

Langlois Reagent: An Efficient Trifluoromethylation Reagent

K. Anusha

Megha Mahesh

Shikha

T. M. Rangarajan*

Sharda Pasricha*

Department of Chemistry, Sri Venkateswara College, University of Delhi,
New Delhi-110021, India
spasricha@svc.ac.in, rt.m@svc.ac.in

Received: 15.12.2022

Accepted after revision: 01.02.2023

Published online: 01.02.2023 (Accepted Manuscript),

20.02.2023 (Version of Record)

DOI: 10.1055/a-2024-1382; Art ID: SO-2022-12-0076-SPOT



License terms:

© 2023. The Author(s). This is an open access article published by Thieme under the terms of the Creative Commons Attribution License, permitting unrestricted use, distribution and reproduction, so long as the original work is properly cited. (<https://creativecommons.org/licenses/by/4.0/>)

Abstract **Key words** Langlois reagent, trifluoromethylation, free radical, fluoroorganic synthesis, sodium trifluoromethanesulfonate or sodium triflate

The fluorine atom has a grand reception in pharmaceutical, material, and agrochemical industries as it dramatically alters the physical, chemical, and metabolic properties of organic compounds in its presence. Hence, its incorporation as an atom or fluorine-containing functional groups in organic molecules remains a huge interest among the organic chemists.¹ Among the fluorine-containing functional groups, the trifluoromethyl (CF₃) group is the most common group that could improve molecular properties and hence predominantly found in pharmaceutical substances. Therefore, the development of novel methods to build the C–CF₃ bond is of great interest, and several other reagents have been developed.²

Among the other available trifluoromethylating reagents, such as Togni, Umemoto, Ruppert–Prakash reagents, etc., Langlois reagent (CF₃SO₂Na) has been extensively focused in the past few decades due to its commercial availability, inexpensiveness, stability, and, importantly, its capability of transferring the CF₃ group into a large variety of substrates via both electrophilic and free-radical mechanistic pathways.^{3,4} Interestingly, this reagent can also be used to install SCF₃, SOCF₃, etc. functions into organic compounds.⁵



Anusha K. (first from left) and **Megha Mahesh** (second from left) are undergraduate life science students, and **Shikha** is an undergraduate chemistry honors student in the Department of Chemistry at Sri Venkateswara College.

Prof. Sharda Pasricha (first from right) is currently working as professor in the Department of Chemistry, Sri Venkateswara College, University of Delhi, India. She has nearly 25 years of teaching experience at the undergraduate level. She has published nearly 20 research articles in journals of national and international reputation. She was a recipient of the prestigious ‘Distinguished Teacher Award’ from University of Delhi in 2009. She has authored two books and several e-modules for UG students. Her current research interests include green chemistry, heterogeneous catalysis, and synthesis of biologically relevant heterocycles.

Dr. T. M. Rangarajan is currently working as assistant professor of chemistry and has nearly 10 years of research experience and 5 years of teaching experience. He has published 23 research articles of international repute and has three national patents and one international patent to his name. His research interests include electrochemical perfluorination, electro-organic synthesis, organic synthesis, cross-coupling methodologies, and medicinal chemistry.

In 1991, the sodium trifluoromethanesulfonate (NaSO₂CF₃) reagent was first introduced by the Langlois group for the introduction of the trifluoromethyl group in an aromatic system and the reagent was first prepared from trifluoromethylchloride and sodium dithionite⁶ as shown in Scheme 1. However, this reagent was unexplored for fifteen years. In