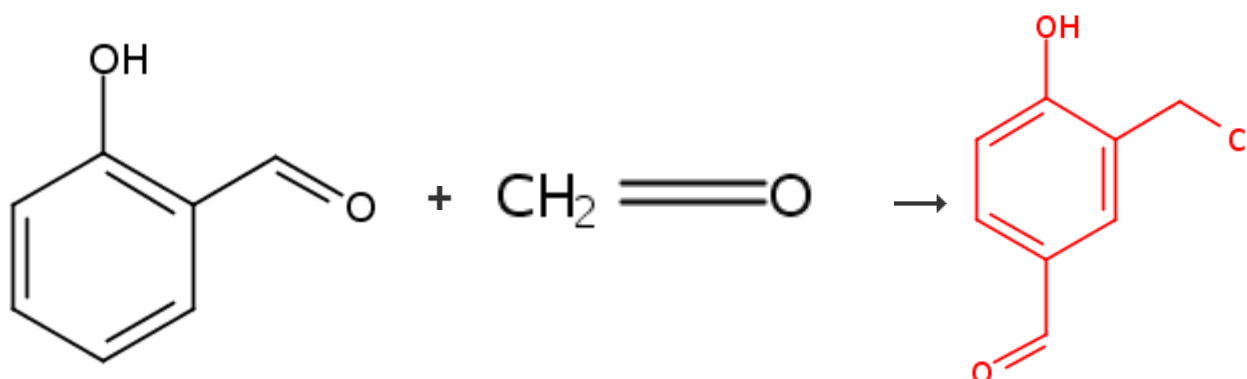


## 1. Single Step



73%

[Overview](#)

## Steps/Stages

1.1 R:HCl, 48 h, rt

## Notes

sealed flask used, paraformaldehyde used,  
Reactants: 2, Reagents: 1, Steps: 1, Stages:  
1, Most stages in a one step: 1

## References

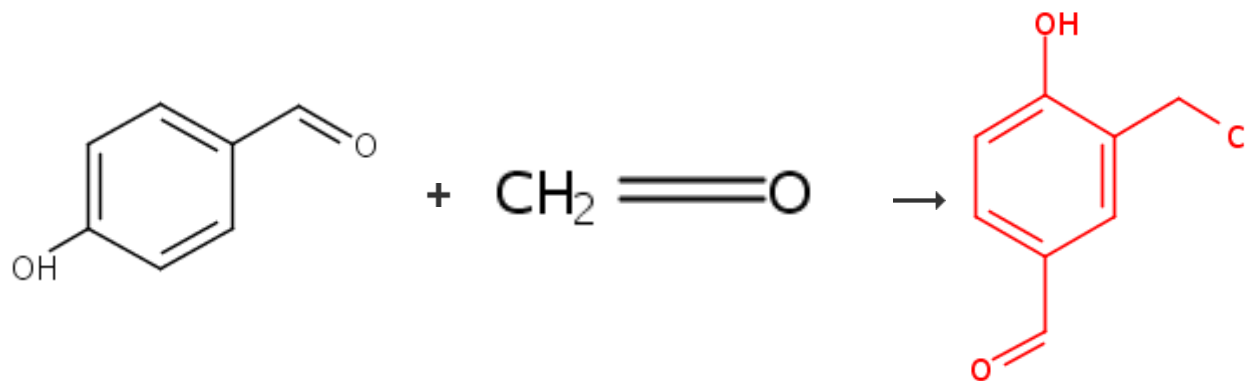
[Synthesis and fluorescence properties of a new fluorescent compound for determination of  \$\text{Co}^{2+}\$  selectively](#)

By Li, Rui-ji et al

From Jingxi Huagong Zhongjianti, 43(5), 46-50; 2013

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## 2. Single Step



56%

[Overview](#)

## Steps/Stages

## Notes

1.1 R:

S:H<sub>2</sub>O

Reactants: 2, Reagents: 1, Solvents: 1, Steps: 1, Stages: 1, Most stages in any one step: 1

**References**

[Diastereoselective oxy-Michael addition of a new chiral water equivalent to electron deficient alkenes - Synthesis of 1,2-amino alcohols and beta-hydroxy malonates](#)

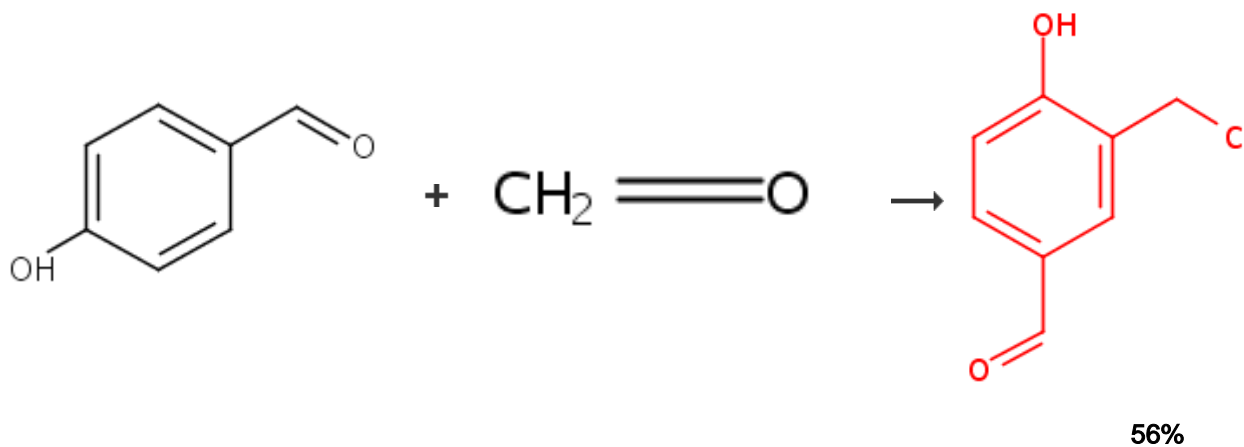
By Buchanan, David Jonathan

From null, , No pp.; 2005

**Experimental Procedure**

3-Chloromethyl-4-hydroxy benzaldehyde, 174: To a stirring solution of HCl (153 mL, 10.2 N, 1.56 mol) and formaldehyde (17 mL, 40 % w/w aq. soln.) at rt was added 4-hydroxy benzaldehyde 173 (23.2 g, 0.19 mol). The reaction mixture was stirred at 65 degC for 3.5 hours before the resulting pink precipitate was filtered and washed with water (20 mL). Water (20 mL) and EtOAc (40 mL) were added and the aqueous layer was separated and extracted with EtOAc (3 x 40 mL). The combined organic layers were washed with brine (15 mL), dried with MgSO<sub>4</sub>, filtered and concentrated in vacuo. Purification by trituration with DCM (60 mL) gave the title compound 174 (18.3 g, 56 %) as a white solid.

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**3. Single Step****Overview****Steps/Stages**

1.1 R:HCl, S:H<sub>2</sub>O, 3.5 h, 65°C

**Notes**

Reactants: 2, Reagents: 1, Solvents: 1, Steps: 1, Stages: 1, Most stages in any one step: 1

**References**

[A short stereoselective synthesis of \(R\)-salmeterol](#)

By Buchanan, David J. et al

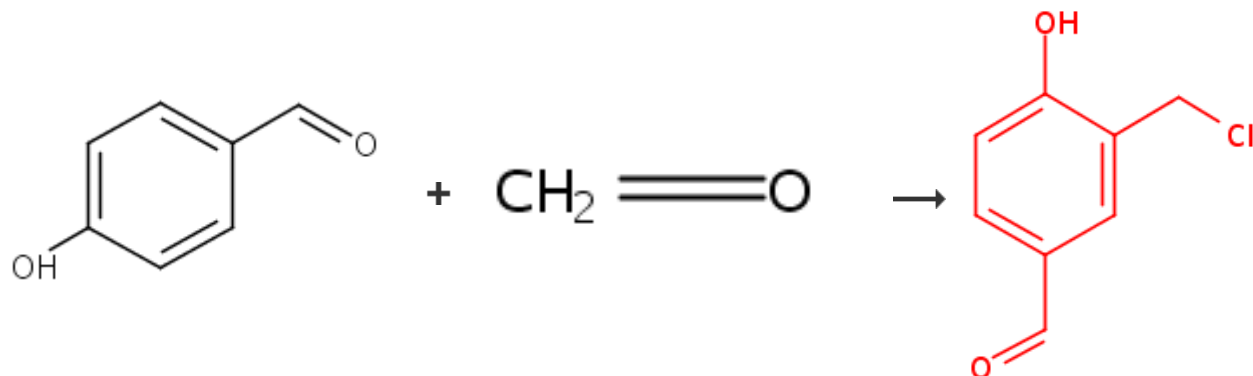
From Synlett, (12), 1948-1950; 2005

**Experimental Procedure**

The synthesis of the key Michael acceptor required four steps beginning from 4-hydroxybenzaldehyde (5). First, chloromethylation using formaldehyde and concentrated hydrochloric acid at 65 °C for 3.5 hours afforded the chloromethyl arene 6 in 56% yield.

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**4. Single Step**

[Overview](#)**Steps/Stages**

1.1 R:HCl, S:H<sub>2</sub>O, 10 h, 20-25°C

**Notes**

regioselective, paraformaldehyde used,  
Reactants: 2, Reagents: 1, Solvents: 1, Steps:  
1, Stages: 1, Most stages in any one step: 1

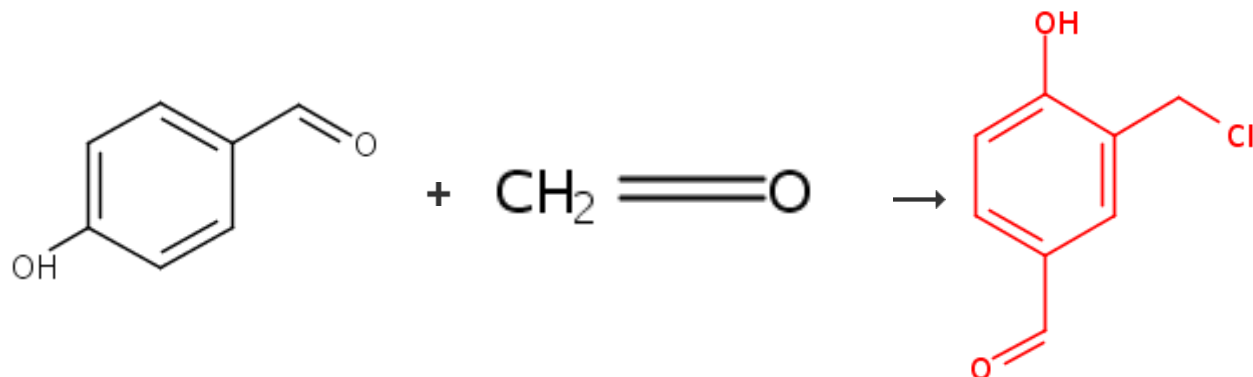
**References**

Process for synthesizing salbutamol and its sulfate salt via chloromethylation, hydrolysis, propylidene protection, epoxidation, aminolysis ring-opening reaction, deprotection of propylidene, and salification

By Yang, Qiaoming

From Faming Zhuanli Shenqing, 103951568, 30 Jul 2014

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**5. Single Step**[Overview](#)**Steps/Stages****Notes**

1.1 R:HCl, R:H<sub>2</sub>SO<sub>4</sub>

paraformaldehyde used, literature prepn., gaseous HCl used, Reactants: 2, Reagents: 2, Steps: 1, Stages: 1, Most stages in any one step: 1

#### References

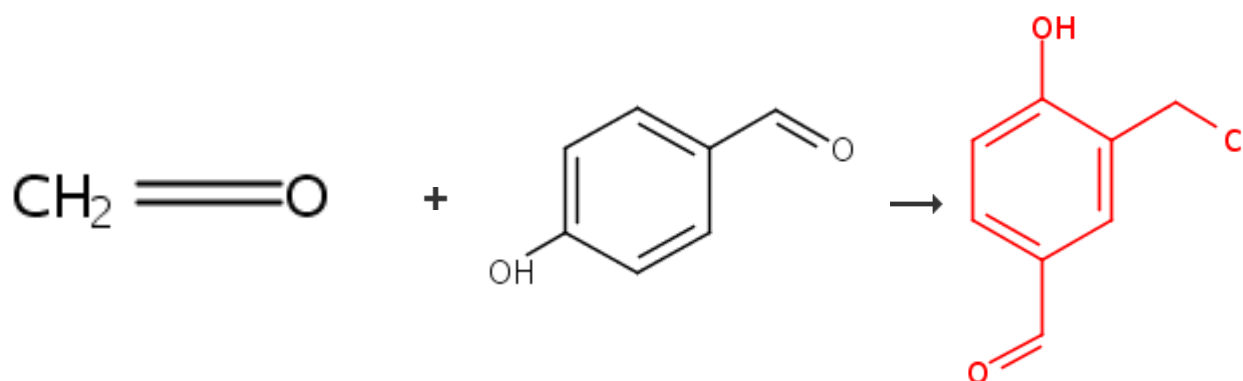
[Enantioselective synthesis of \(R\)-salmeterol employing an asymmetric Henry reaction as the key step](#)

By Guo, Zong-Liang et al

From Tetrahedron: Asymmetry, 22(13), 1395-1399; 2011

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#### 6. Single Step



#### Overview

#### Steps/Stages

1.1

#### Notes

Reactants: 2, Steps: 1, Stages: 1, Most stages in any one step: 1

#### References

[Preparation of phosphorylated derivatives of L-DOPA and compositions and methods for increasing the melanin content in mammalian skin and hair](#)

By Pawelek, John M. and Osber, Michael P.

From PCT Int. Appl., 9012016, 18 Oct 1990

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