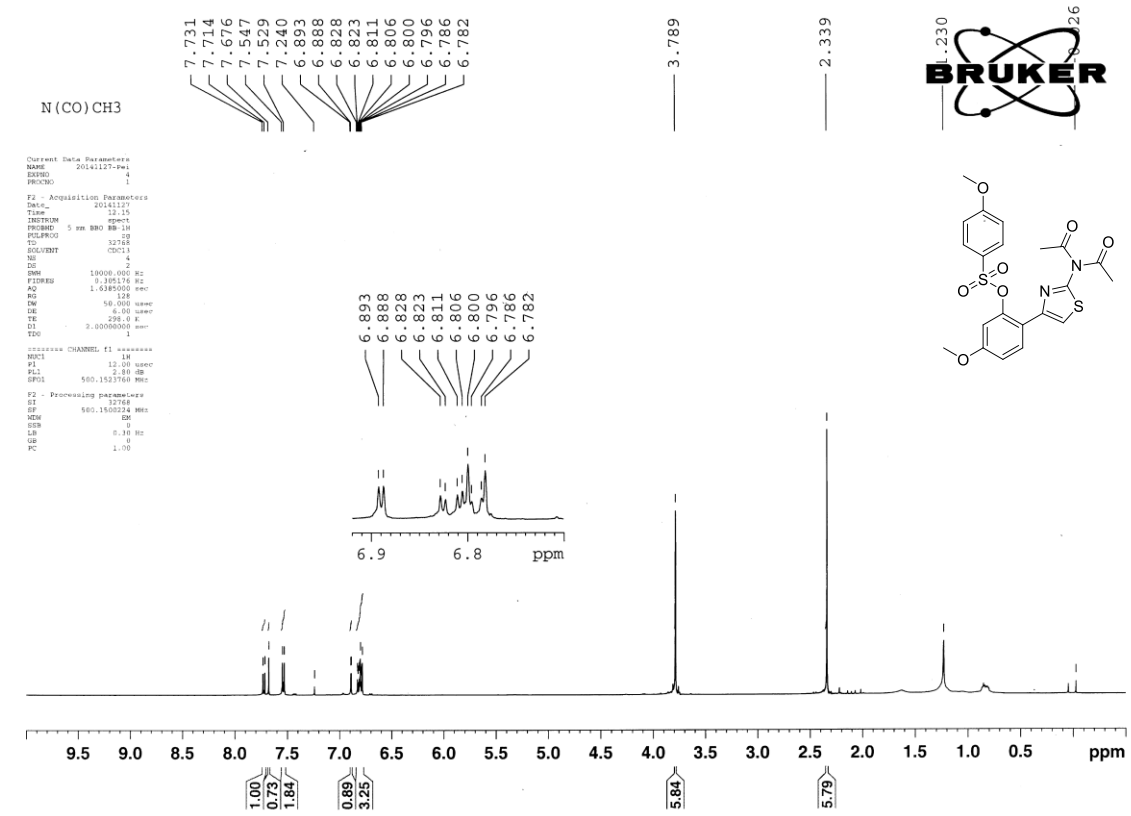
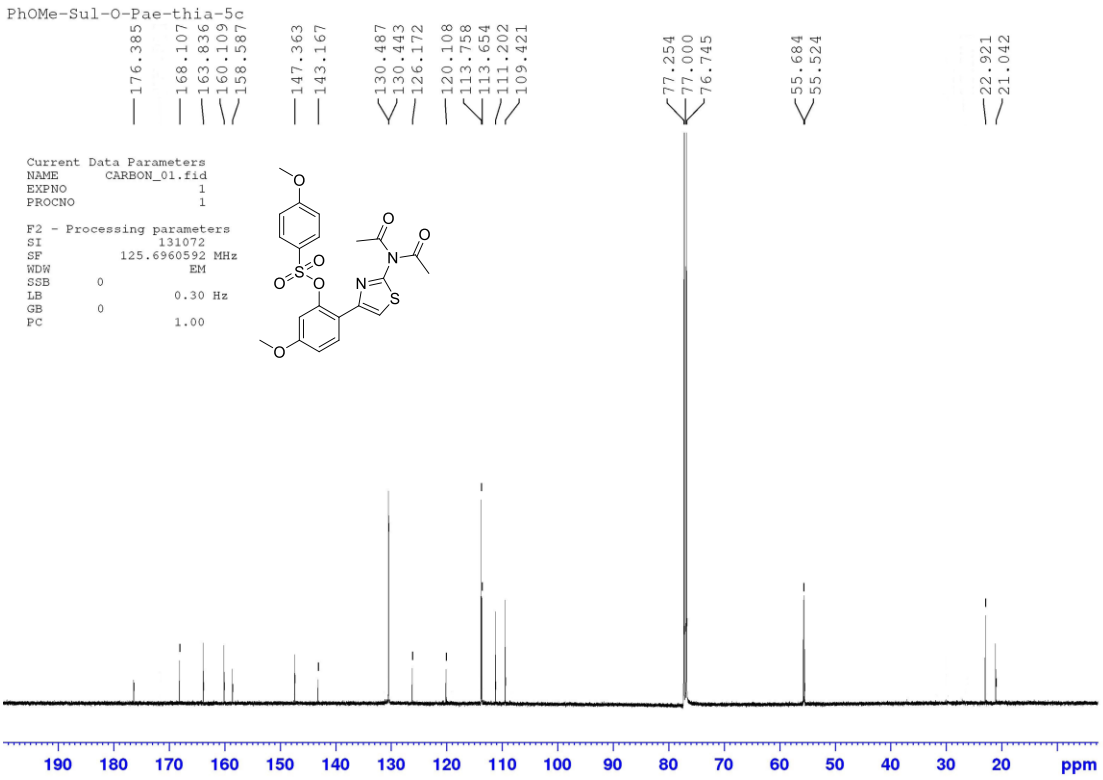
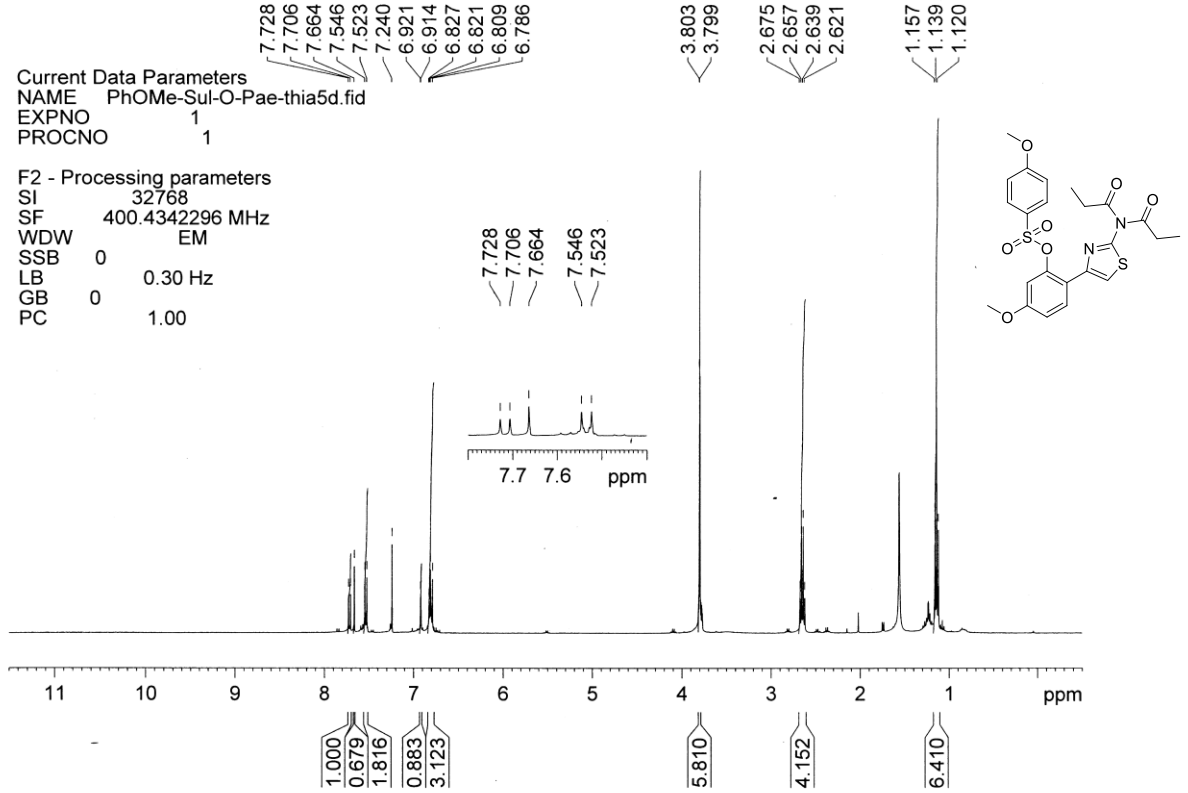
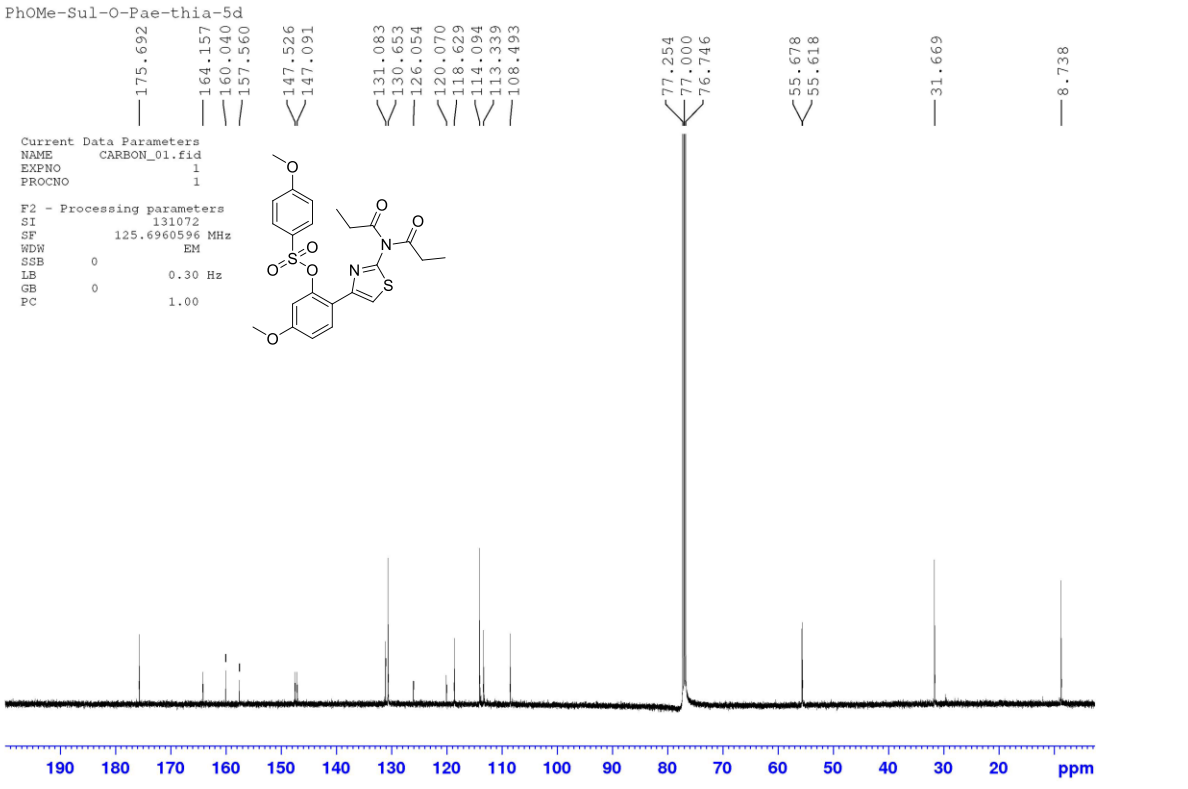




^{1}H and ^{13}C NMR of 9d





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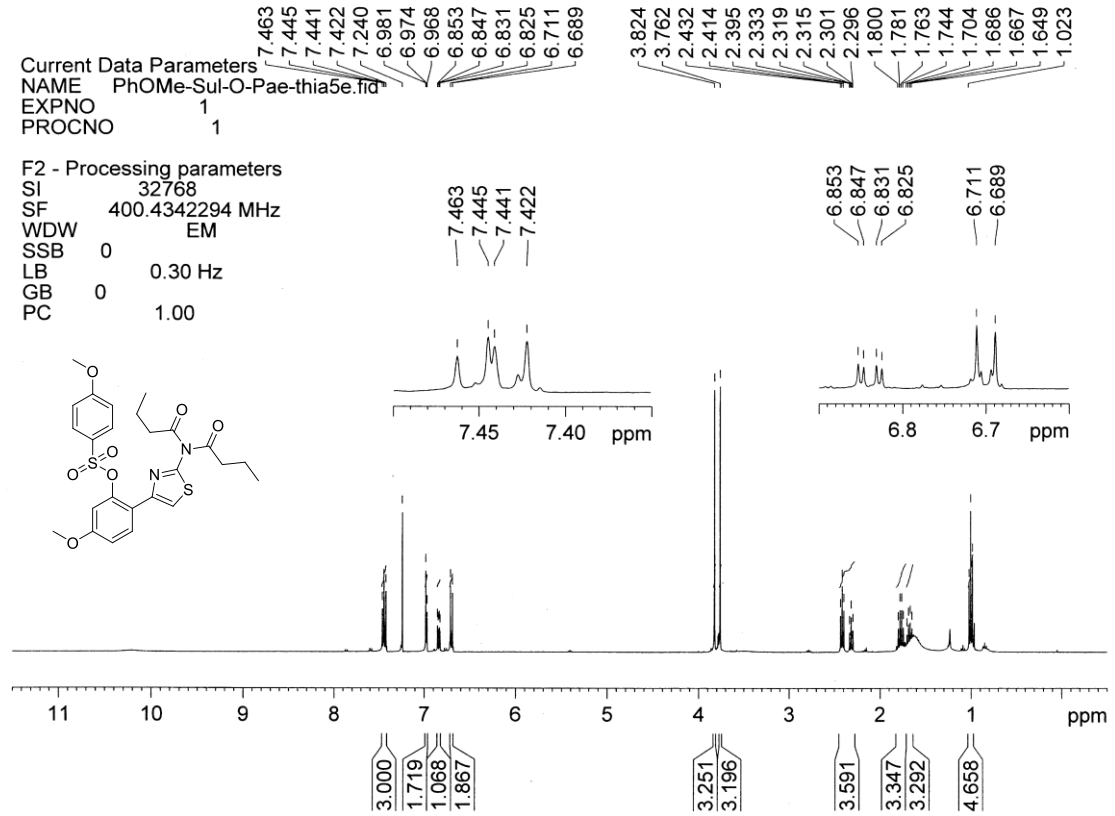
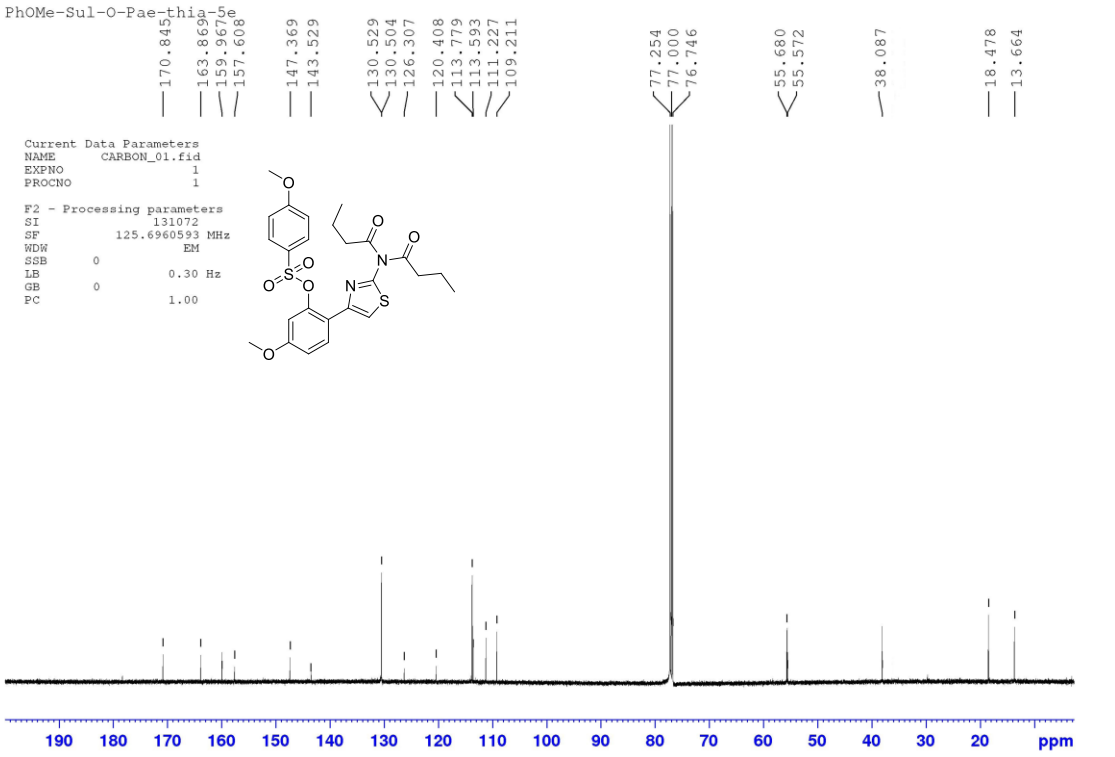
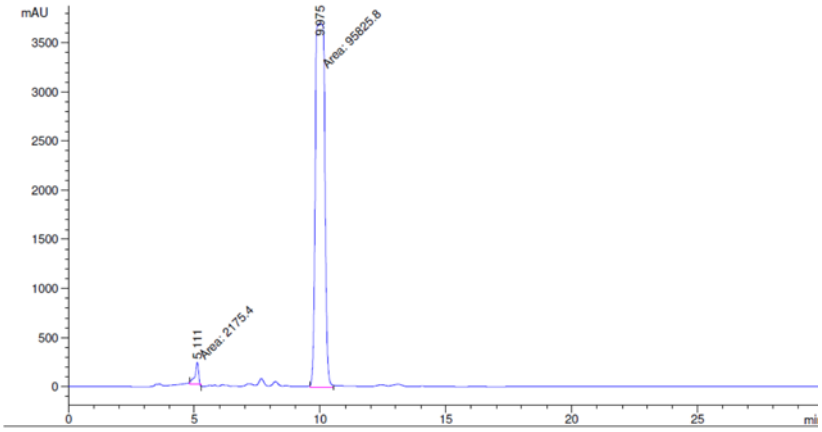




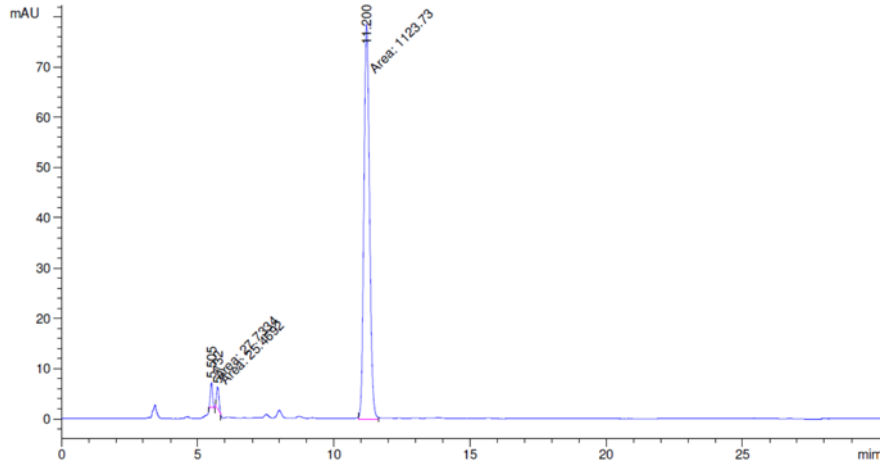
HPLC analysis of 8a



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.11	0.1632	2175.398	2.22
2	9.98	0.4316	9.58e4	97.78

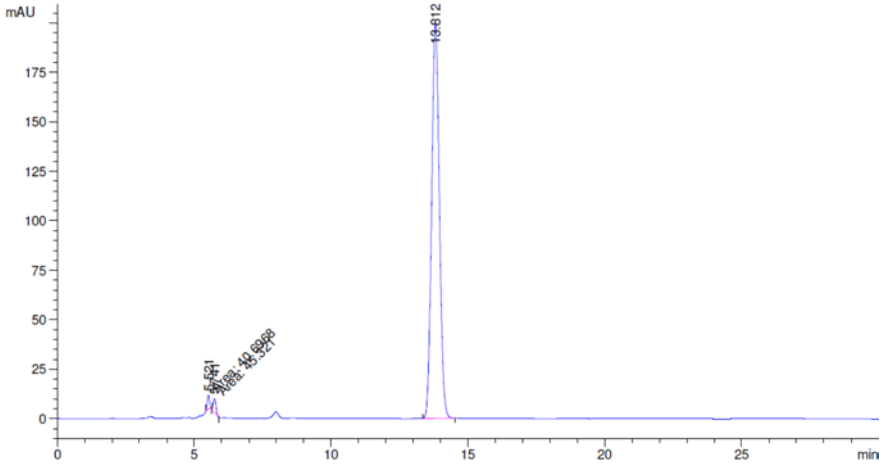
HPLC analysis of 8b



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.51	0.0938	27.733	2.36
2	5.73	0.0968	25.469	2.16
3	11.20	0.2383	1123.734	95.48

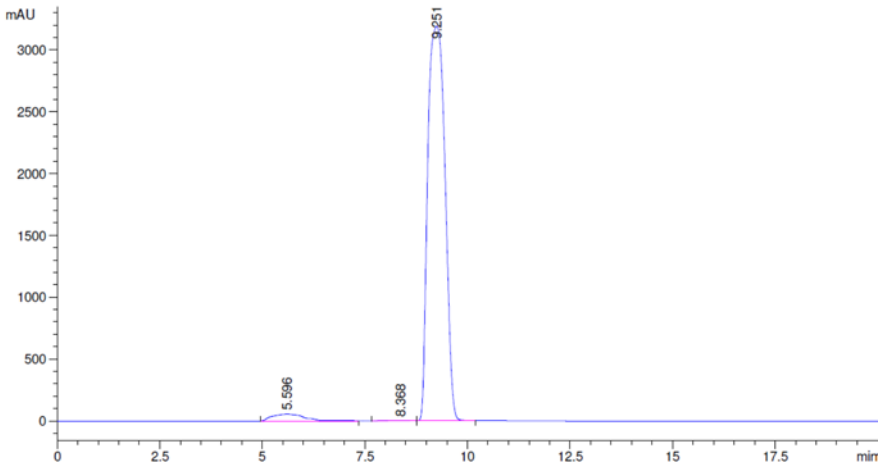
HPLC analysis of 8c



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.21	0.0960	40.697	1.02
2	5.74	0.1075	45.321	1.13
3	13.81	0.3081	3922.876	97.85

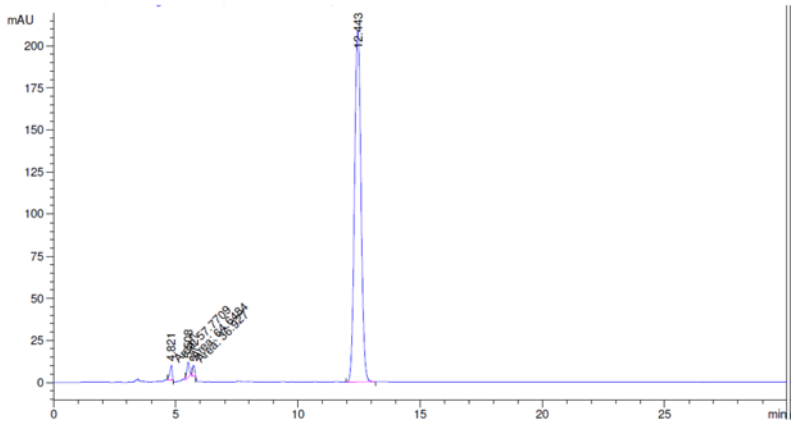
HPLC analysis of 8d



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.60	0.7979	2794.841	2.93
2	8.37	0.4470	54.367	0.06
3	9.25	0.4812	9.26e4	97.01

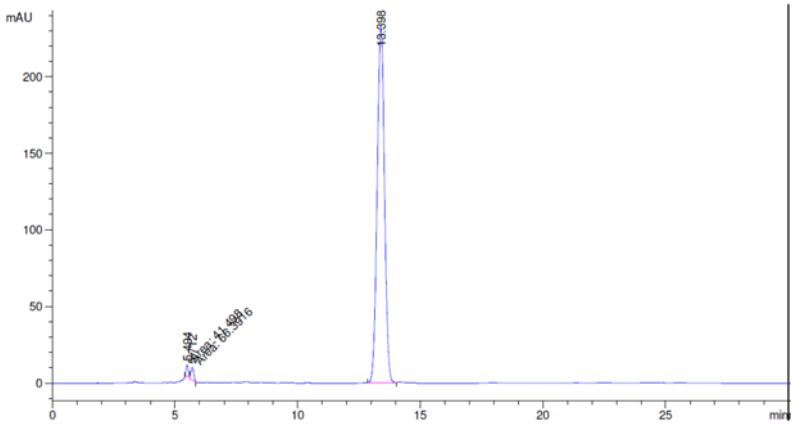
HPLC analysis of 8e



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	4.82	0.1099	57.771	1.44
2	5.51	0.1188	64.648	1.61
3	5.73	0.0959	36.927	0.91
4	12.44	0.2882	3862.217	96.04

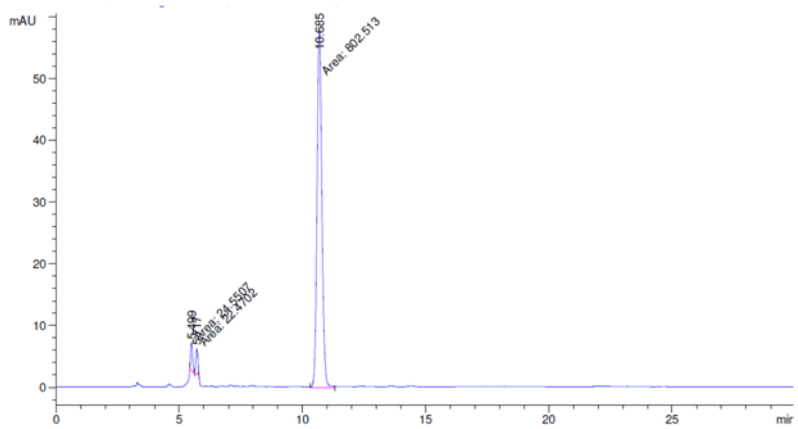
HPLC analysis of 8f



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.49	0.0962	41.498	0.84
2	5.71	0.1302	66.392	1.34
3	13.40	0.3276	4832.661	97.82

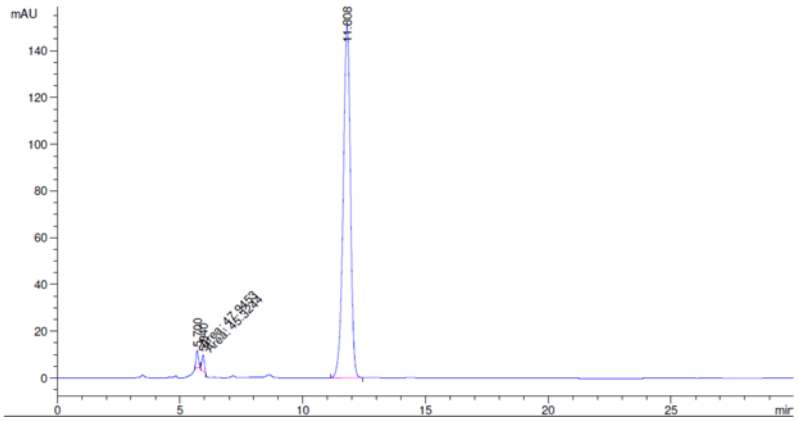
HPLC analysis of 8g



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.50	0.0905	24.551	2.80
2	5.72	0.0928	22.470	1.89
3	10.69	0.2315	802.513	95.31

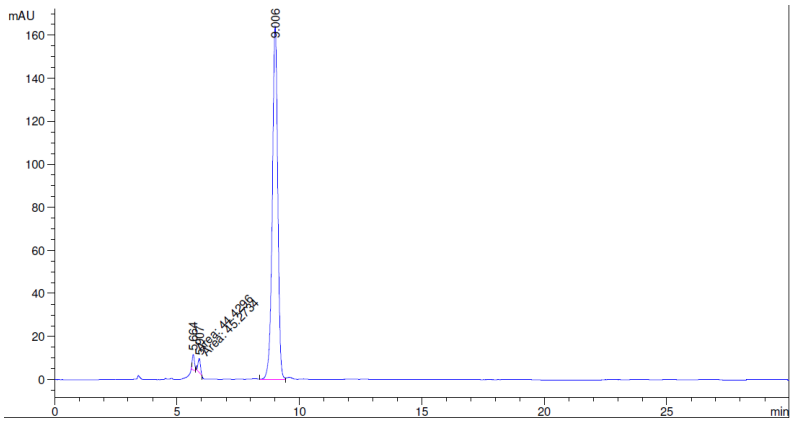
HPLC analysis of 8h



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.70	0.1072	47.945	1.51
2	5.94	0.1108	45.325	1.42
3	11.81	0.3157	3092.286	97.07

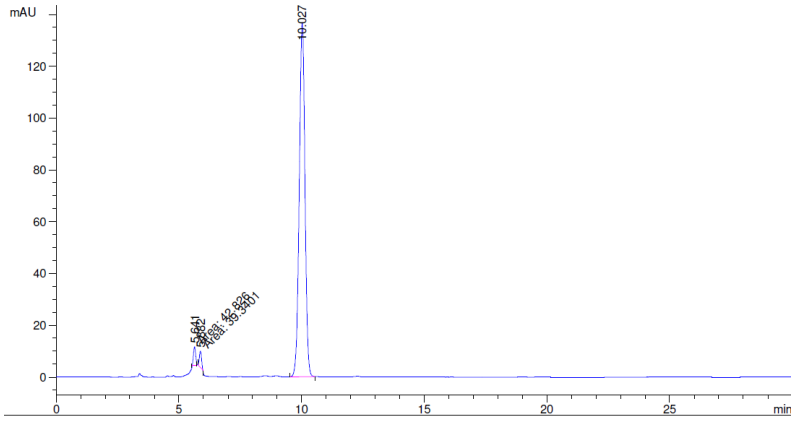
HPLC analysis of 9a



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.66	0.1022	44.430	1.70
2	5.91	0.1126	45.273	1.74
3	9.01	0.2345	2521.588	96.56

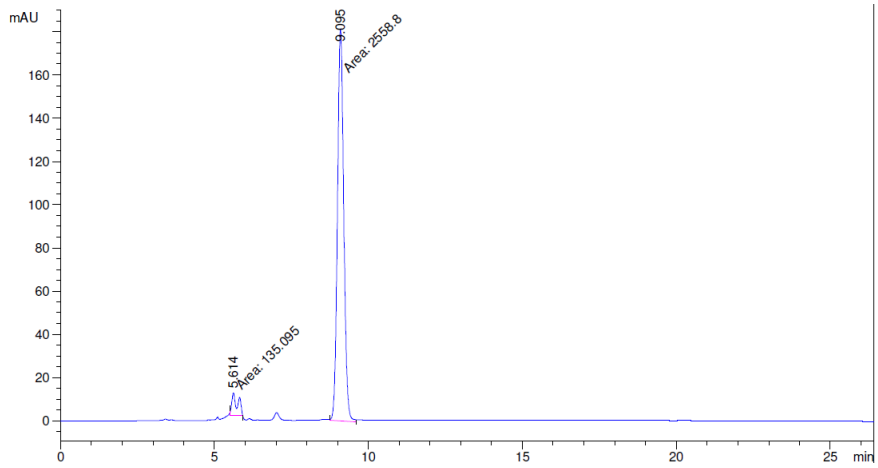
HPLC analysis of 9b



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.64	0.1006	42.826	1.87
2	5.88	0.1042	39.340	1.71
3	10.03	0.2501	2213.741	96.42

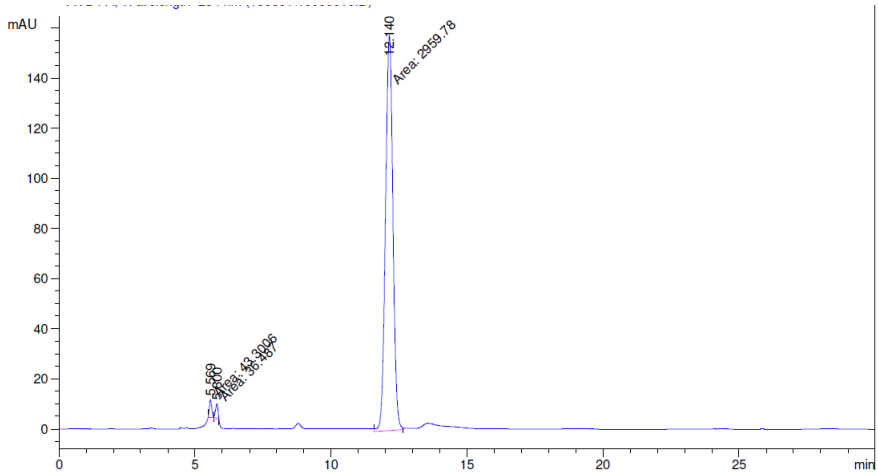
HPLC analysis of 9c



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.61	0.2178	135.095	5.01
2	9.10	0.2353	2558.805	94.99

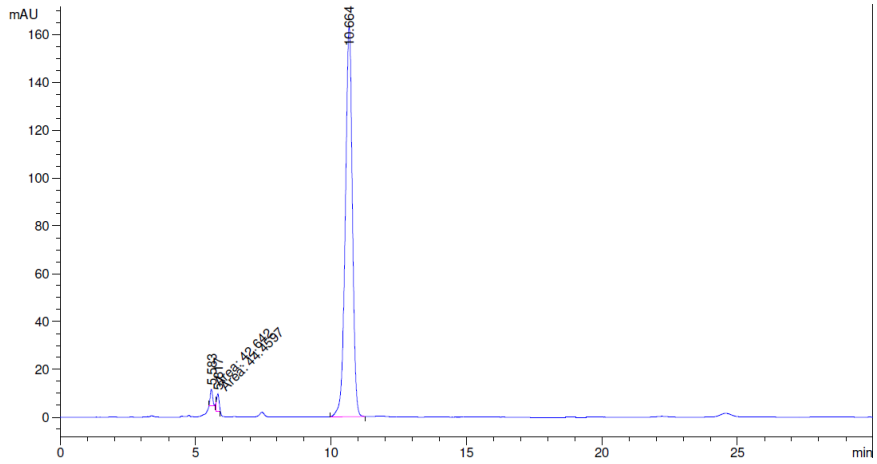
HPLC analysis of 9d



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.57	0.1014	43.301	1.42
2	5.80	0.1037	36.487	1.20
3	12.14	0.3133	2959.780	97.38

HPLC analysis of 9e



VWD1 A, Wavelength=254 nm; eluent: 20% DI water in acetonitrile; flow rate=0.5mL/min

Peak #	Ret Time (min)	Width (min)	Area mAu*s	Area %
1	5.58	0.1020	42.642	1.40
2	5.82	0.1030	44.460	1.46
3	10.66	0.2784	2961.159	97.14

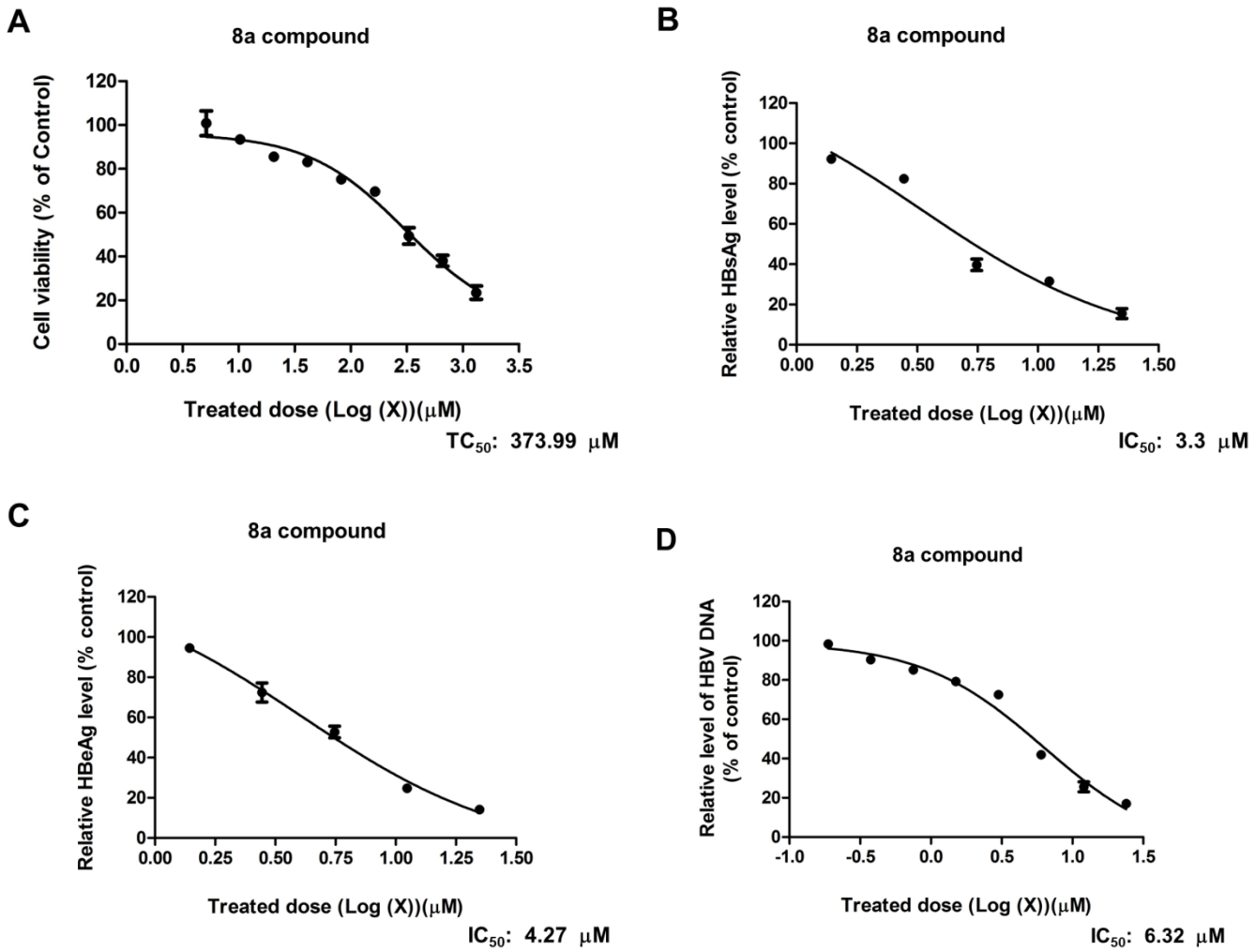
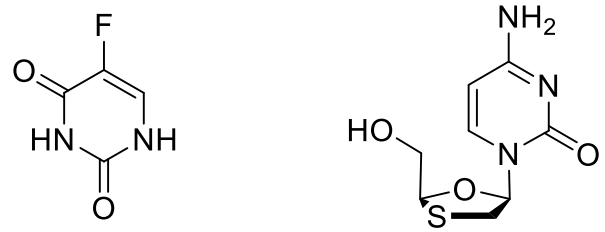


Figure S1 The representative dose-response graphic results of Paeonol-Phenylsufonyl derivative **8a** on (A) HepG2 2.15 cells' cytotoxicity, (B) hepatitis B virus (HBV) surface antigen (HBsAg) levels, (C) HB e antigen (HBeAg) levels, and (D) HBV viral DNA levels. The TC_{50} or IC_{50} value was analyzed with nonlinear regression fitting analysis using GraphPad Prism software (GraphPad Software Inc.). The presented values are known as mean**±SD.**



5-Fluorouracil (5-Fu) Lamivudine (3TC)

Figure S2 The structures of 5-Fluorouracil and Lamivudine, the reference compounds listed in Table 1.