

Supporting Information

**Asymmetric Fluorination of Indanone-2-carboxylates Using a
Polystyrene-supported Diphenylamine-linked Bis(oxazoline)
Complex**

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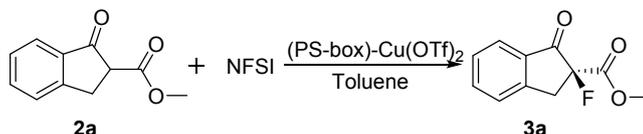
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1. Screening of different proportions of Cu(OTf)₂/1d in batch condition

Table S1 Catalytic performance of different proportions of Cu(OTf)₂/1d



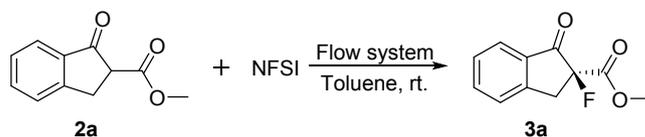
Entry	Cu(OTf) ₂ /1d	Yield[%] ^b	ee[%] ^c
1	0.5	95	91
2	0.6	96	92
3	0.7	95	92
4	0.8	96	95
5	0.9	96	94
6	1.0	97	92
7	1.2	94	90
8	1.5	96	88
9 ^d	-	76	0
10 ^{d, e}	-	90	0

^a Typical reaction conditions: to a tube a mixture of PS-box **1d** (0.005mmol, 0.2 mol%) and Cu(OTf)₂ with 25.0 mL toluene were added, then **2a** (2.5 mmol, 1.0 equiv.) and NFSI (1.2 equiv.) were added successively. The reaction mixture was shaken for 120 min. ^b Isolated yields. ^c Determined by chiral HPLC. ^d Without PS-box and Cu(OTf)₂. ^e The reaction mixture was shaken for 240 min.

Before the screening of solvent and catalyst loading, different proportions Cu(OTf)₂/1d were also carried out in batch condition. As can be seen in Table S1, the optimum proportion of Cu(OTf)₂/1d was 0.8, which achieved 97% yield and 96% ee. To be noted, the background reaction occurred obviously without the catalyst.

2. Optimization of flow rate

Table S2 Optimization of the fluorination of 1-indanone-2-carboxylate **2a** catalyzed by **1d** in continuous flow condition ^a



Entry	Flow rate (μL min ⁻¹)	Yield[%] ^b	ee[%] ^c
1	10	100	97
2	20	98	96
3	30	99	96
4	40	96	95
5	50	94	94

6	60	90	95
7	70	87	93

^a Typical continuous flow conditions: the reactions were performed with PS-box **1d** (0.068 mmol, 0.12 g) and celite (0.8 g) in a column which was charged with Cu(OTf)₂ at room temperature, **2a** (0.2 M, 1.0 equiv.) and NFSI (0.24 M, 1.2 equiv.) were pumped into the column from respective toluene solutions. The same PS-box was used for all the substrates. ^b Isolated yield. ^c Determined by chiral HPLC.

As can be seen in Table S2, the proper flow rate is 30 μL min⁻¹, which achieved 99% yield and 96% ee.

3. Long period experiment in toluene

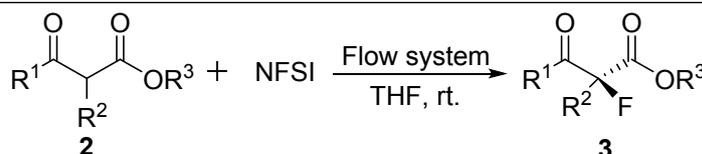
Table S3 Long period experiment in toluene^a

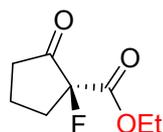
Time(h)	Yield[%] ^b	ee[%] ^c	Time(h)	Yield[%] ^b	ee[%] ^c
12-12.5	-	-	60-60.5	97	97
24-24.5	97	96	72-72.5	96	96
36-36.5	98	96	84-84.5	97	95
48-48.5	95	95	96-96.5	98	96

^a The reactions were performed with PS-box **1d** (0.12 g, 0.068 mmol) and celite (0.8 g) in a column which was charged with Cu(OTf)₂ at room temperature, and then **2a** (0.2 M, 1.0 equiv.) and NFSI (0.24 M, 1.2 equiv.) were pumped into the column at 30 μL min⁻¹ from respective toluene solutions (60 μL min⁻¹ total flow rate, 0.1 M concentration of **2a** in the reactor). ^b Isolated yields. ^c Determined by chiral HPLC.

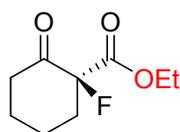
4. Scope of the enantioselective fluorination of aliphatic cyclic and acyclic β-keto esters

Table S4 Scope of the enantioselective fluorination of aliphatic cyclic and acyclic β-keto esters in continuous flow with immobilized catalyst **1d**^a and in batch condition with its parent ligand **1f**^b

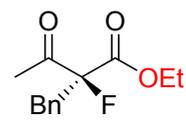




3p
^a17 yield, rac.
^b20 yield, rac.



3q
^atrace, nd
^btrace, nd



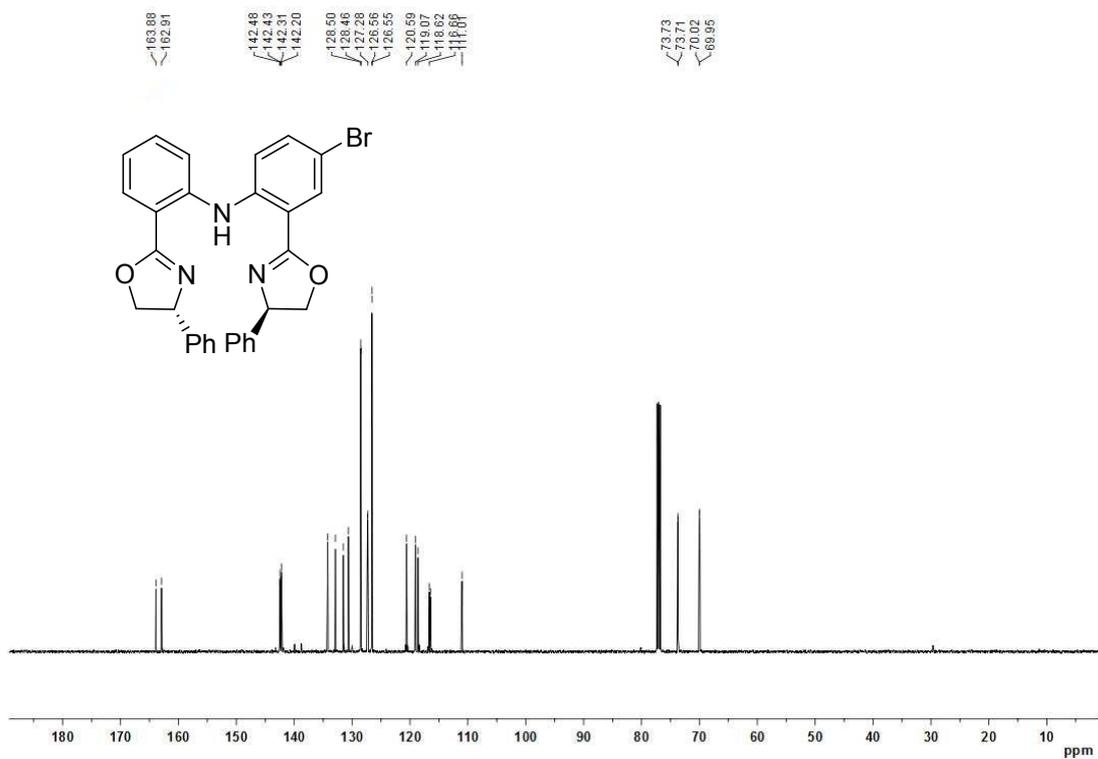
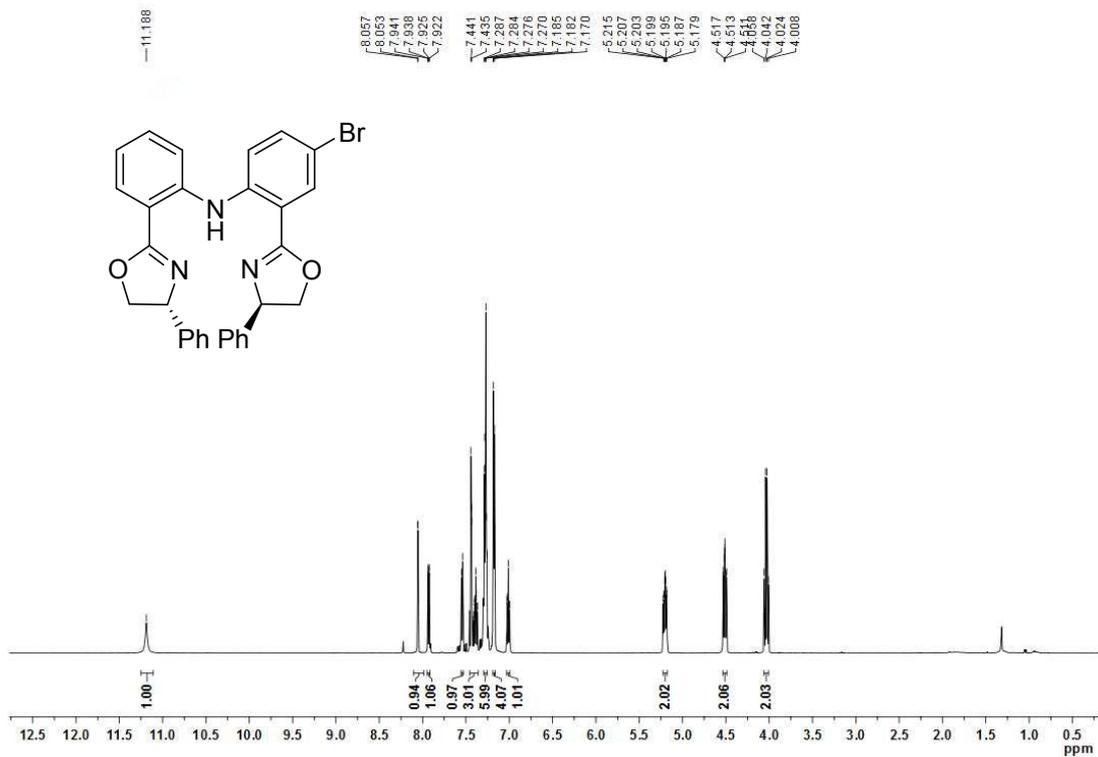
3r
^atrace, nd
^btrace, nd

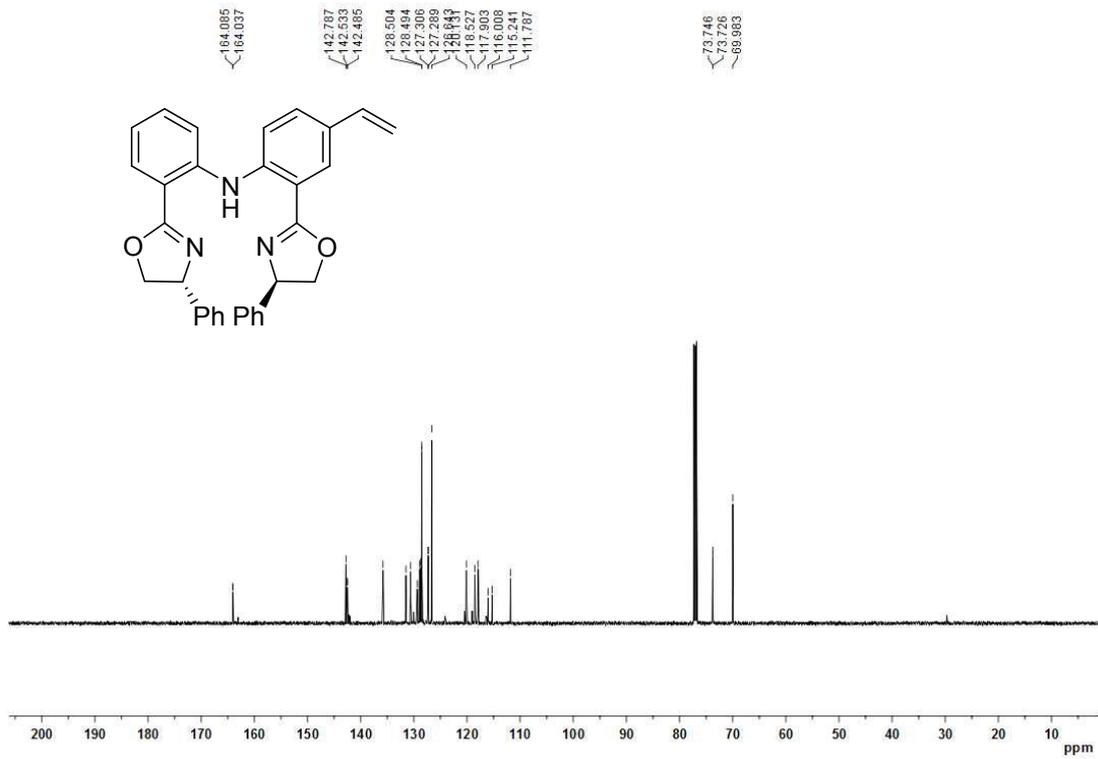
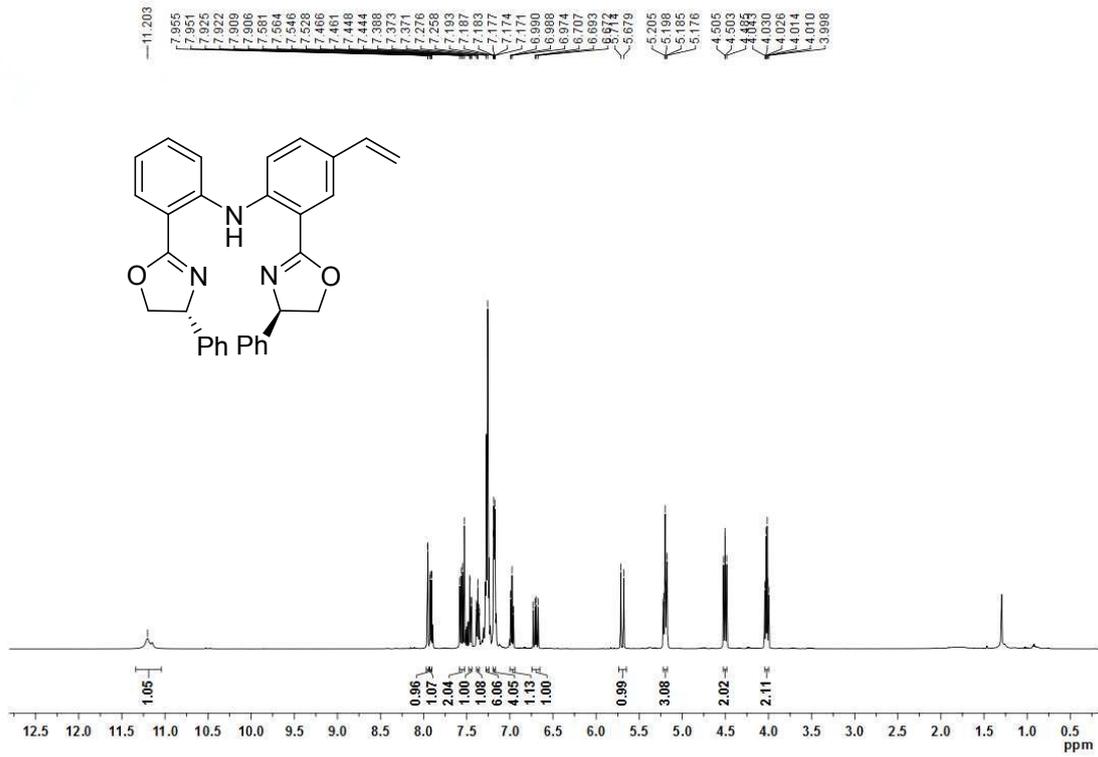
^a Typical continuous flow conditions: the reactions were performed with PS-box **1d** (0.12 g, 0.068 mmol) and celite (0.8 g) in a column which is charged with Cu(OTf)₂ at room temperature, and then **2** (0.2 M, 1.0 equiv.) and NFSI (0.24 M, 1.2 equiv.) were pumped into the column at 30 μL min⁻¹ from respective THF solutions (60 μL min⁻¹ total flow rate, 0.1 M concentration of **2** in the column). Each substrate carried out for 12 h. The same PS-box was used for all the substrates. Residence time under these conditions was 28 min. ^b Typical batch conditions: to a tube a complex of **1f**-Cu(OTf)₂ (0.05 mmol) with 5.0 mL toluene were added, and then **2** (0.5 mmol, 1.0 equiv.) and NFSI (0.6 mmol, 1.2 equiv.) were added into the tube. The reaction mixture was shaken for 12 h.

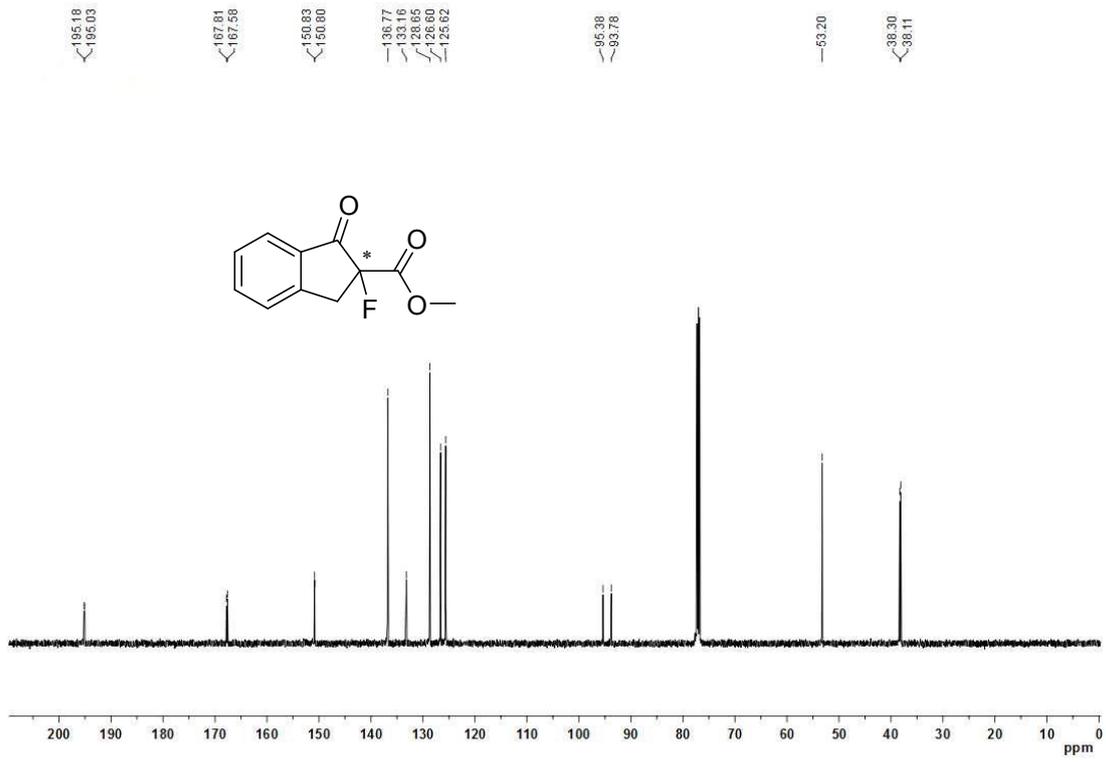
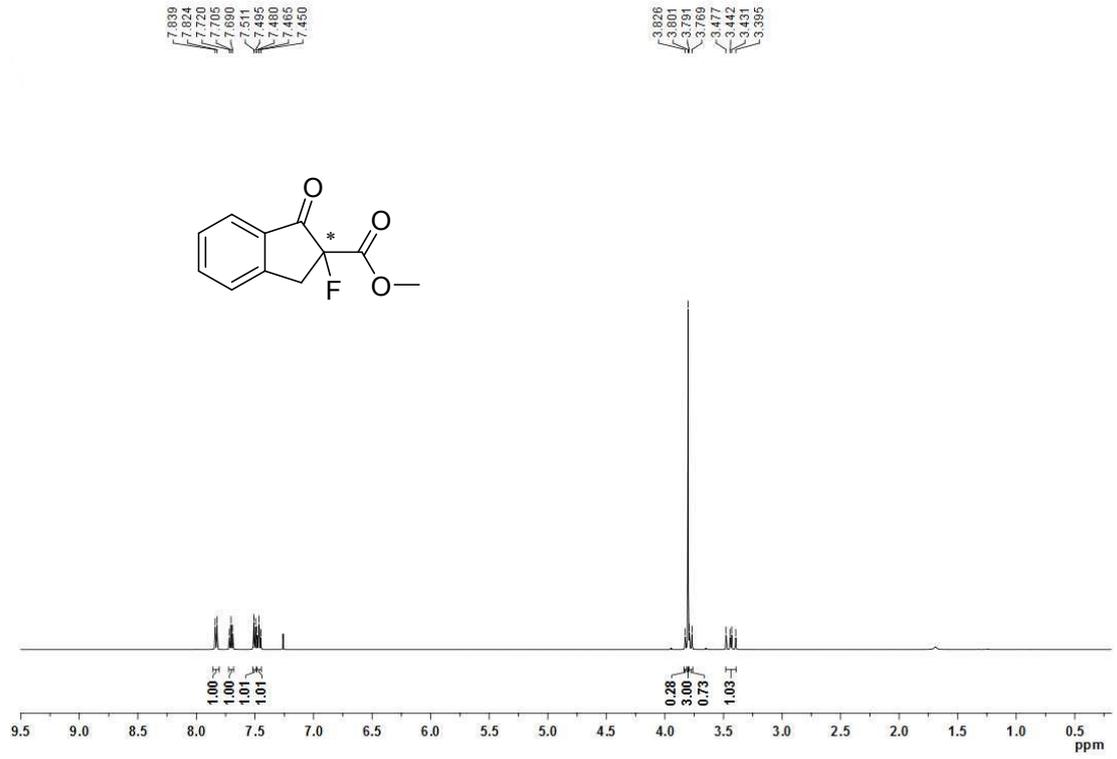
5. Characterization results

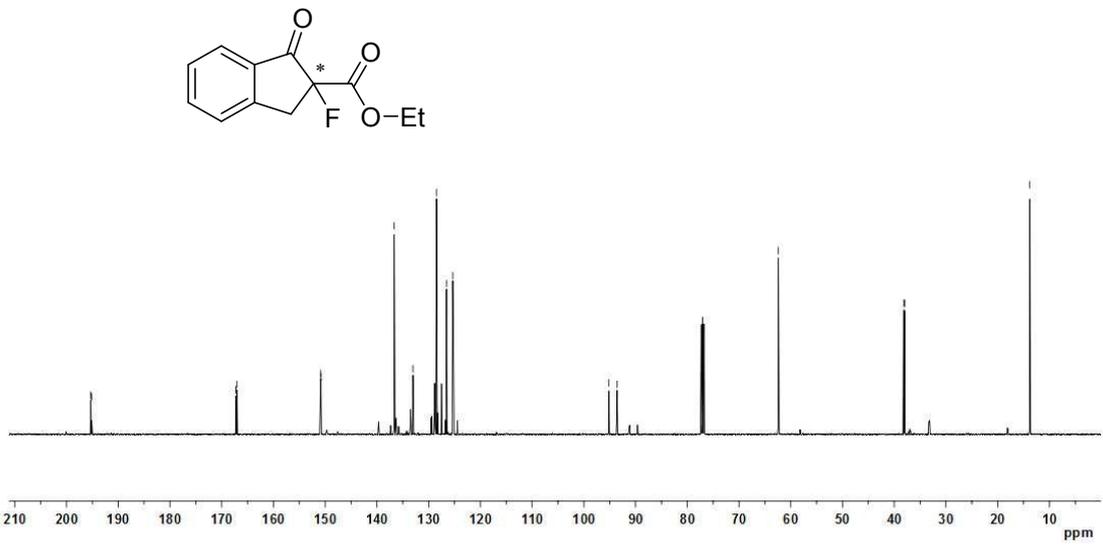
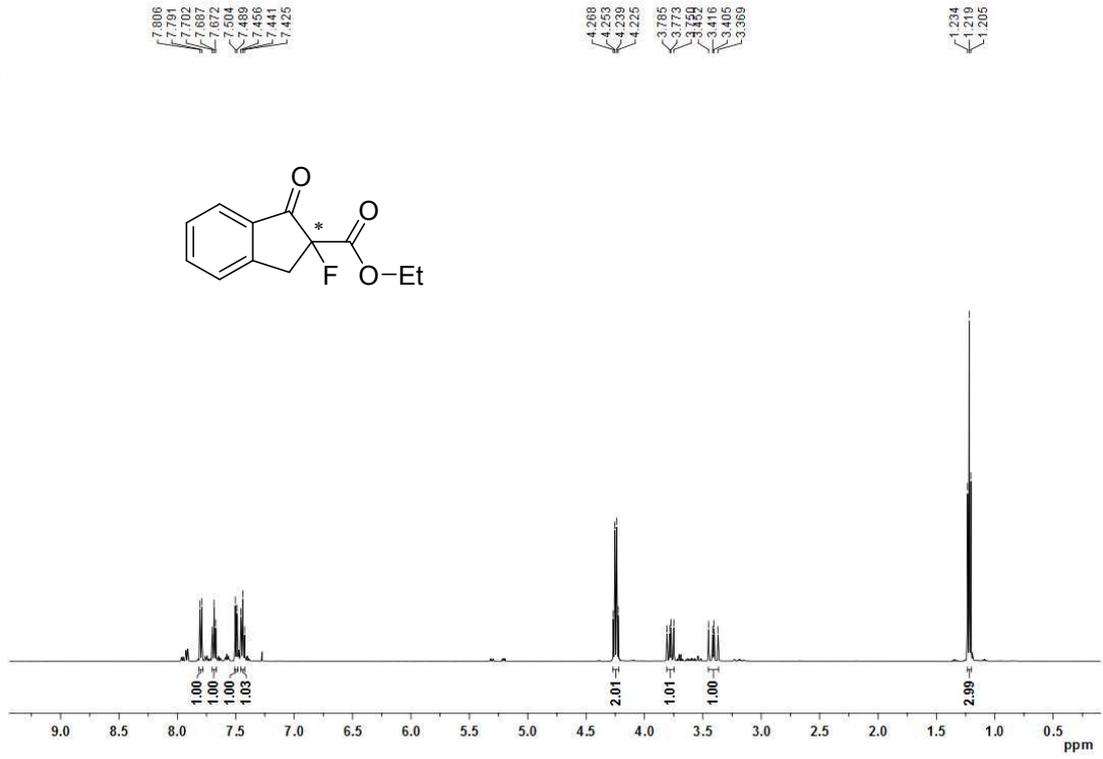
The substrates **2a-o** and the corresponding products **3a-o** are known compounds that exhibited ¹H and ¹³C NMR data in agreement with those reported in the literature.¹⁻³. The absolute configuration of the fluorination products was assigned by the chiral HPLC chromatograms with those reported in the literature under the same conditions.⁴

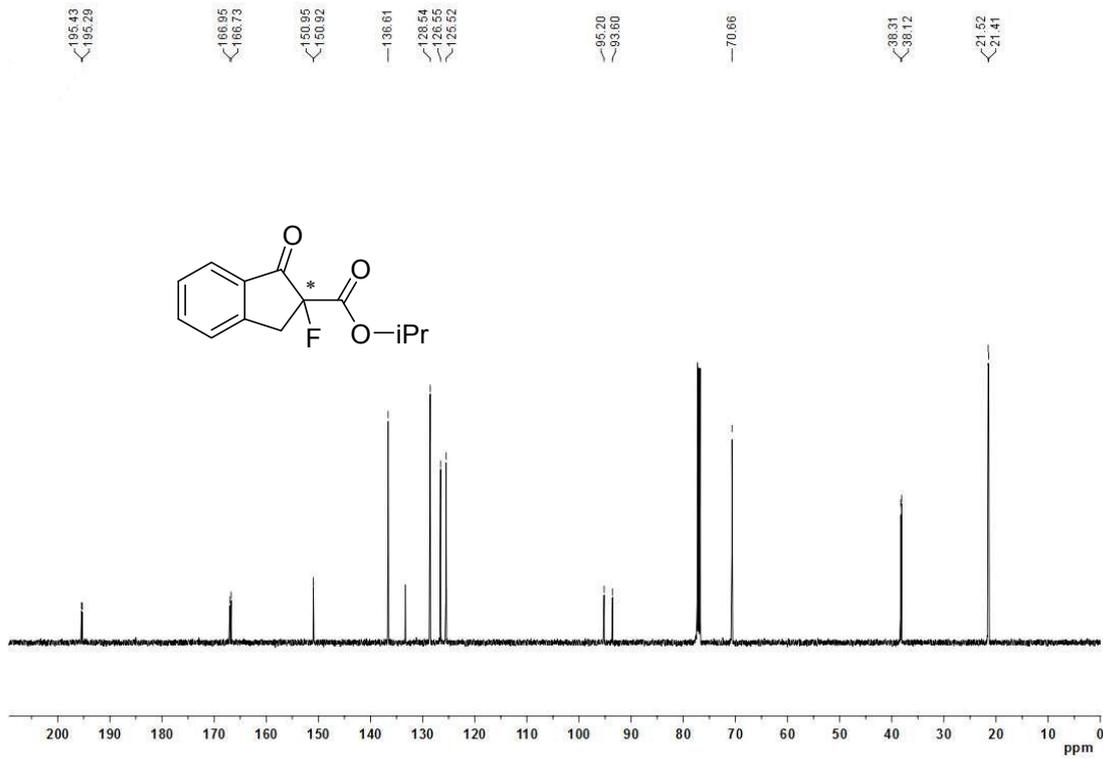
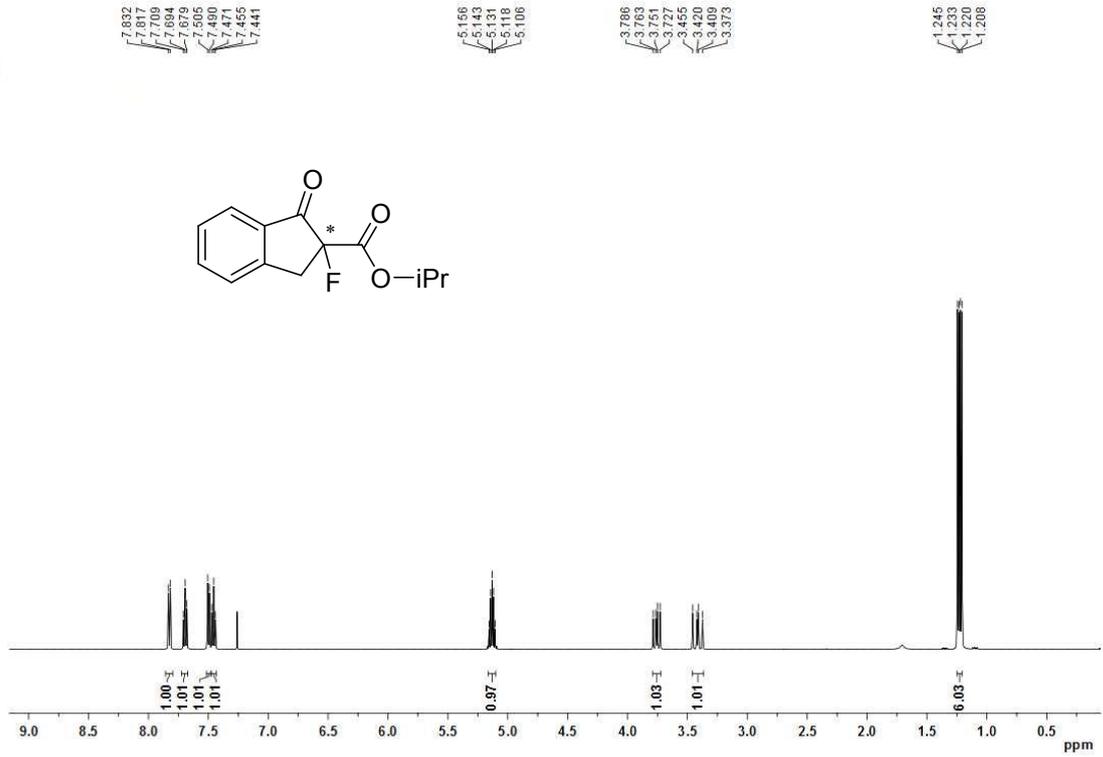
6. NMR spectra

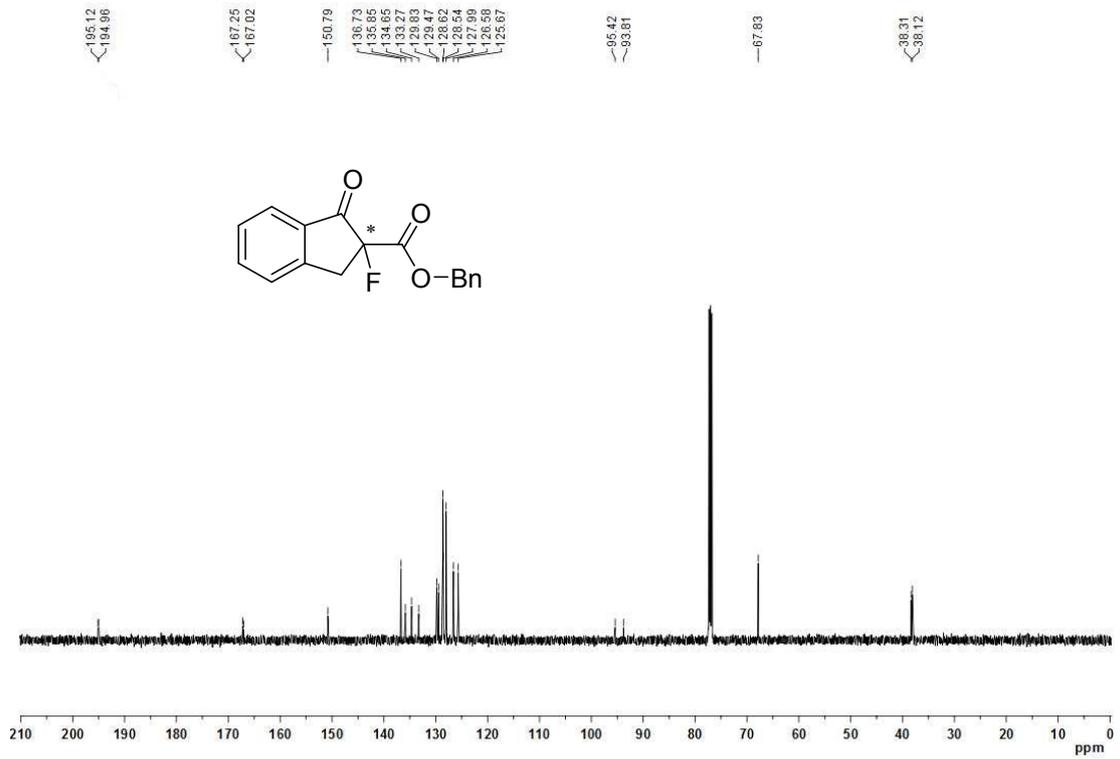
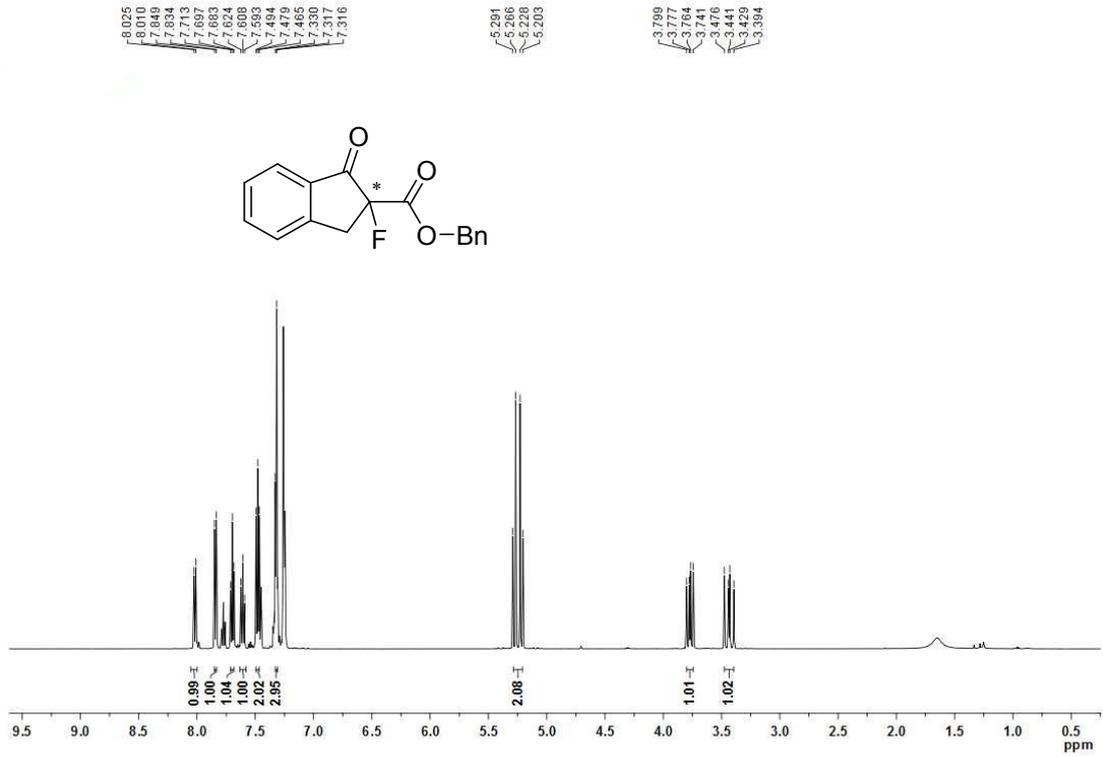


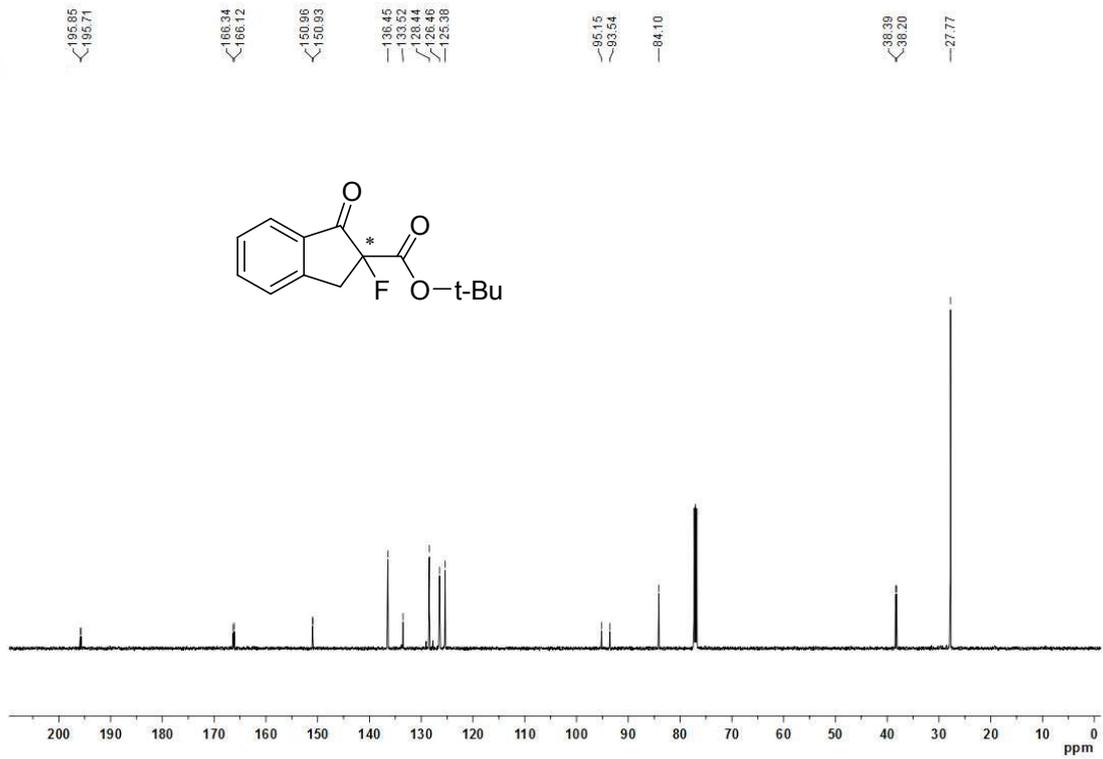
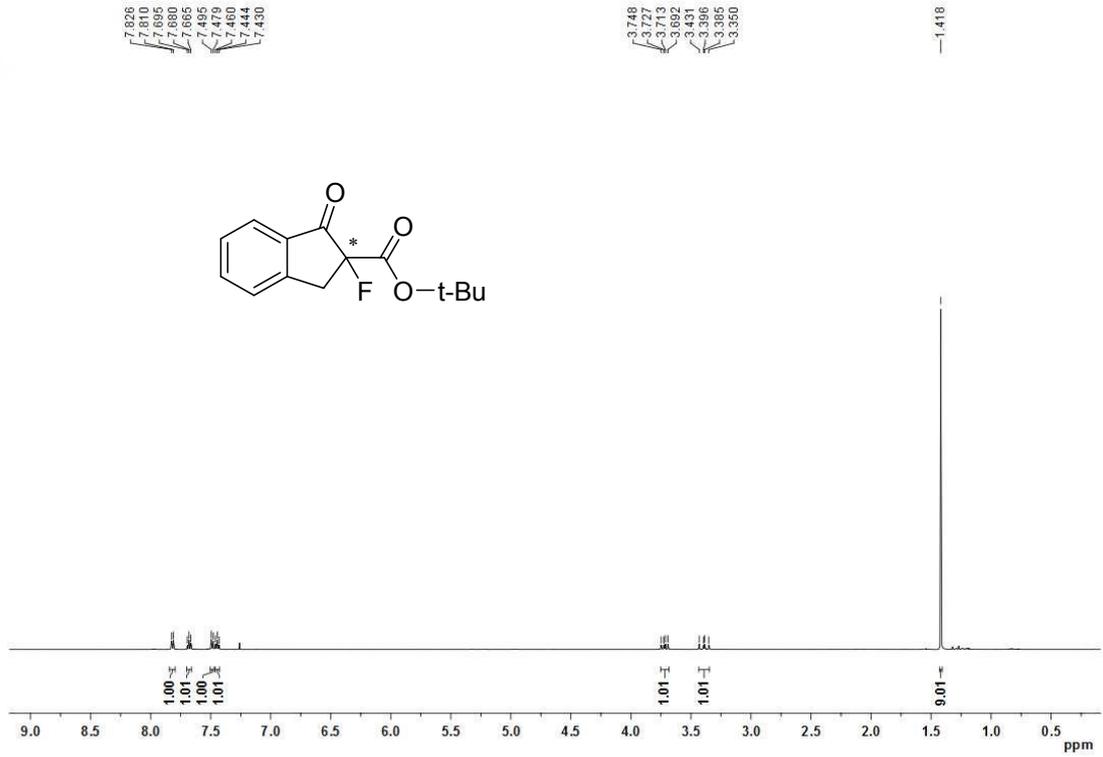








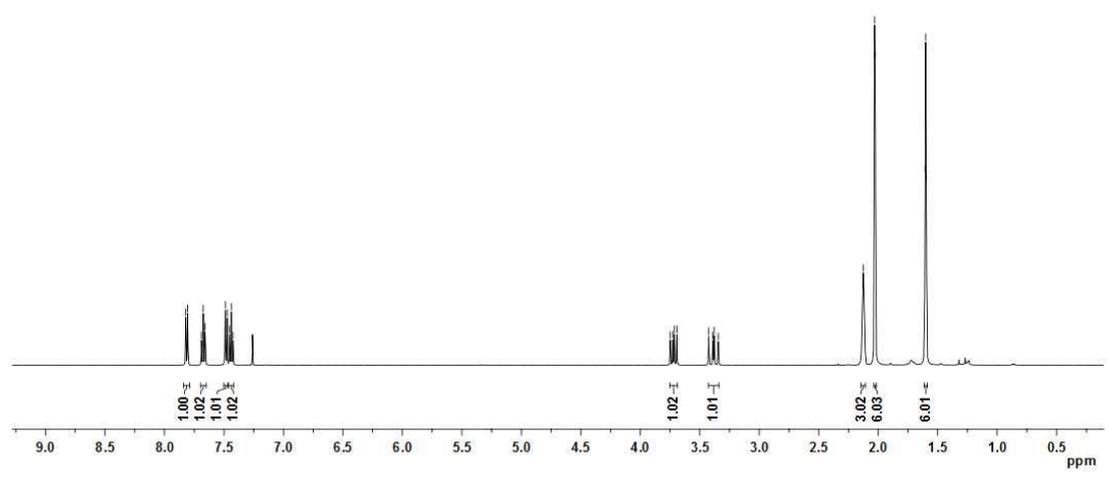
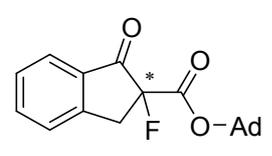




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7.81
7.69
7.67
7.66
7.49
7.47
7.45
7.44
7.42

3.75
3.71
3.69
3.42
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3.38
3.34

2.13
2.03
2.02
1.61
1.60
1.59



195.88
165.74

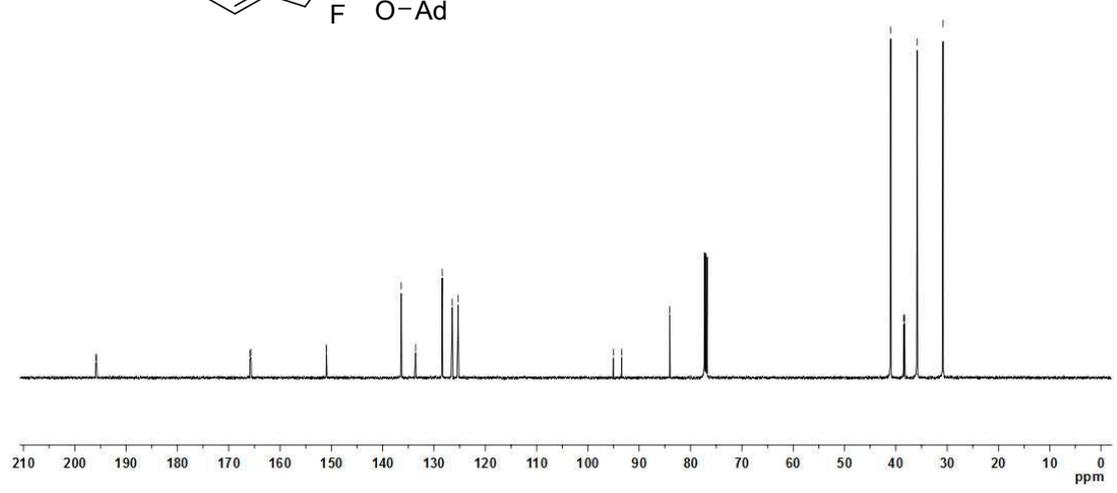
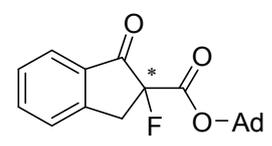
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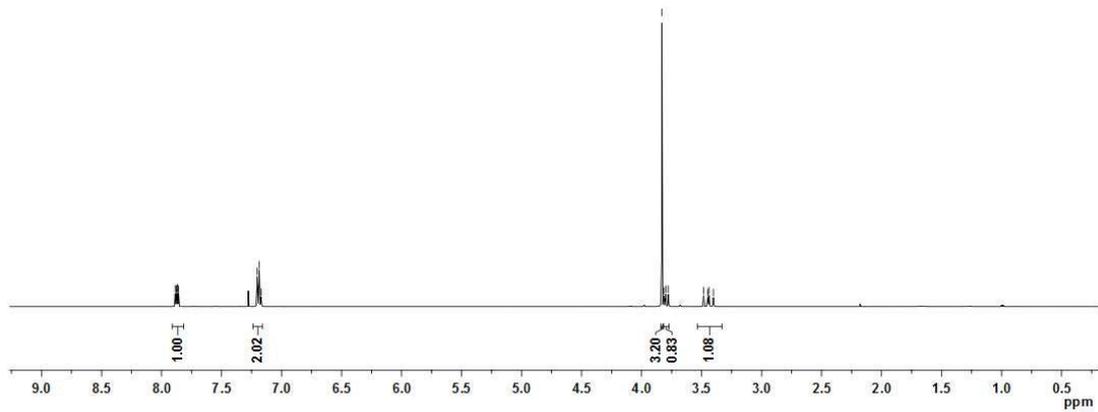
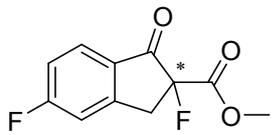
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30.81



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7.171

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3.463
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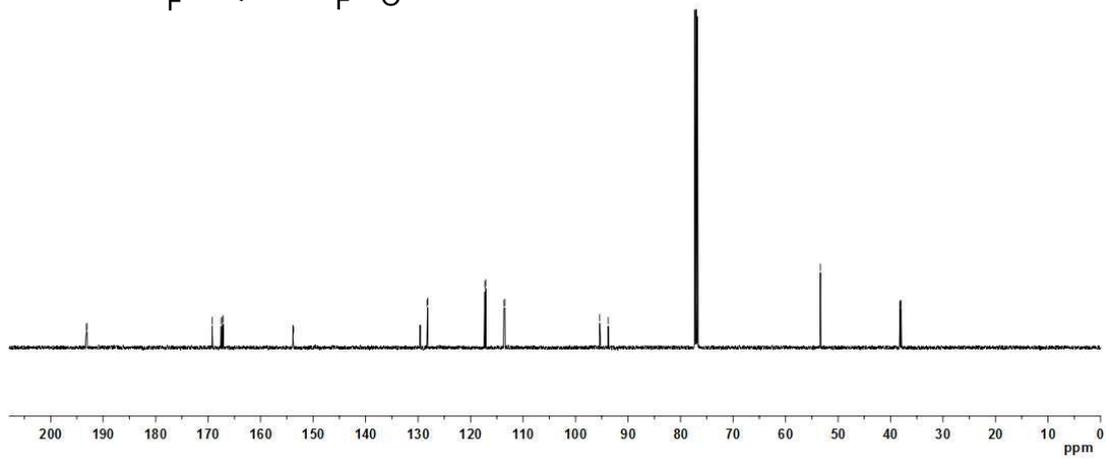
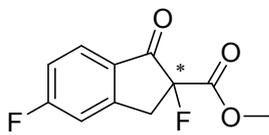
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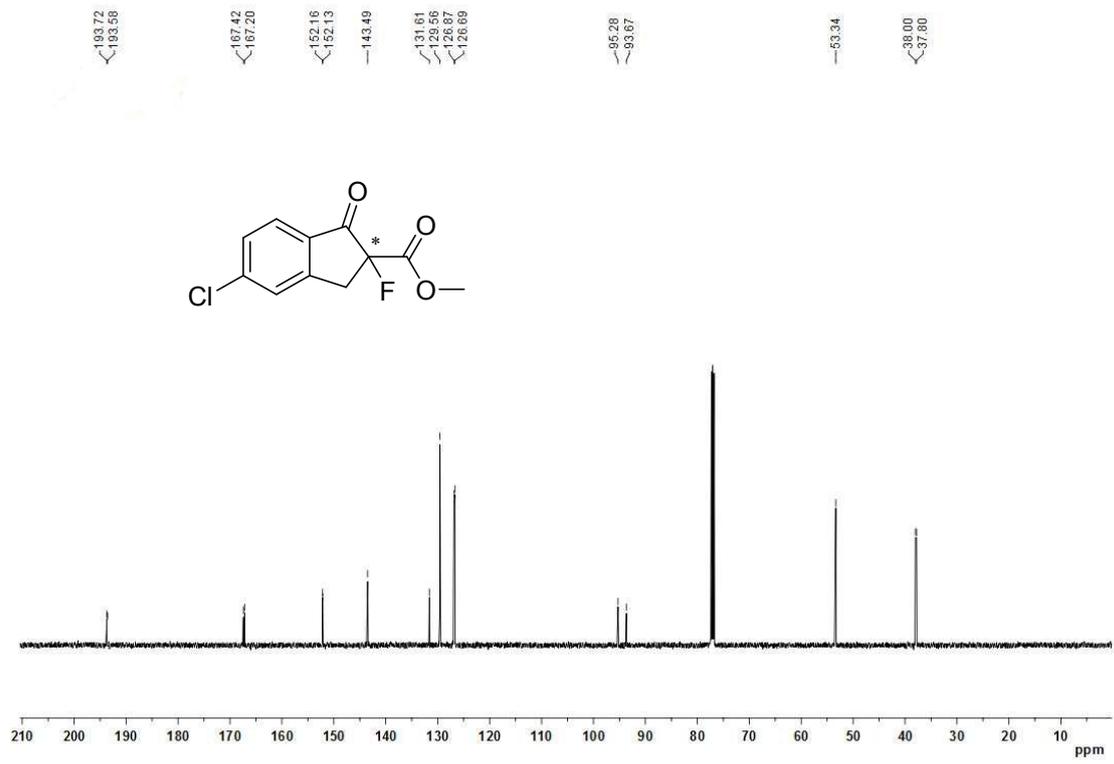
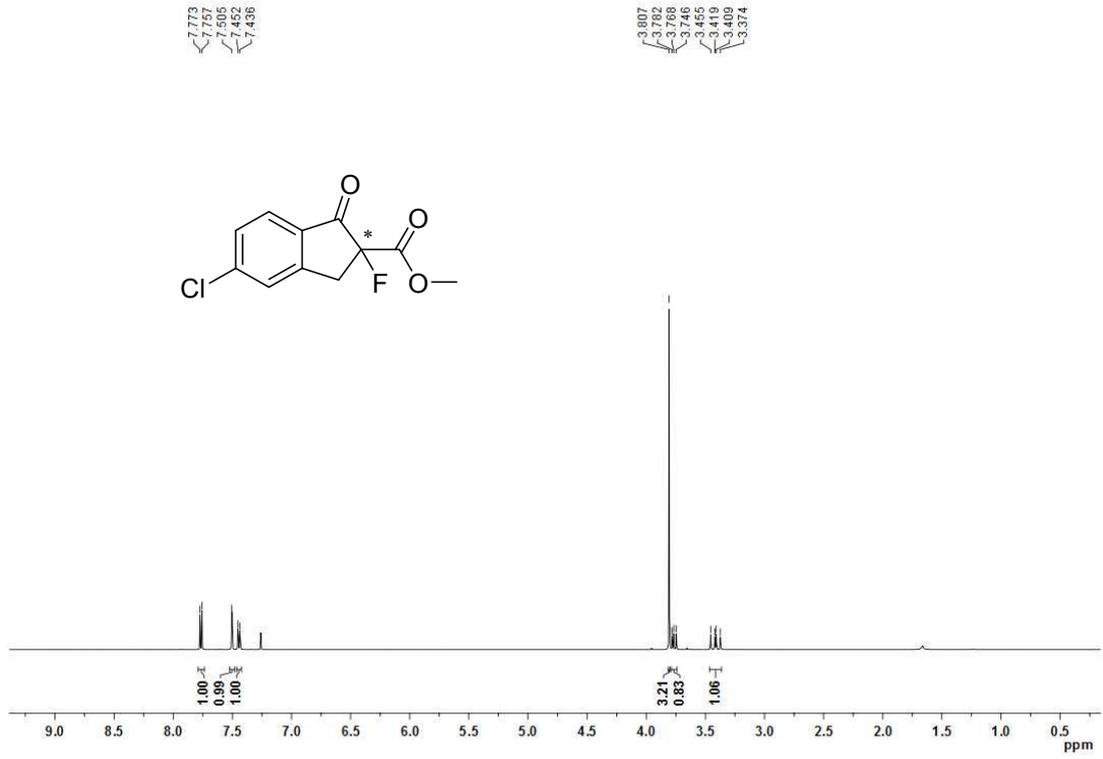
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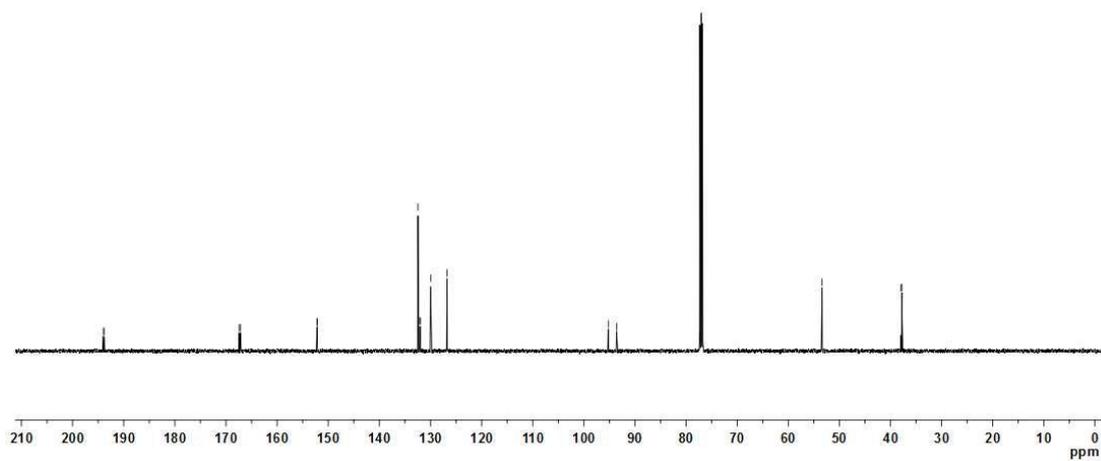
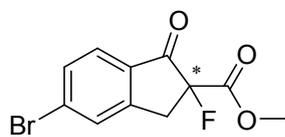
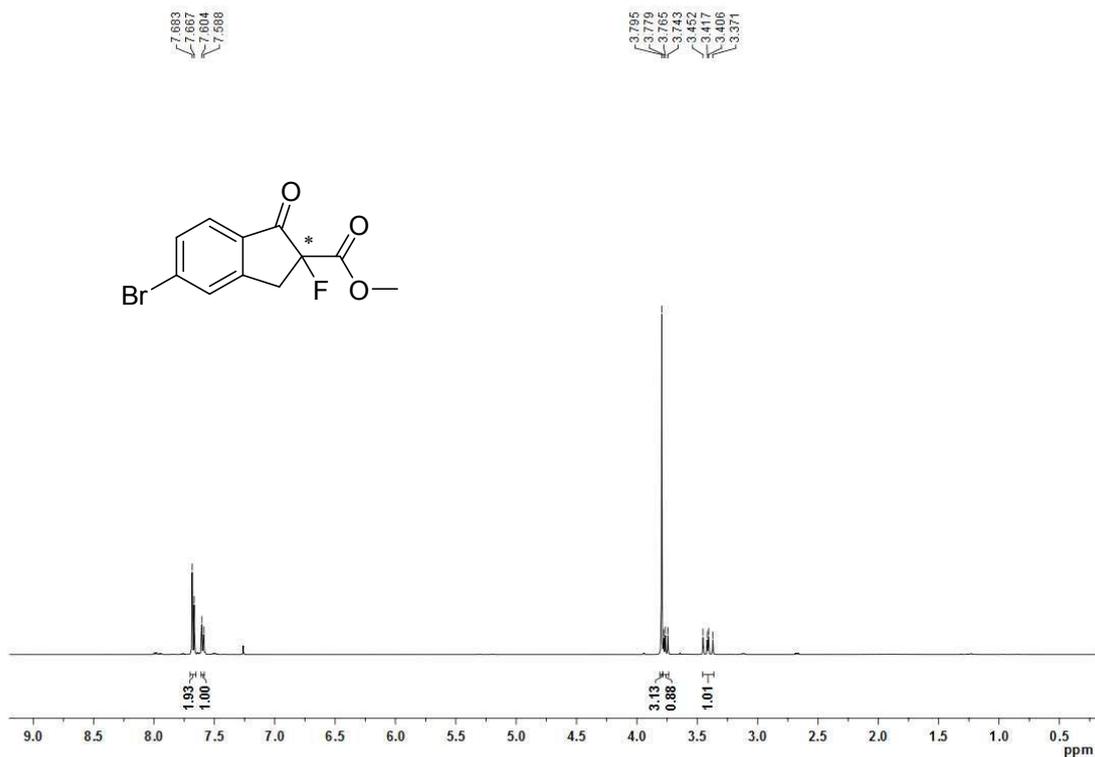
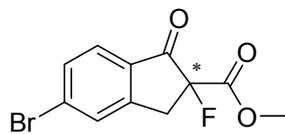
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37.80

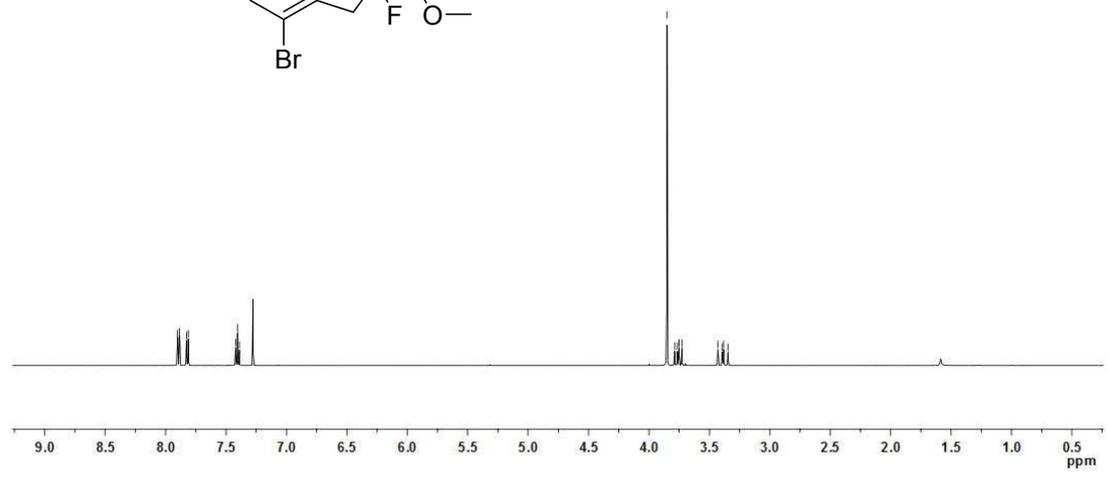
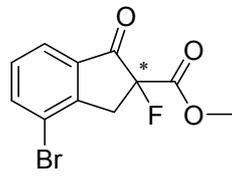






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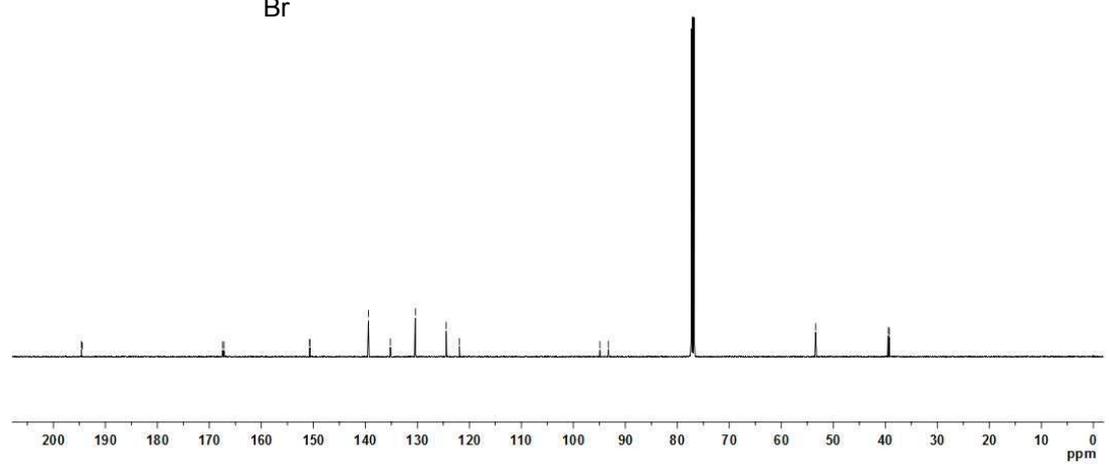
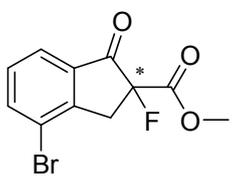
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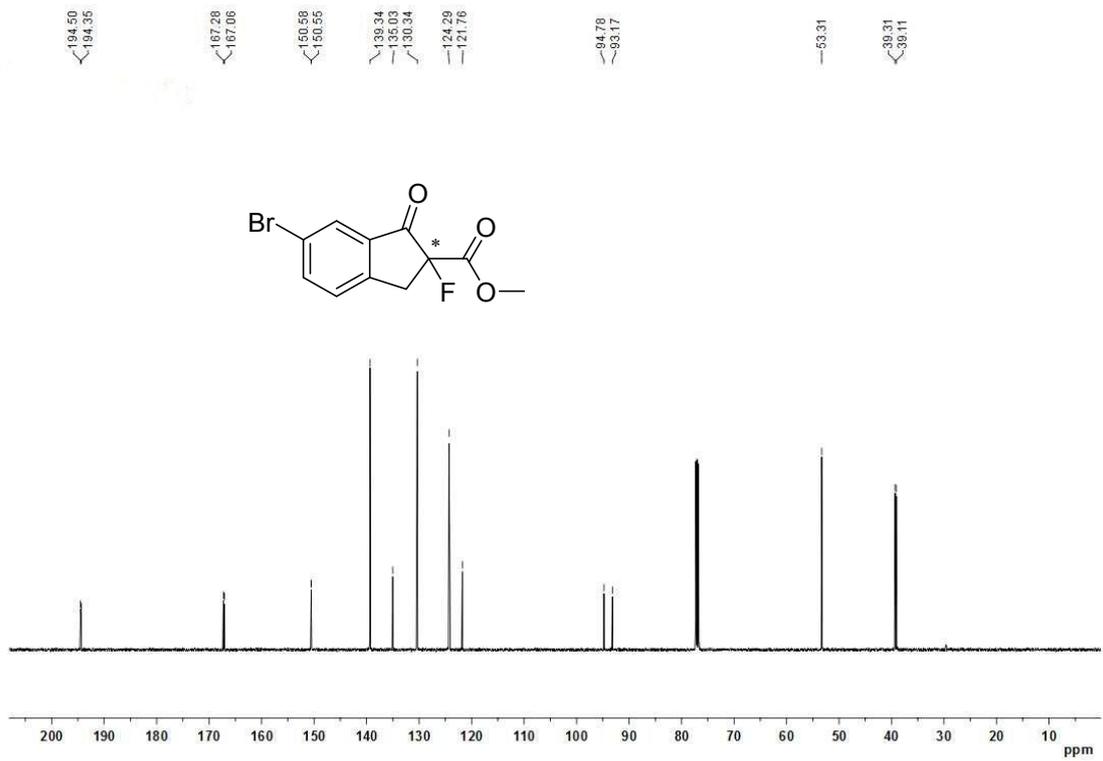
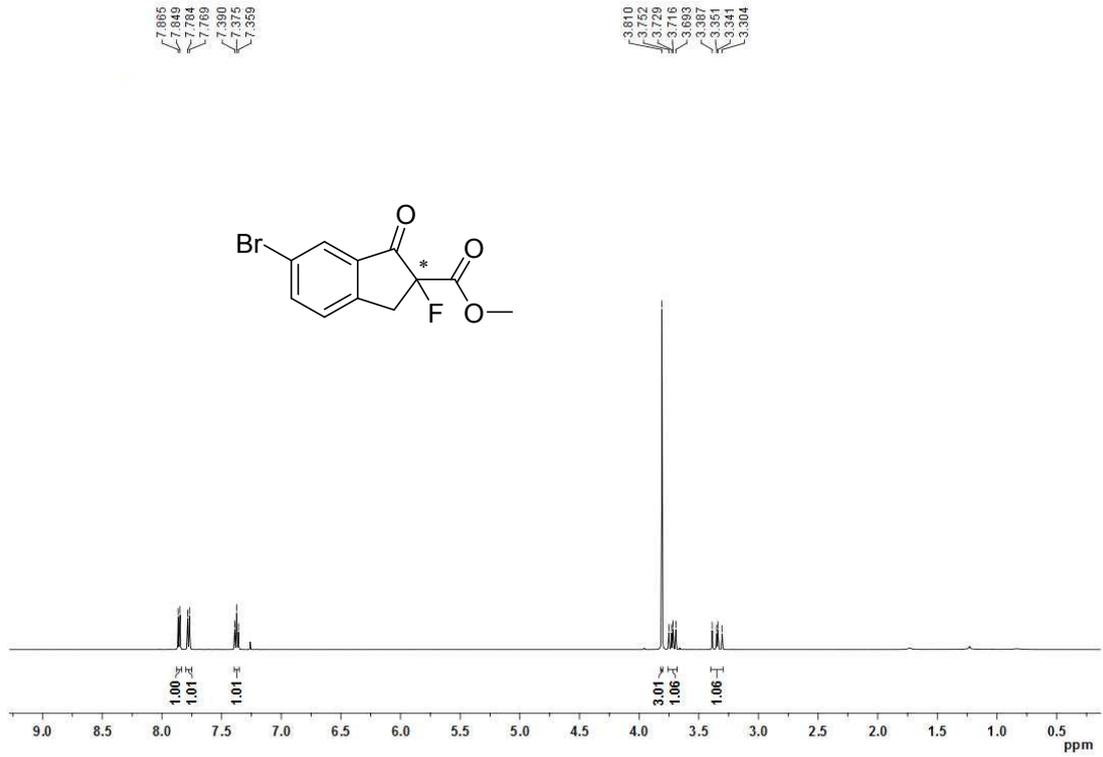
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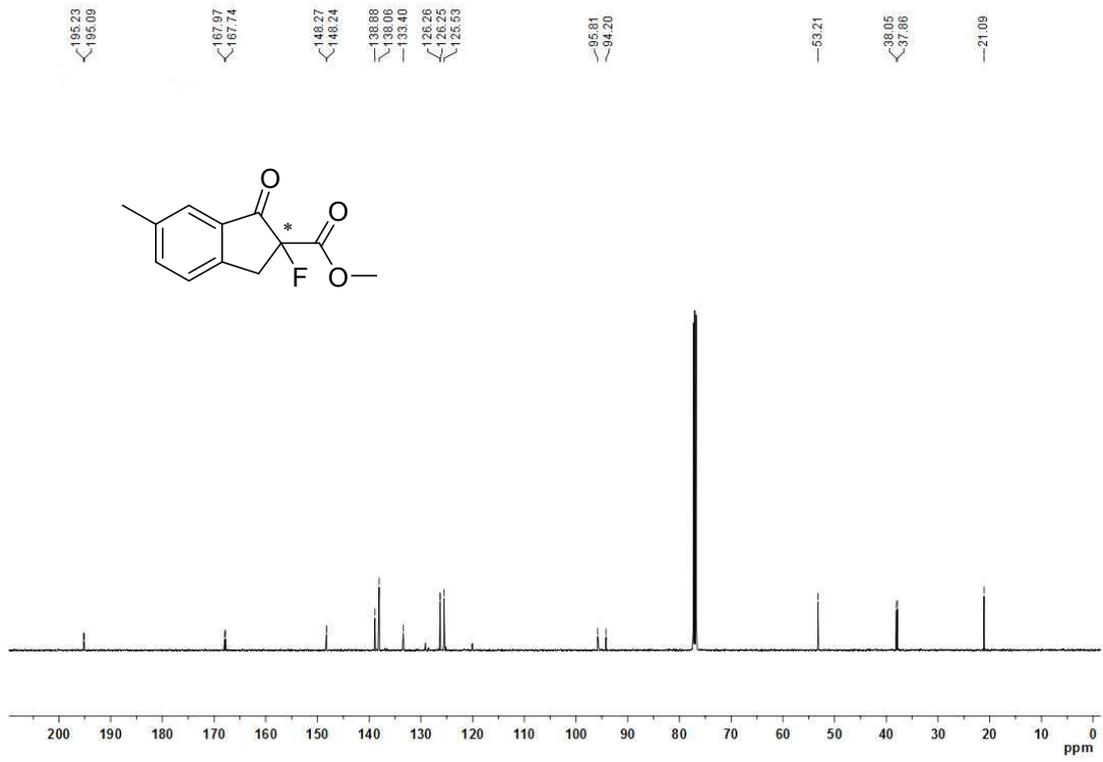
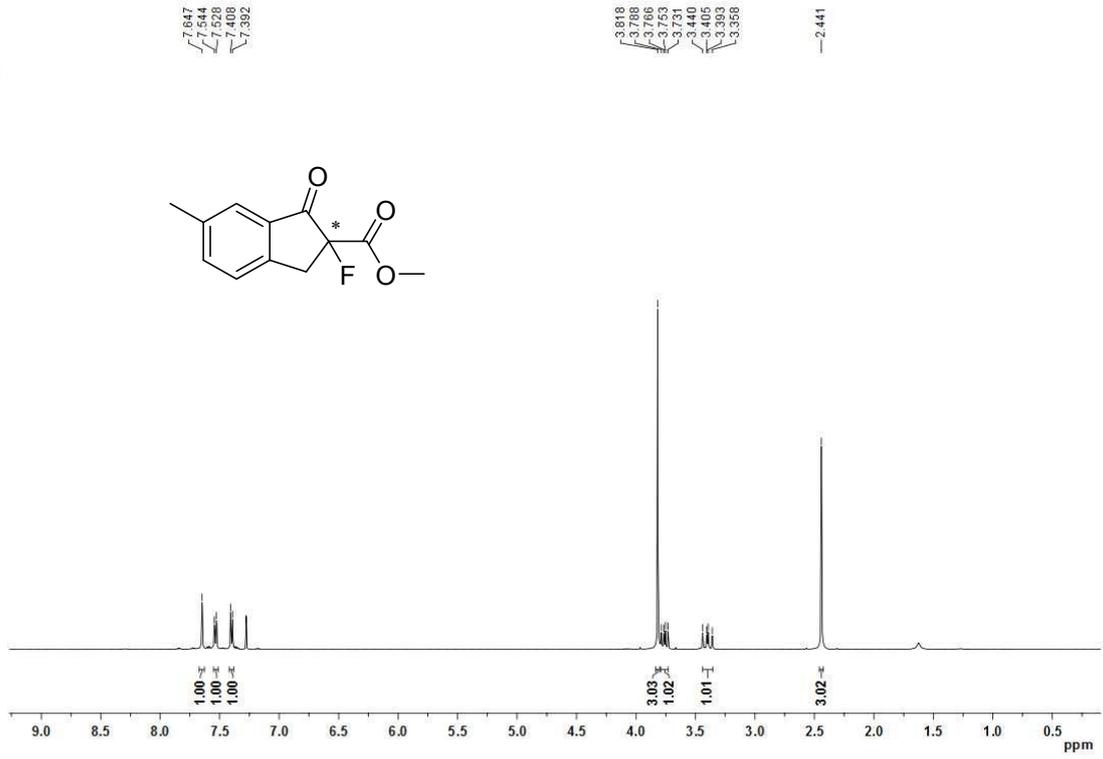
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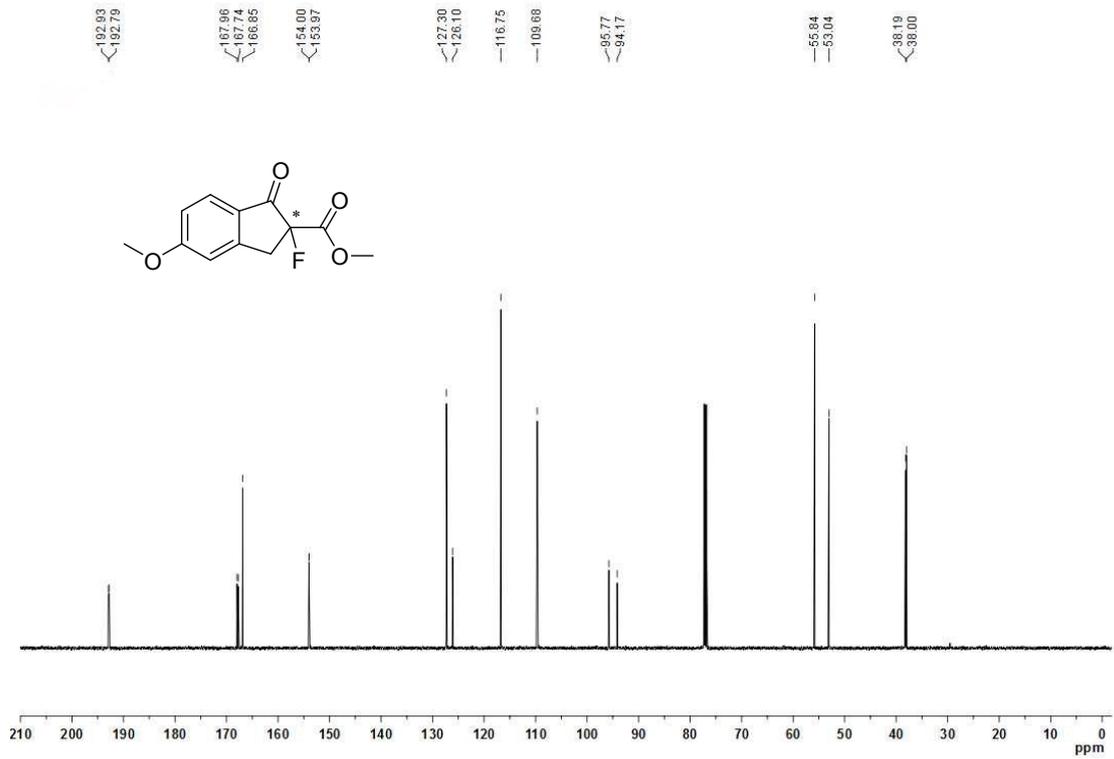
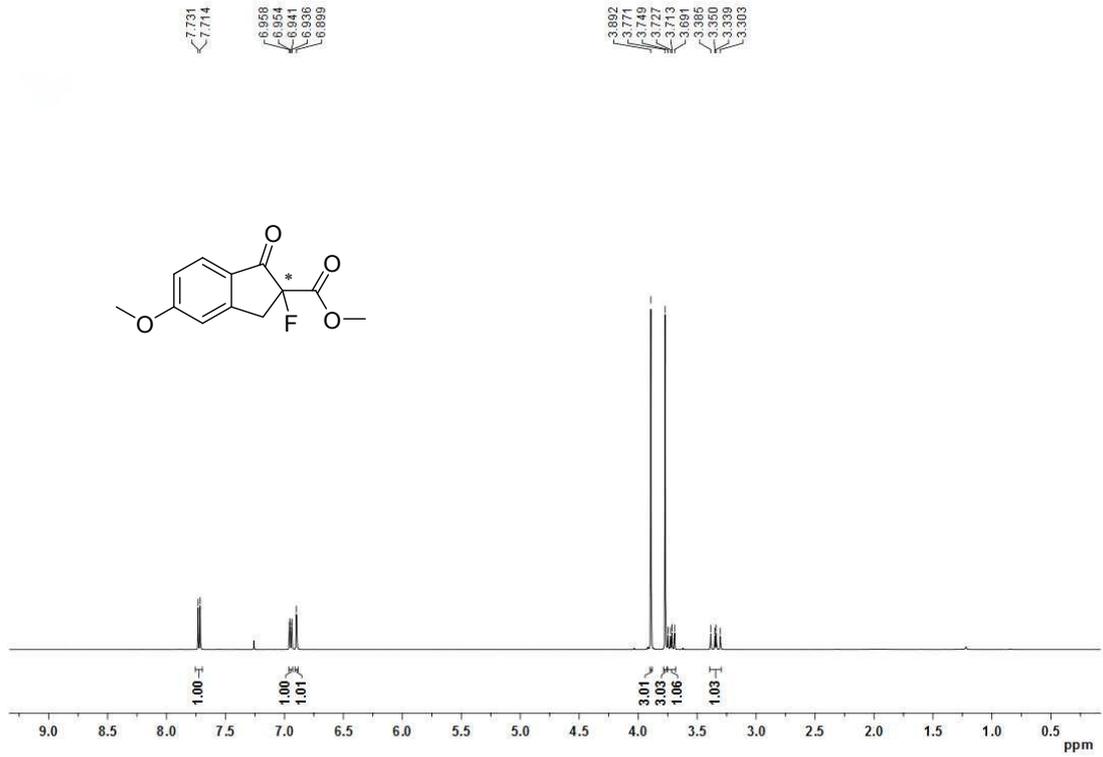
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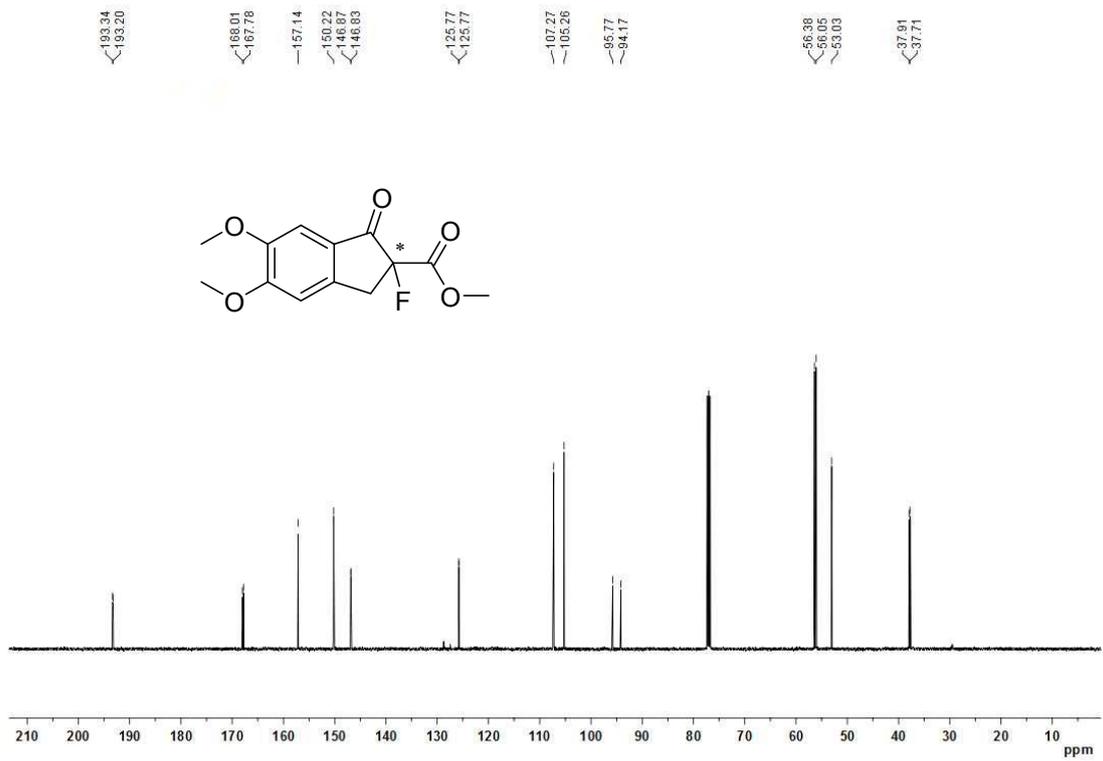
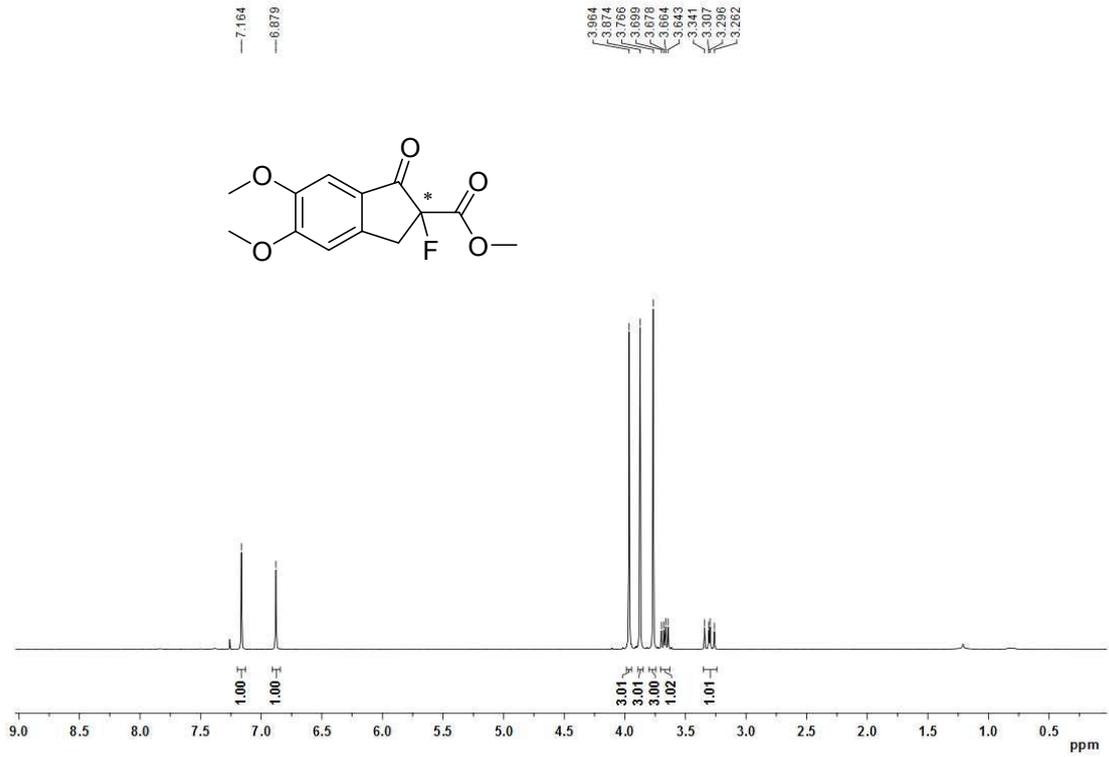
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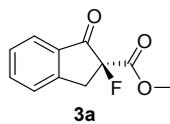




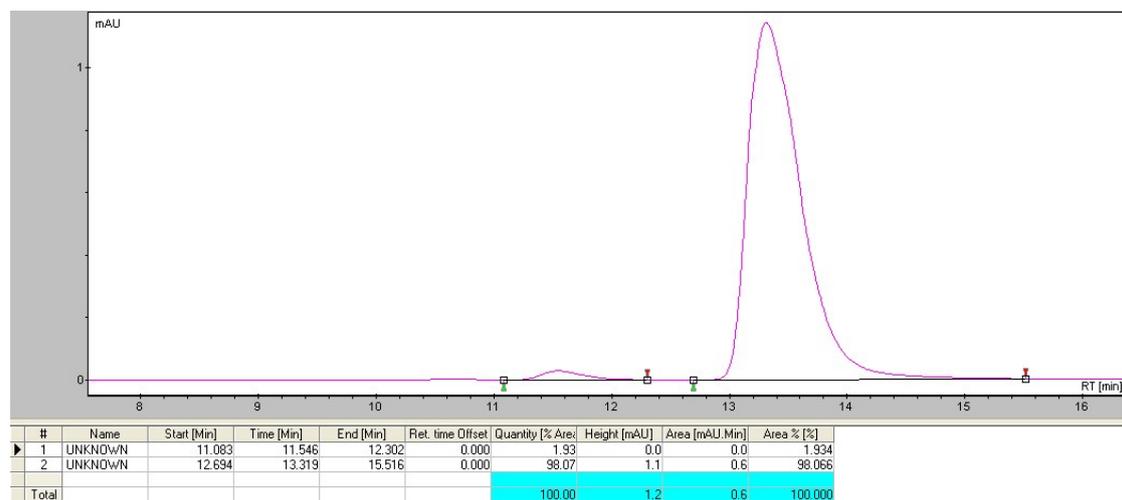




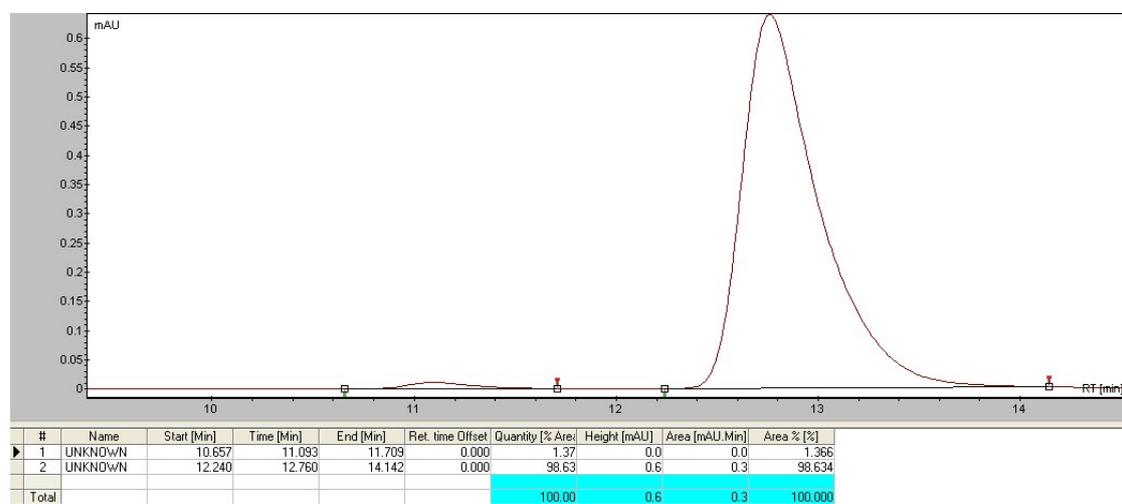
7. HPLC traces of the fluorinated product and racemic standards



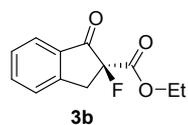
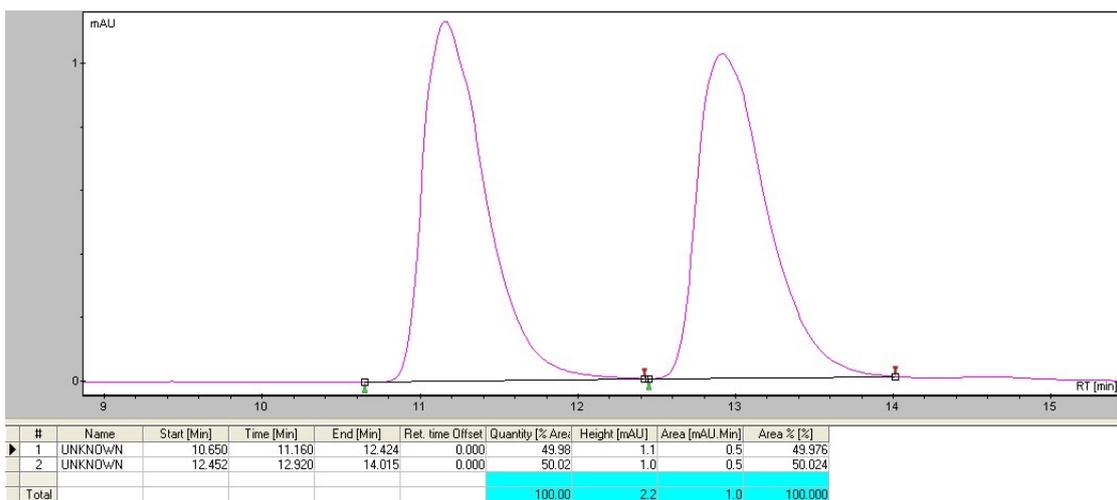
In batch condition



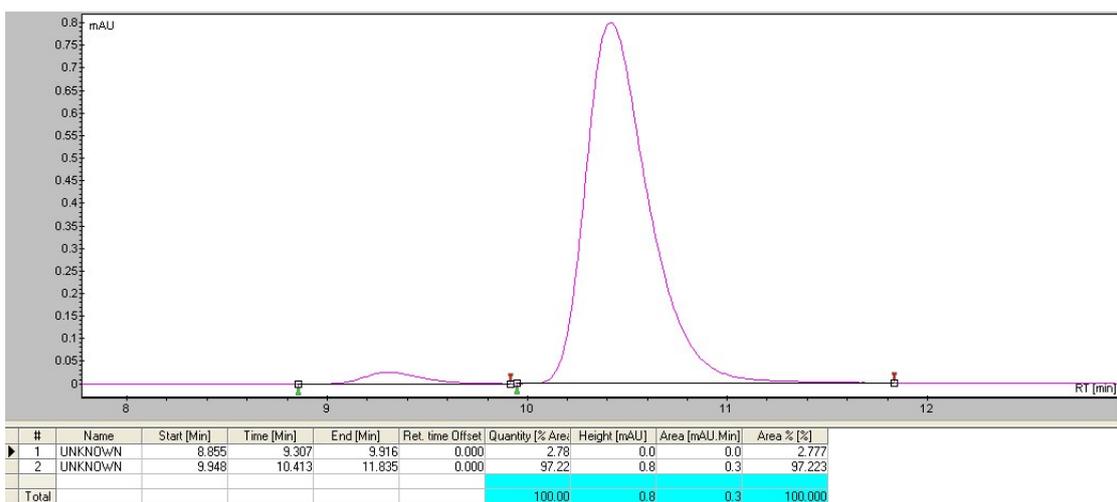
In continuous flow condition



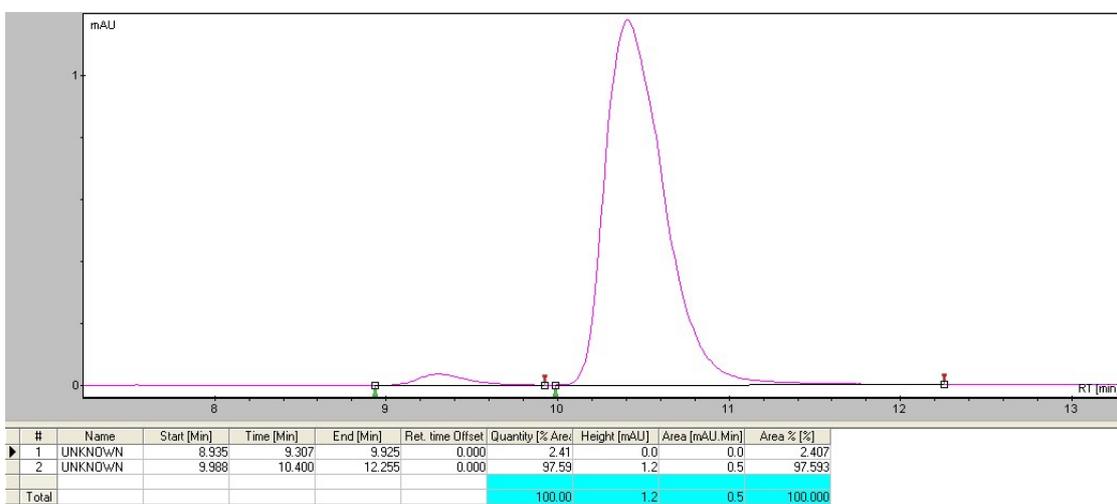
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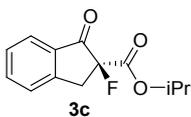
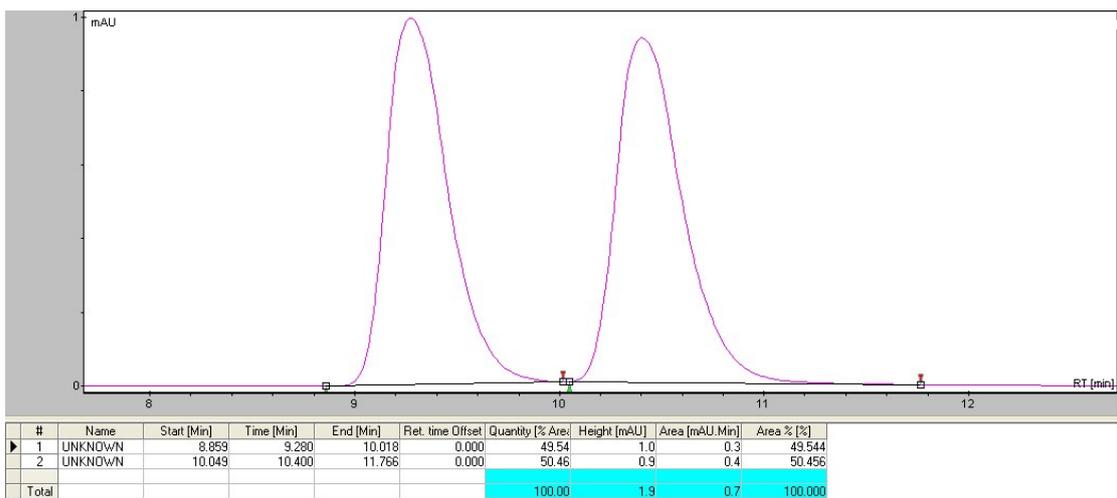
In batch condition



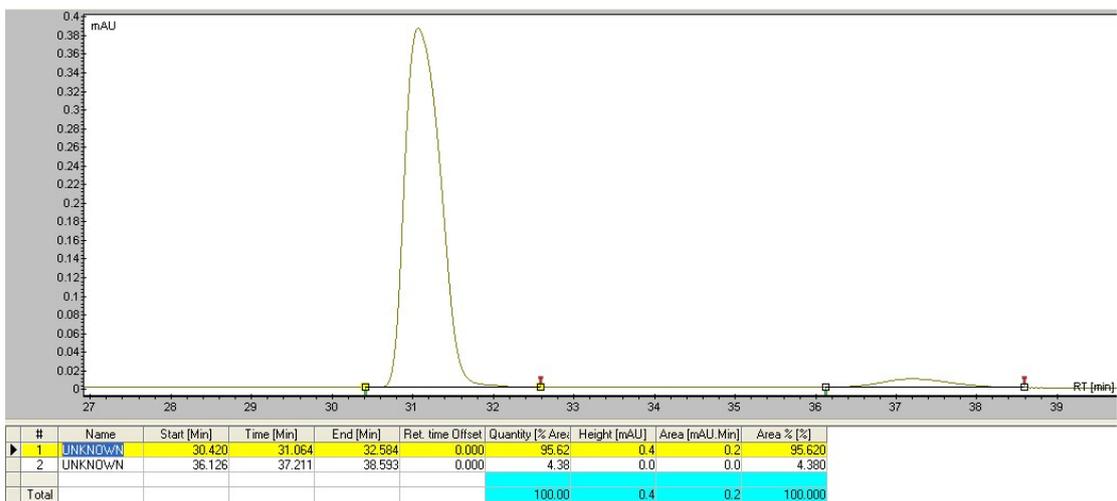
In continuous flow condition



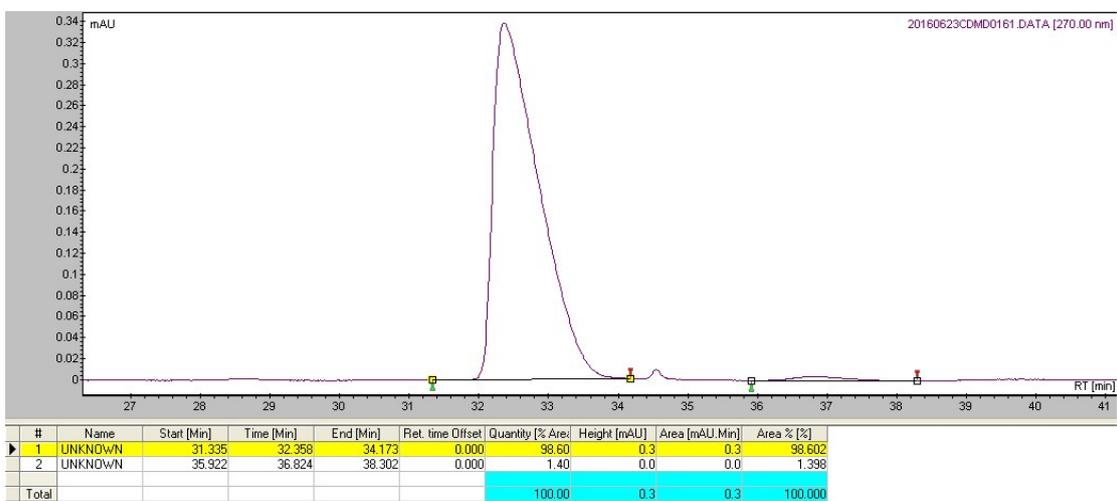
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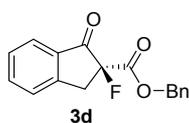
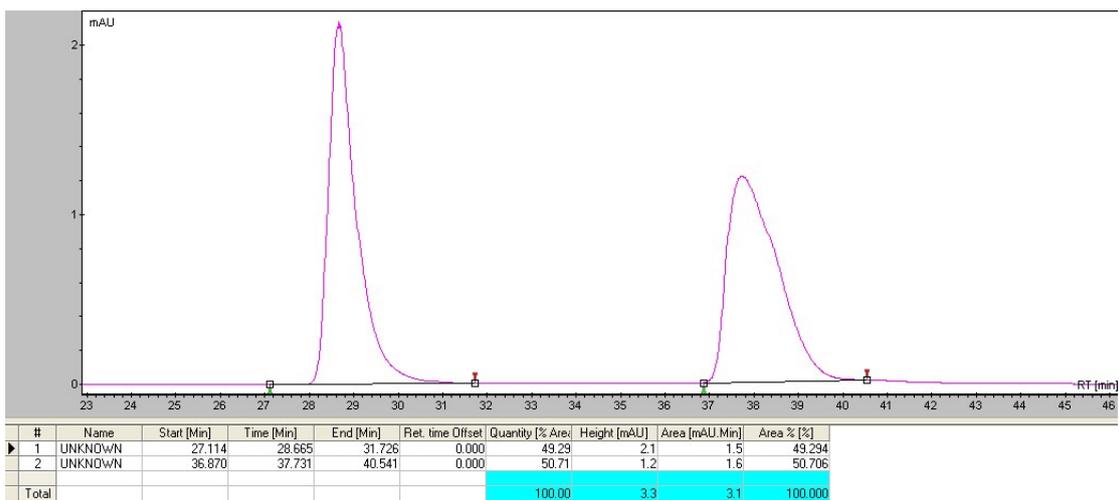
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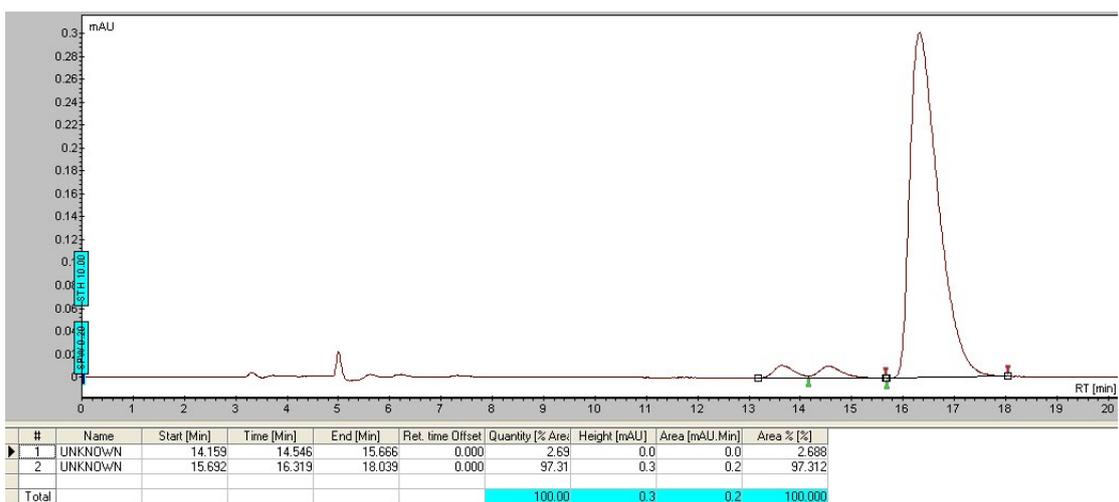
In continuous flow condition



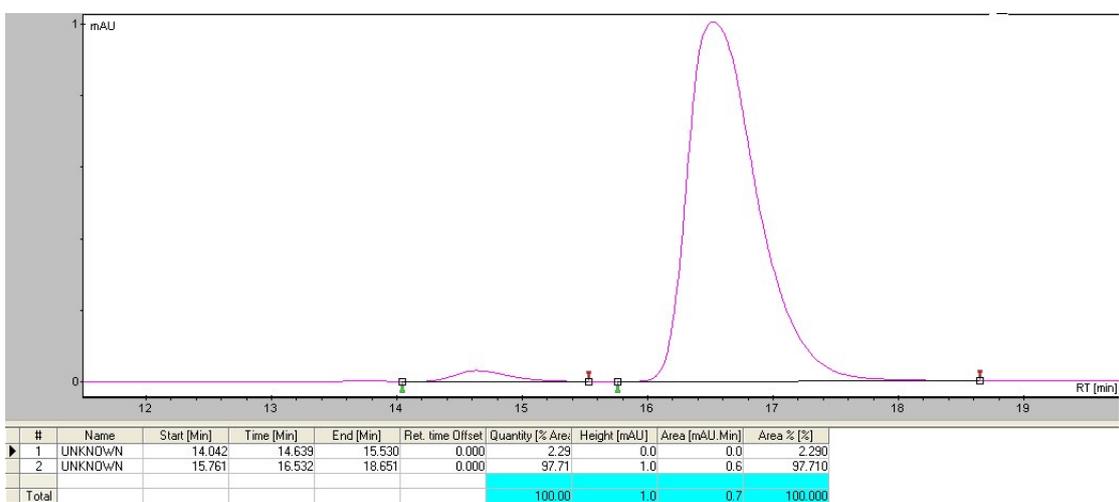
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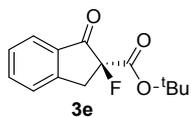
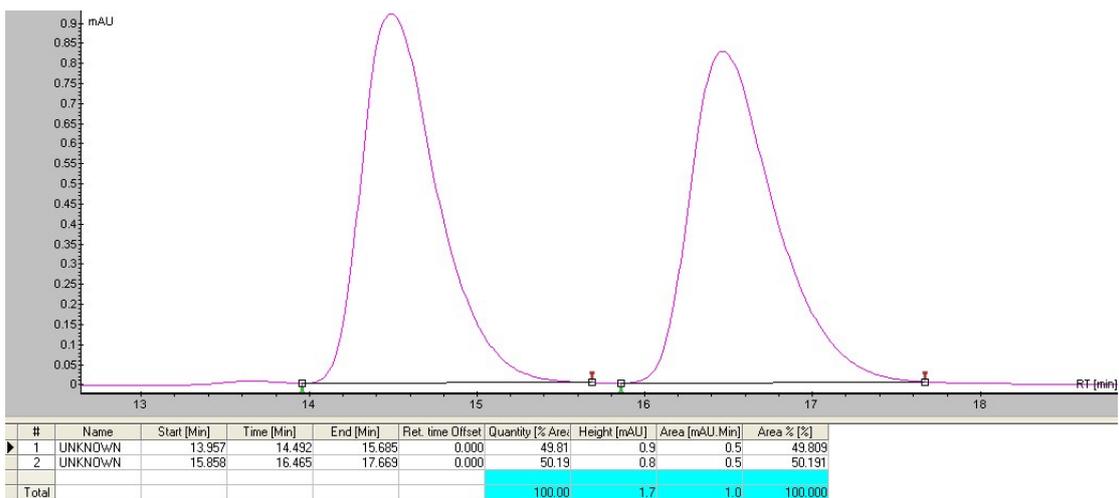
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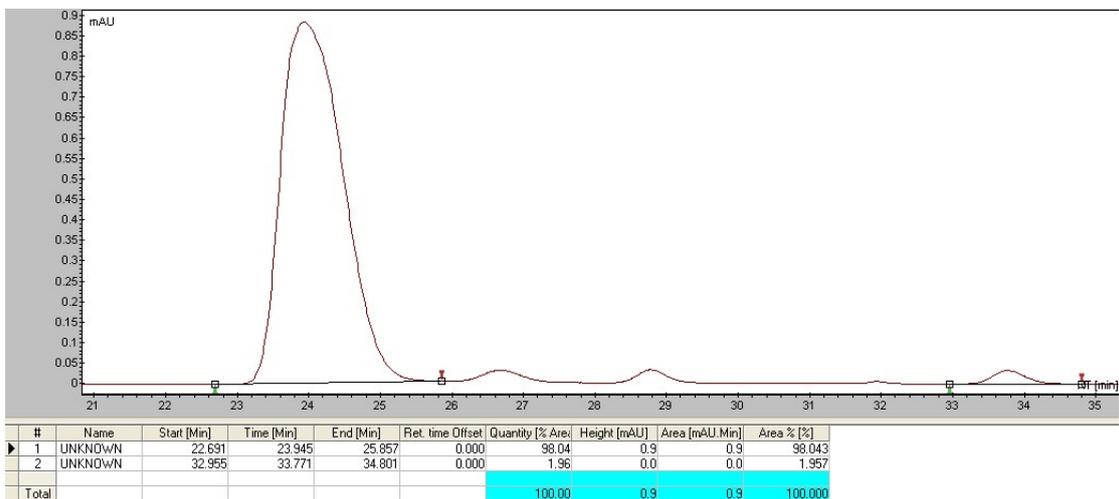
In continuous flow condition



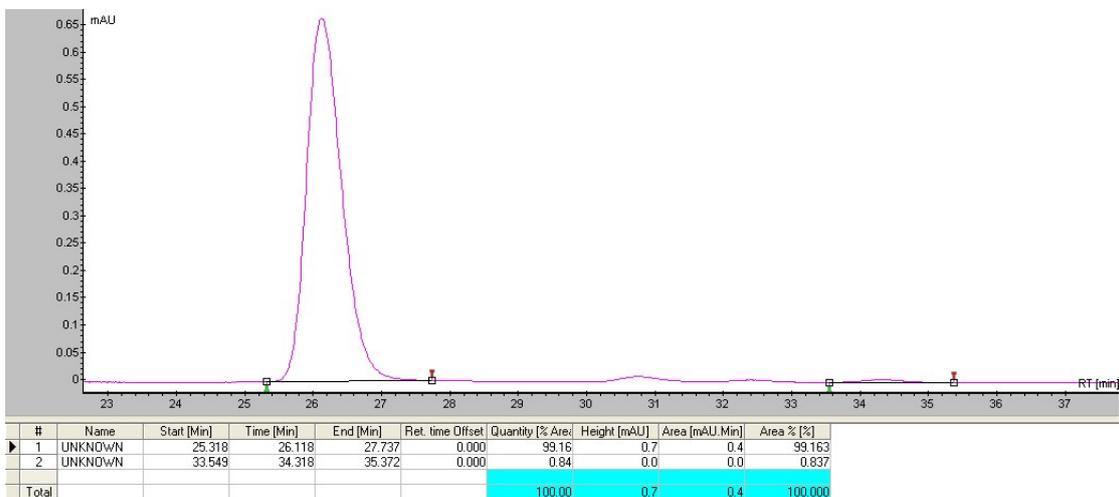
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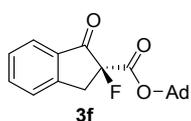
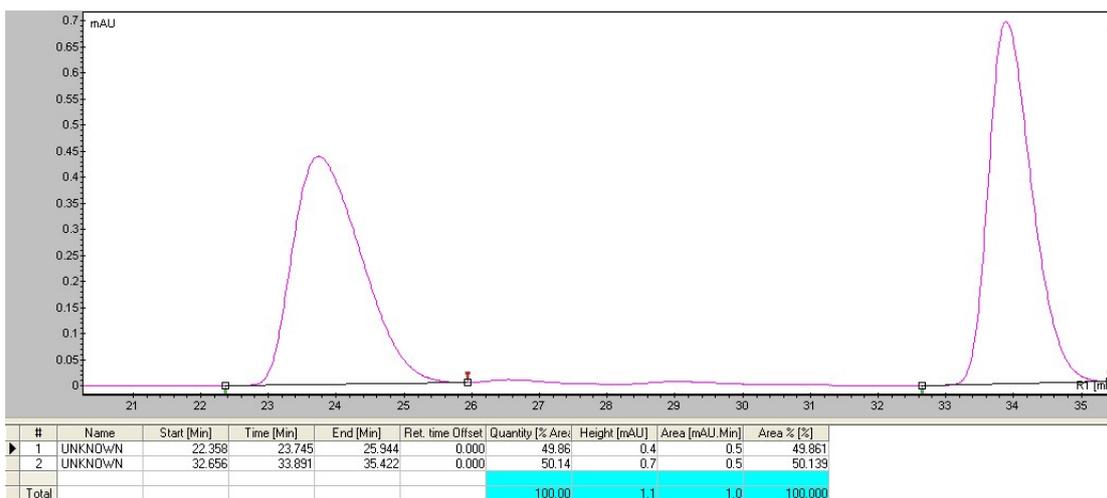
In batch condition



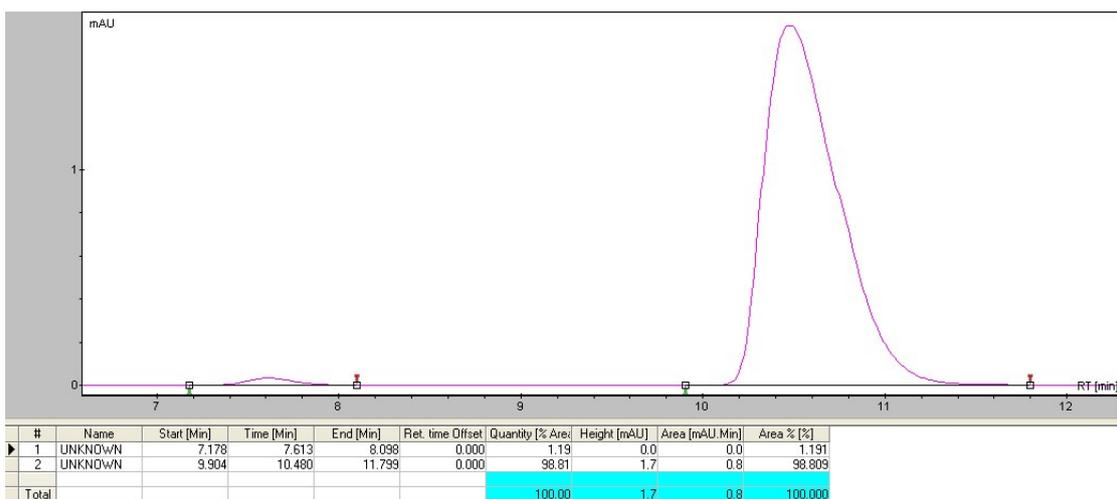
In continuous flow condition



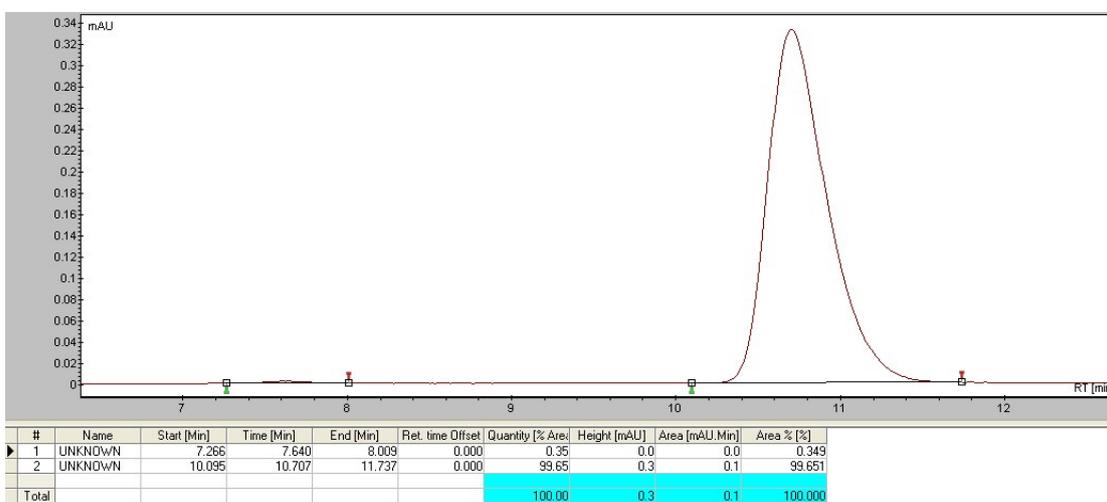
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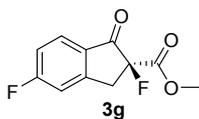
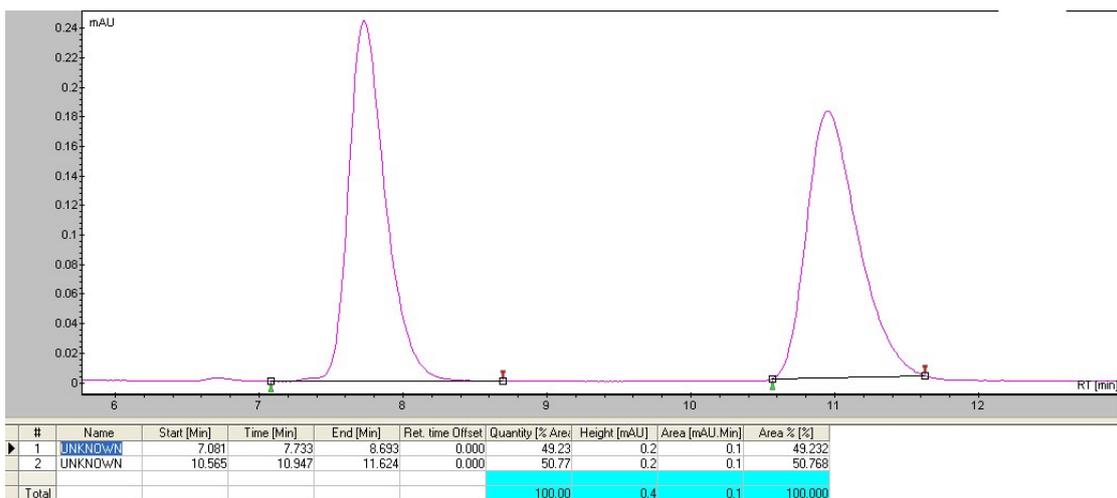
In batch condition



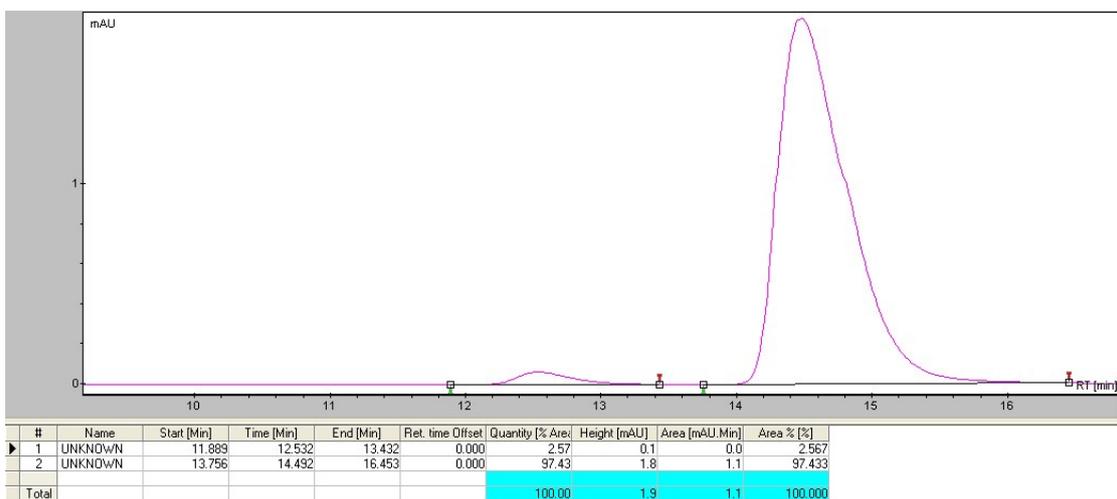
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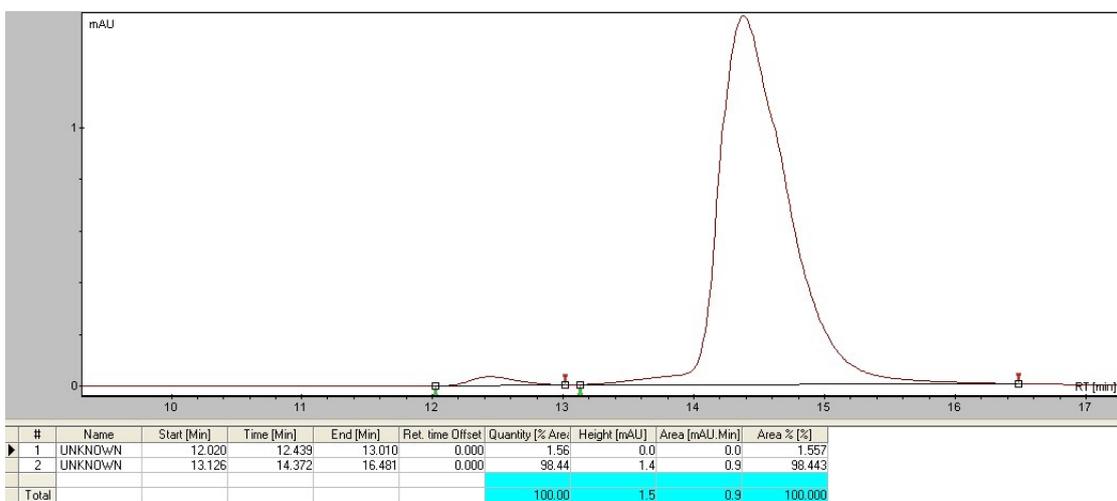
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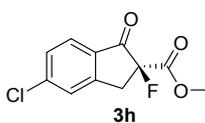
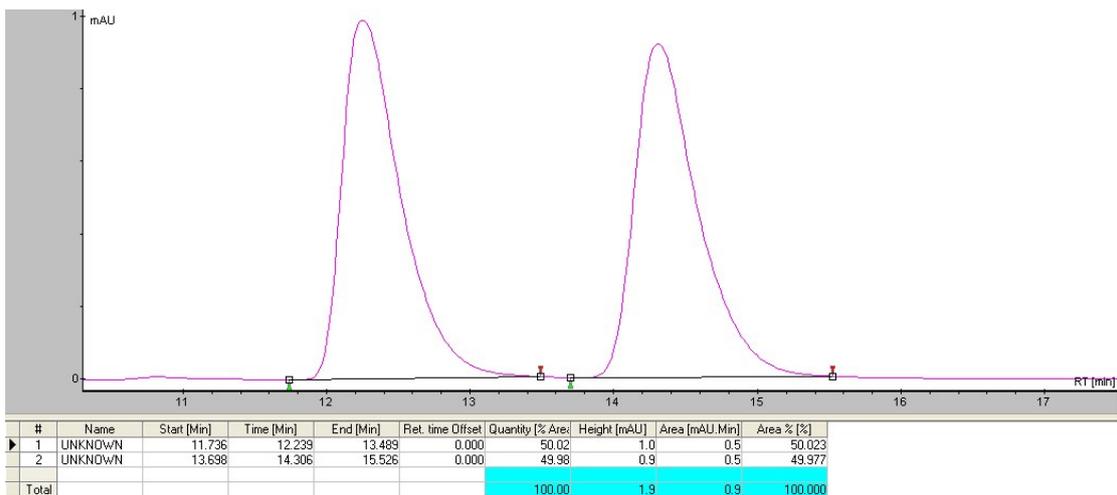
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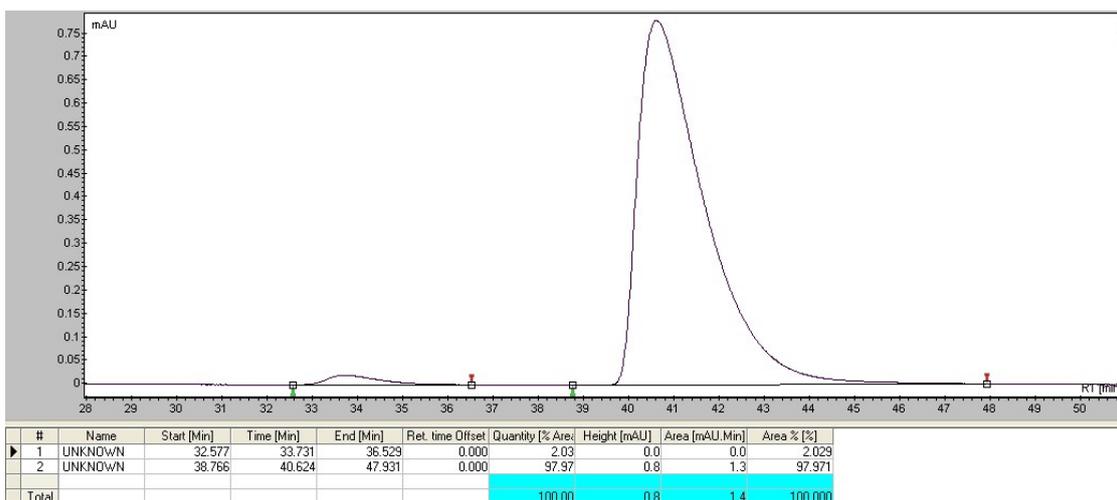
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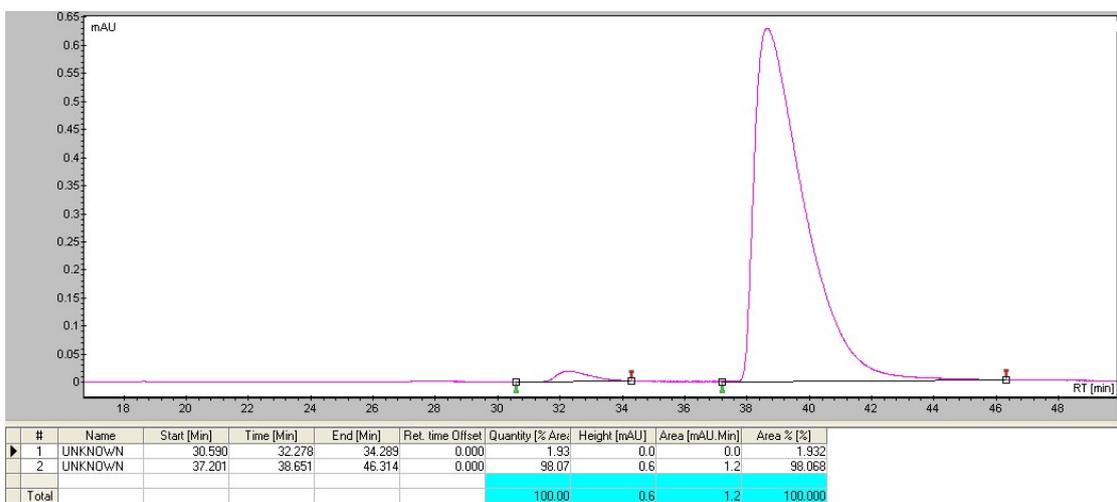
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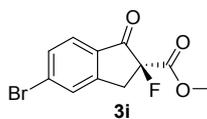
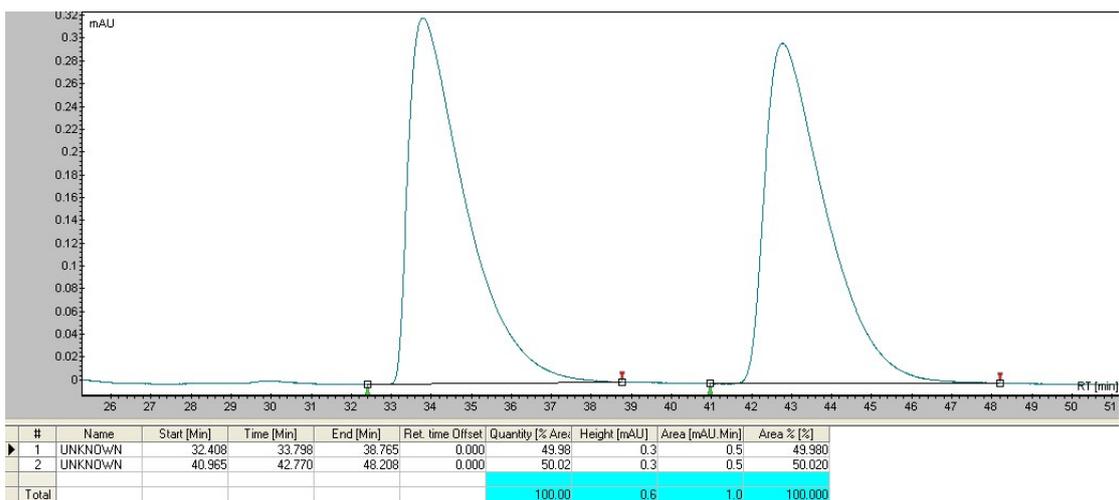
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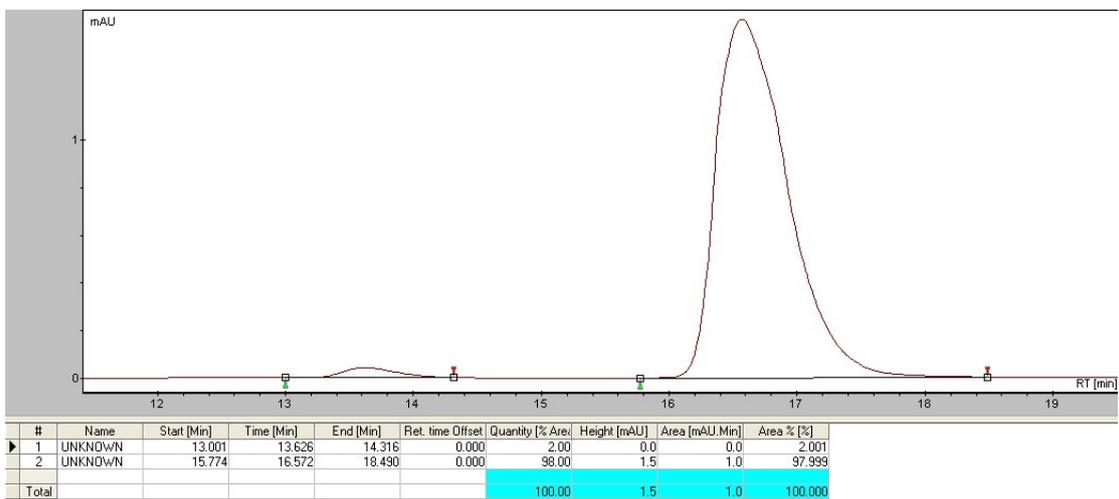
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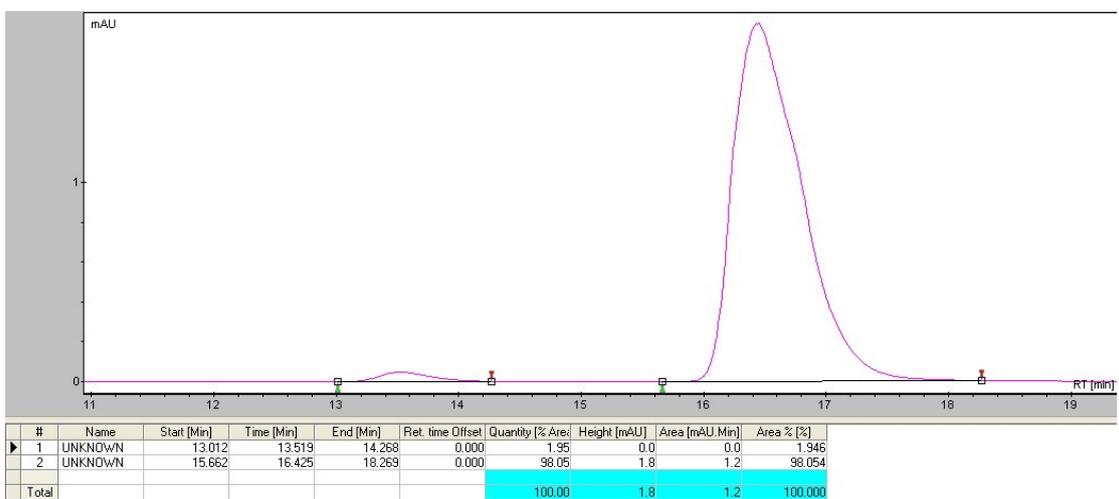
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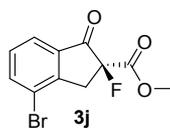
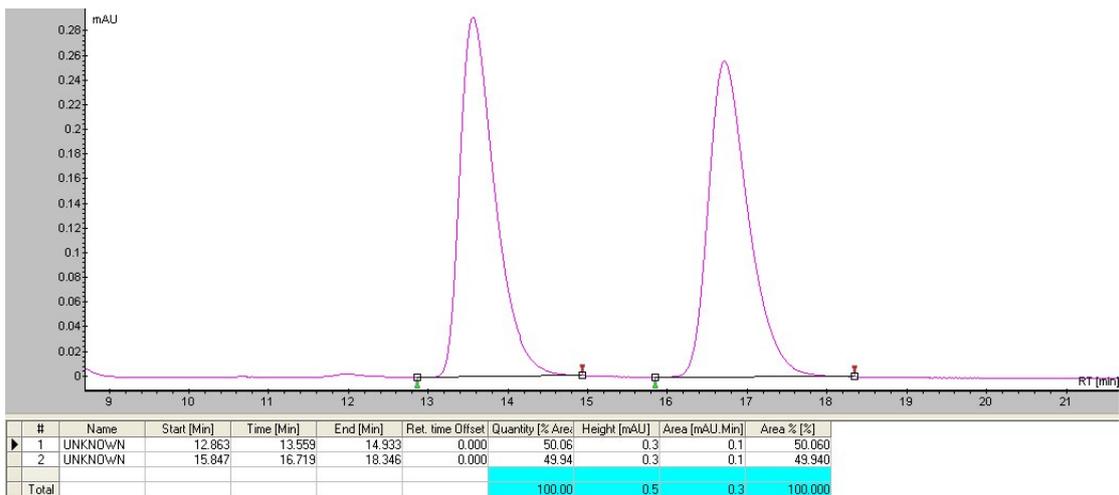
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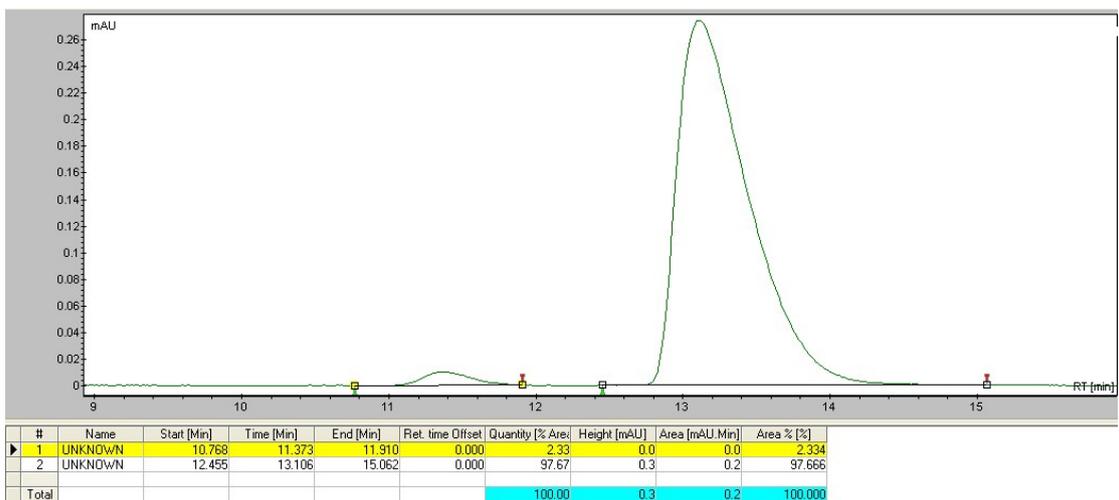
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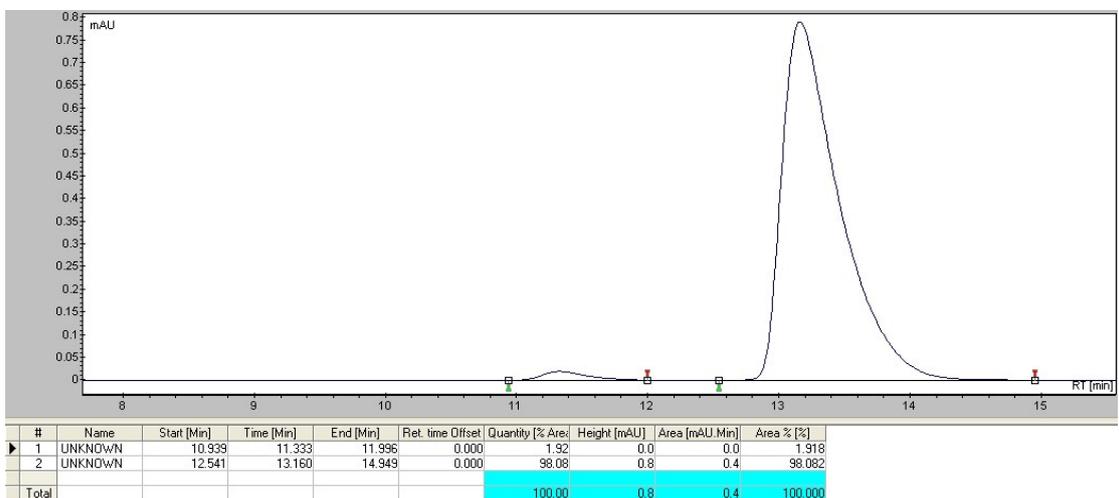
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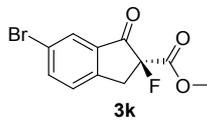
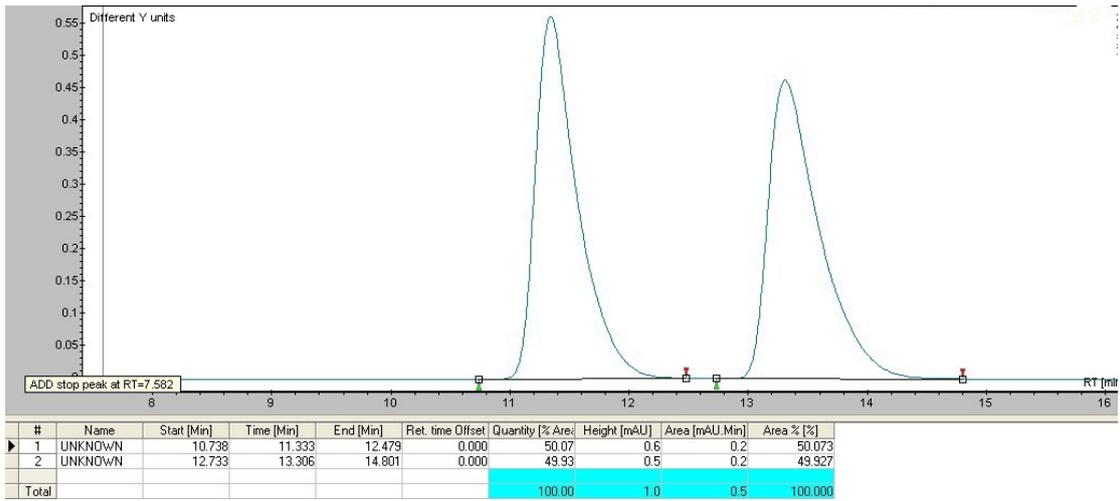
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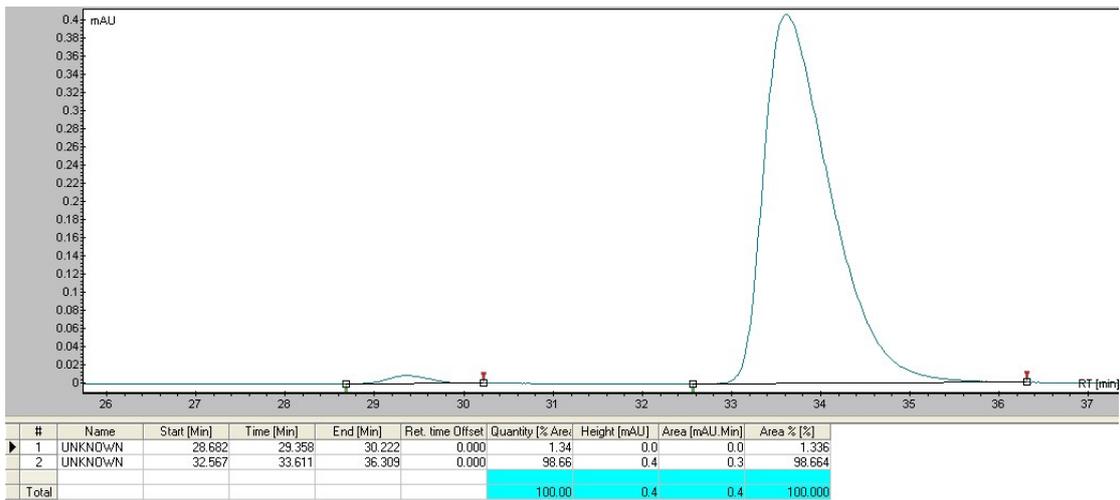
In continuous flow condition



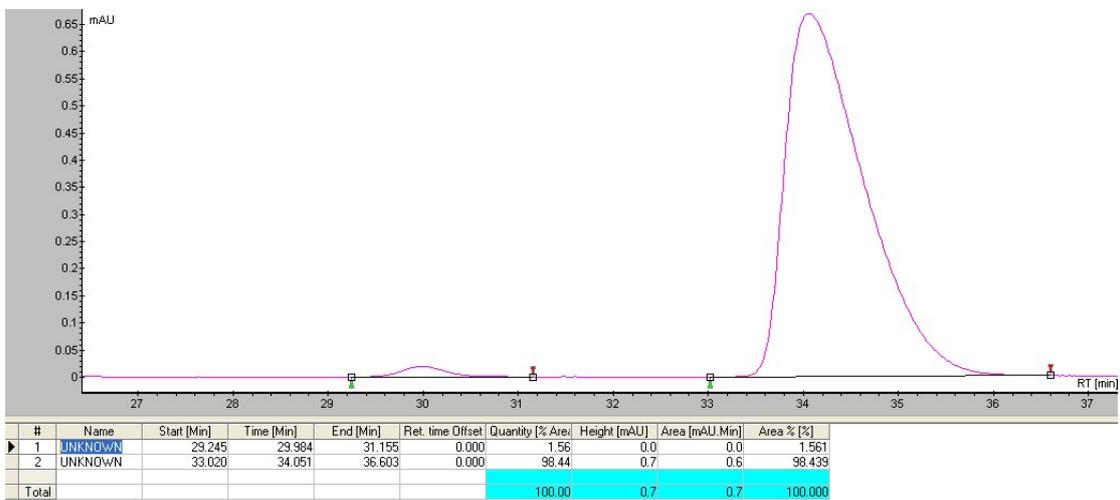
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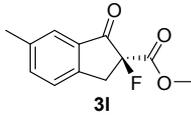
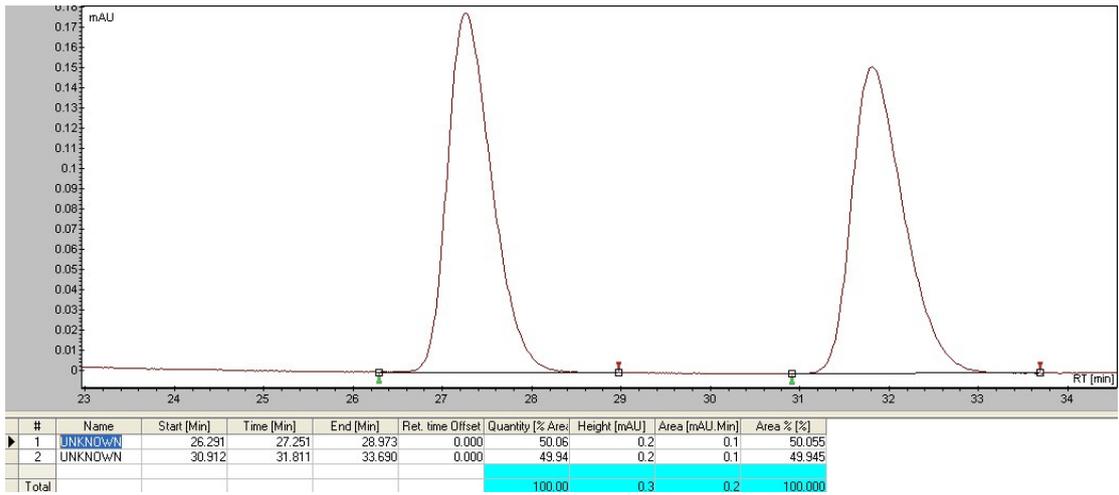
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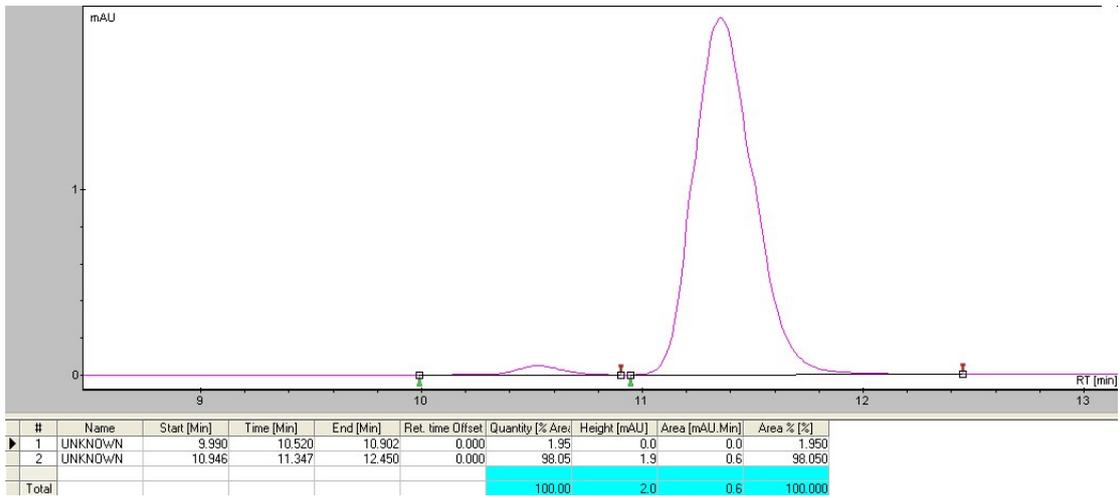
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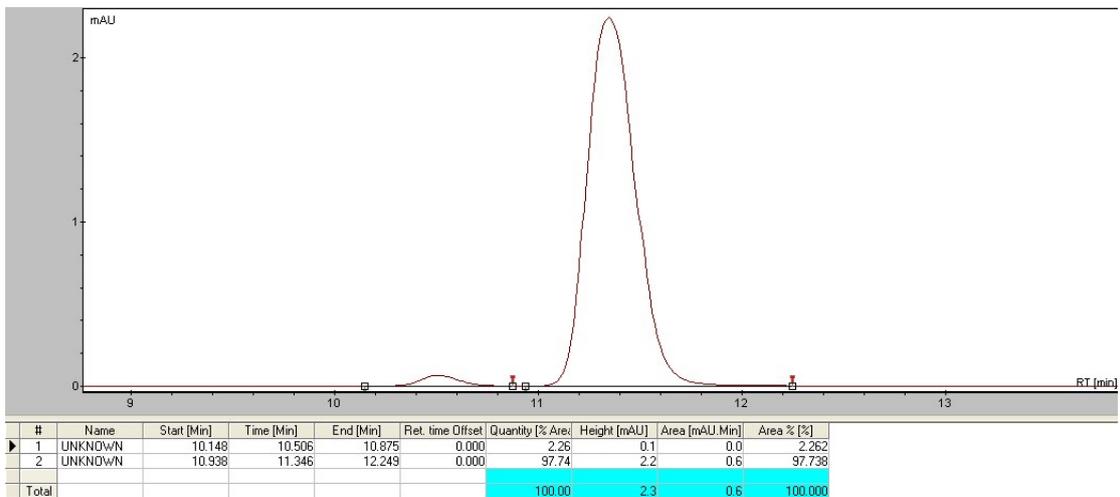
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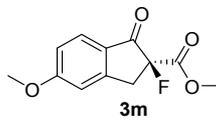
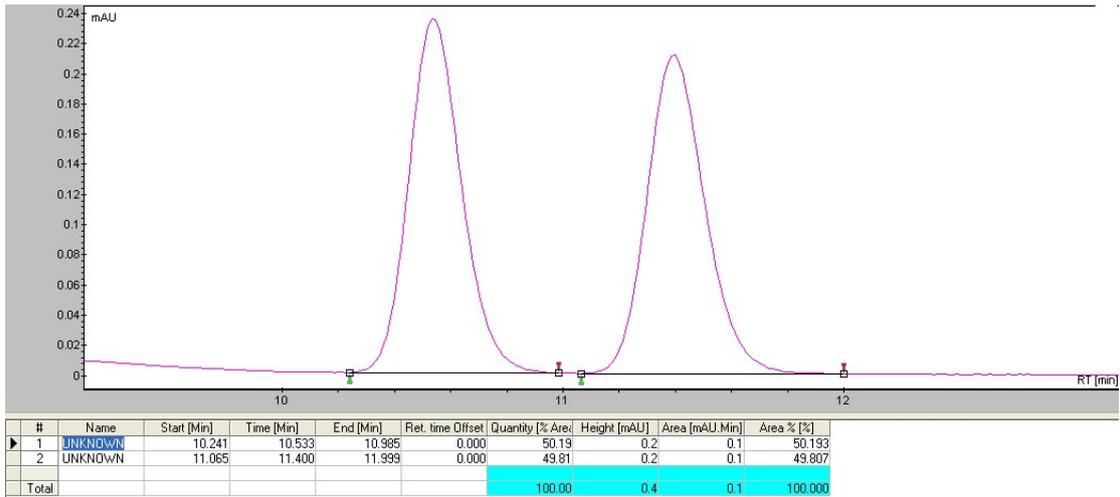
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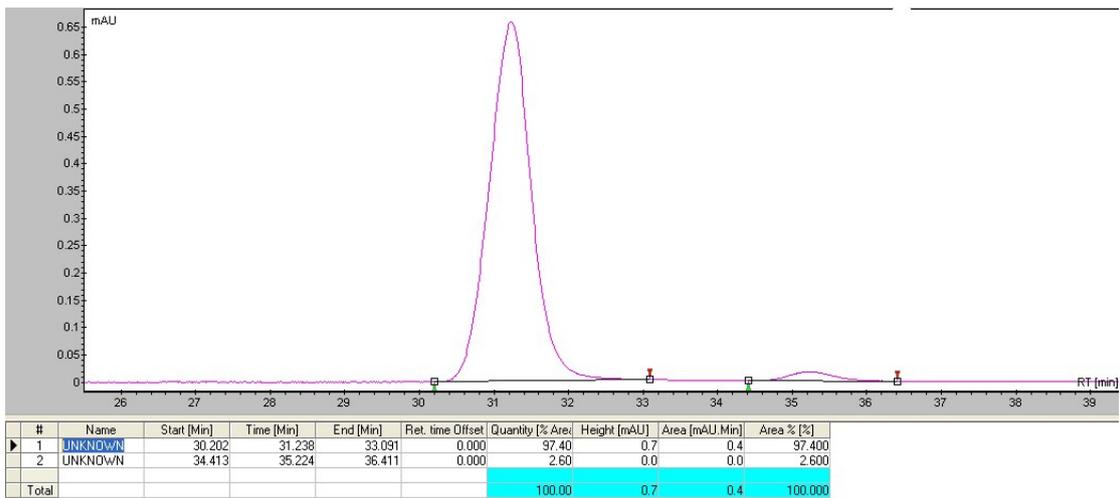
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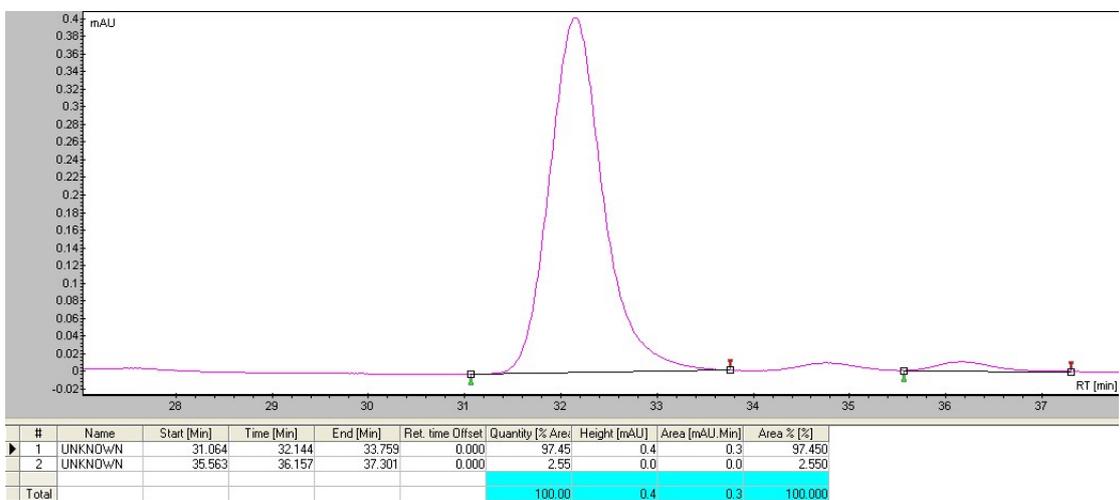
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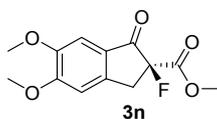
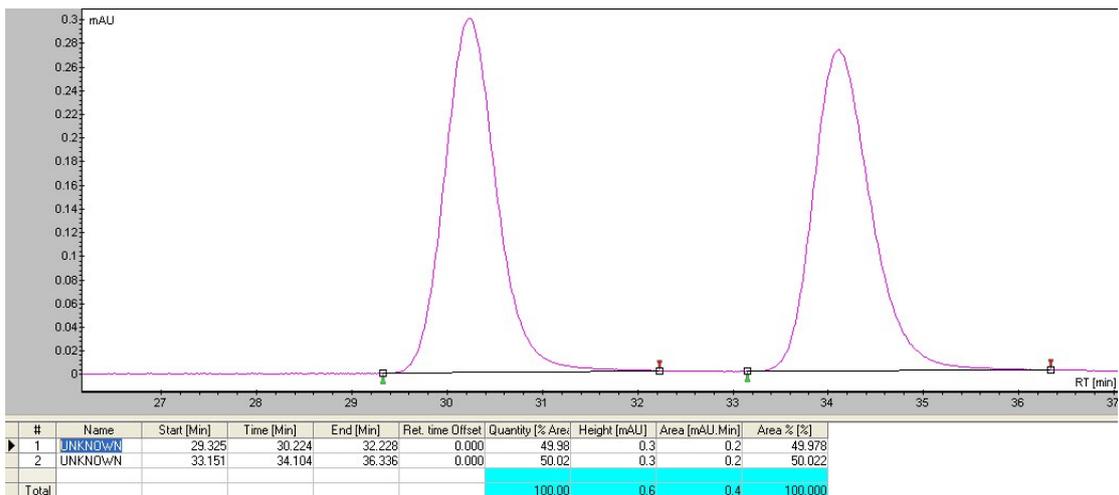
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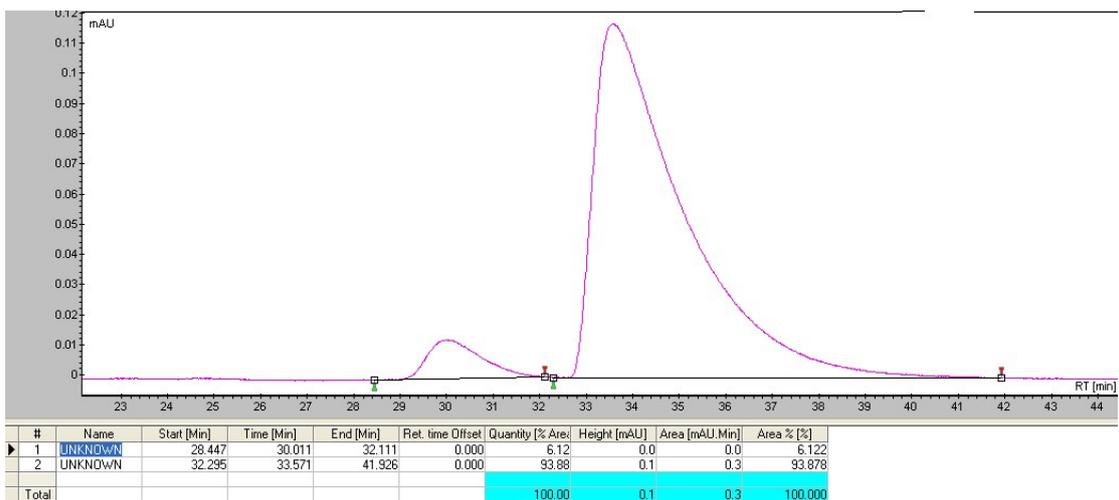
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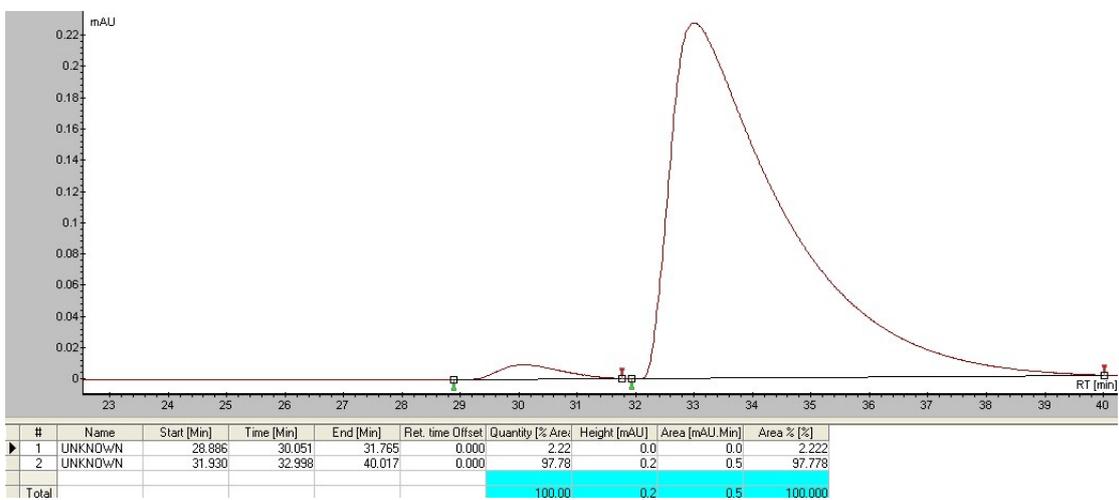
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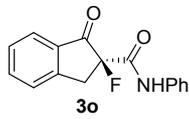
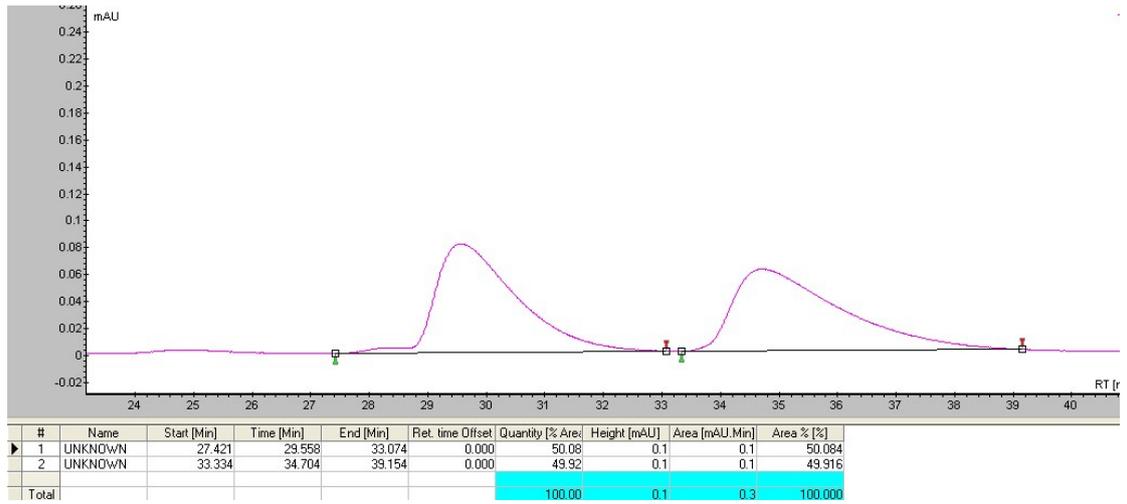
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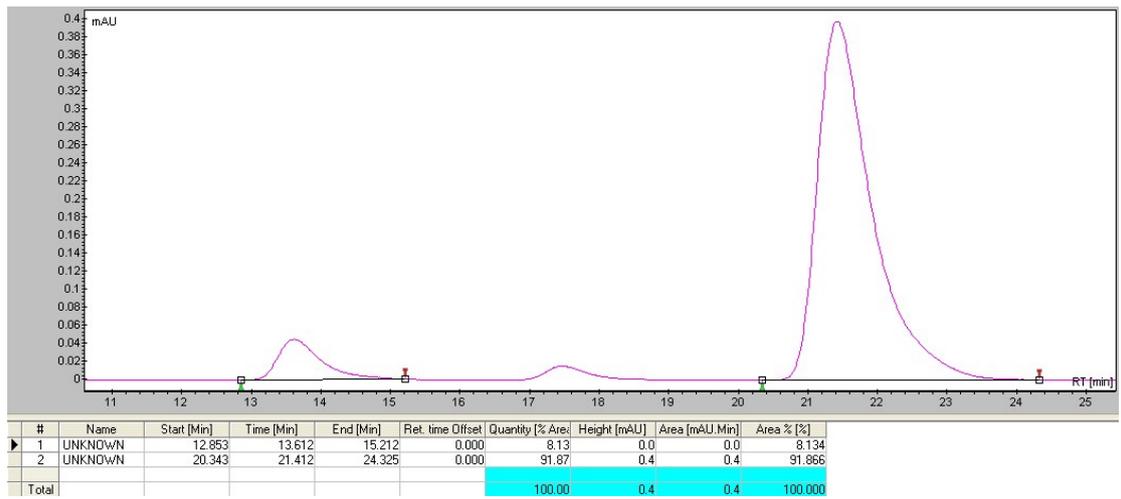
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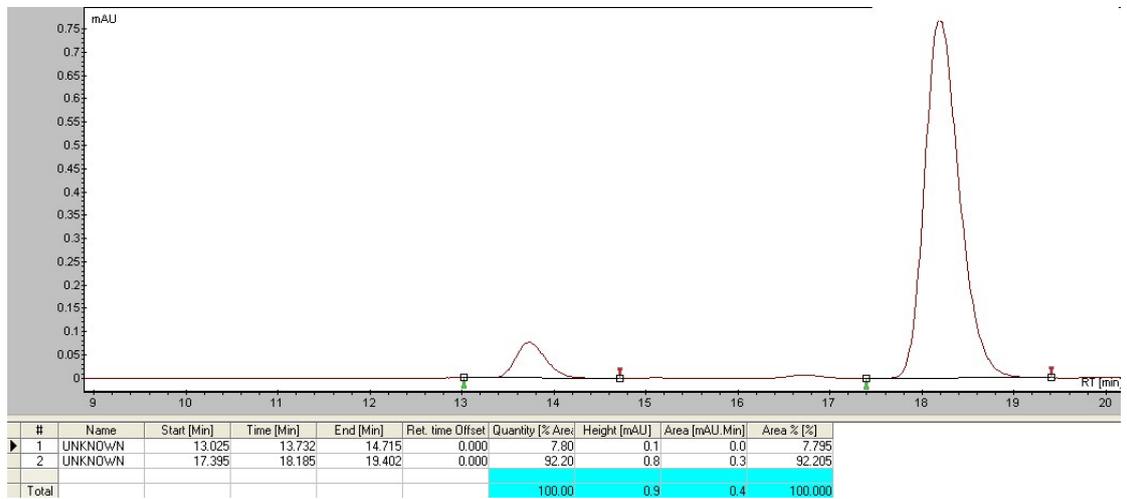
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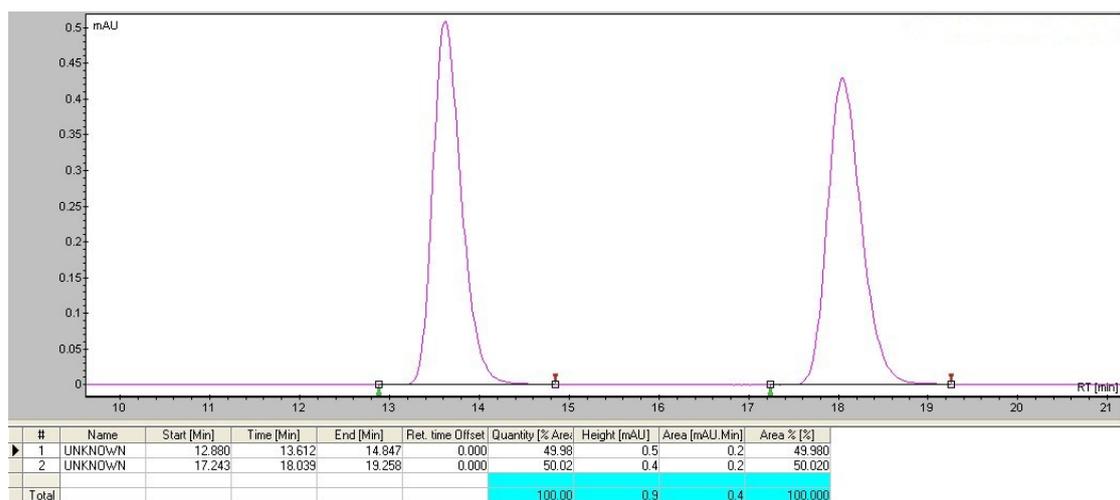
In batch condition



In continuous flow condition



rac



8. References

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