



HAL
open science

Expanding Radical Chloropentafluorosulfanylation of Alkynes

Thi Mo Nguyen, Lucas Popek, David Matchavariani, Nicolas Blanchard, Vincent Bizet, Dominique Cahard

► **To cite this version:**

Thi Mo Nguyen, Lucas Popek, David Matchavariani, Nicolas Blanchard, Vincent Bizet, et al. Expanding Radical Chloropentafluorosulfanylation of Alkynes. *Organic Letters*, 2024, 26 (1), pp.365-369. <10.1021/acs.orglett.3c04043>. <hal-04631759>

HAL Id: hal-04631759

<https://cnrs.hal.science/hal-04631759v1>

Submitted on 2 Jul 2024

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire HAL, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.



HAL Authorization

Expanding Radical Chloropentafluorosulfanylation of Alkynes

Thi Mo Nguyen, Lucas Popek, David Matchavariani, Nicolas Blanchard, Vincent Bizet,* and Dominique Cahard*

Cite This: <https://doi.org/10.1021/acs.orglett.3c04043>

Read Online

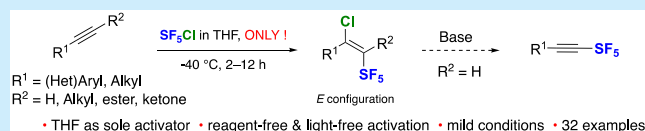
ACCESS |

Metrics & More

Article Recommendations

Supporting Information

ABSTRACT: The chloropentafluorosulfanylation of alkynes is a delicate but crucial operation for accessing SF₅-alkynes that serve as substrates in numerous transformations. Dolbier's procedure using Et₃B/O₂ was the most efficient approach, while recent efforts make use of other initiators and light activation. We found that THF, as a single stimulus, is sufficient to trigger the reaction of SF₅Cl with alkynes. We determined the configuration of Cl/SF₅ products and



clarified the structure of side-products.

In the field of organofluorine chemistry, the search for new chemical motifs offering improved physicochemical or biological properties is very active. To this end, combining fluorine with sulfur has given rise to the blossoming of compounds featuring either (Het)ArSF₄ or sulfur pentafluoride SF₅ groups; these motifs are considered truly emerging and are currently generating an abundance of bibliographic data.^{1–3} As a rationale for the preparation of pentafluorosulfanylated molecules, particular mention should be made of its larger volume in comparison with the CF₃ group, a square pyramidal geometry, its high electronegativity and high lipophilicity, a metabolic robustness and chemical stability.^{4–8} The construction of SF₅-containing compounds by means of inorganic gaseous pentafluorosulfanyl chloride reagent SF₅Cl constitutes the main synthetic approach.⁹ Besides, the SF₅ group can be built from thiols and disulfides by oxidative fluorination to afford SF₅-(het)aromatics.¹⁰ SF₅-Alkynes are central in downstream functionalizations toward a wide range of heterocyclic and acyclic compounds (Figure 1A).¹¹ SF₅-Alkynes are typically obtained in a two-step sequence from terminal alkynes by chloropentafluorosulfanylation with SF₅Cl followed by HCl elimination with the aid of a base (Figure 1B). Ethynylbenziodoxolone reagents are also substrates for the reaction of SF₅Cl under blue light irradiation.¹² Early radical additions of SF₅Cl onto unsaturated carbon compounds were achieved through thermal or photochemical processes, requiring special equipment. In 2002, Dolbier's breakthrough consisted of the use of BEt₃/O₂ as a catalytic initiator of free radical chain addition of SF₅Cl onto alkenes and alkynes.¹³ The reactions were performed in the liquid phase below the boiling point of SF₅Cl (bp –21 °C) in standard glassware. With terminal alkynes, the chloropentafluorosulfanylation is regio- and stereoselective, leading to the *E*-alkenes, although the configuration was not definitively ascertained. Besides the desired product **2a** obtained from phenyl acetylene **1a**, a 2:1 adduct (**3a**) was concomitantly formed (**2a**/**3a** = 49:27) (see Table 1). So far, this synthetic route was the method of choice

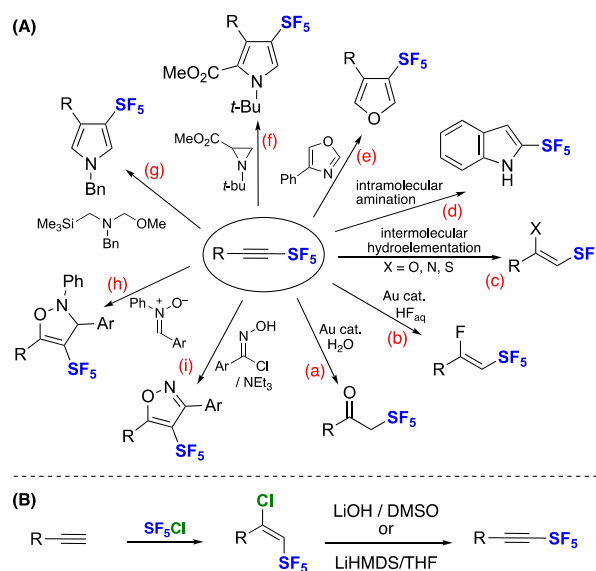


Figure 1. (A) Applications of SF₅-alkynes in (a) Gold-catalyzed hydration,¹⁴ (b) Gold-catalyzed hydrofluorination,¹⁵ (c) Hydroelementation,¹⁶ (d) Intramolecular amination,¹⁷ (e) Diels–Alder reaction,¹⁸ (f, g) Pyrrole syntheses,^{19,20} (h) Isoxazoline synthesis,²¹ (i) Isoxazole synthesis.²² (B) Synthesis of SF₅-alkynes from SF₅Cl and terminal alkynes.

to prepare Cl/SF₅-alkenes and subsequently SF₅-alkynes. Nevertheless, the pyrophoric properties of triethylborane led Paquin's group to evaluate amine-borane complexes as stable

Received: November 30, 2023

Revised: December 26, 2023

Accepted: December 29, 2023

Table 1. Screening of Solvents^a

Entry	Solvent	Conv.	2a/3a yield (%) ^b
1	THF	100	92 (99)/(<1)
2	Et ₂ O	100	(72)/(9)
3	THP	100	73 (94)/(6)
4	Hexane	0	0/0
5	CH ₂ Cl ₂	54	0/0
6	Toluene	30	0/0
7	CH ₃ CN	30	0/0
8	THF/dark	100	(86)/(1)
9	THF/dark/O ₂ free	90	(90)/0
10	THF/dark/O ₂ free/Nalgene flask	100	93 (97)/(3)
11	THF nondistilled	100	(92)/(8)

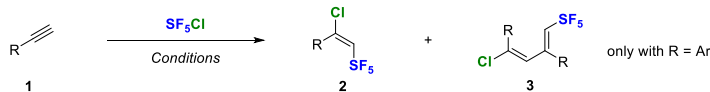
^aReaction conditions: **1a** (1 mmol), solvent (2 mL), $-40\text{ }^{\circ}\text{C}$, 2 h.

^bYields of isolated products (the number in brackets refers to calculated yields by ¹⁹F NMR using PhSCF₃ as internal standard and by GC-MS).

radical initiators.^{23,24} With the same motivation, they developed an electron donor–acceptor (EDA)-complex with visible light irradiation.²⁵ The latest development by this group was the activation by means of a blacklight compact fluorescent lamp.²⁶ This synthetic approach represents noticeable progress en route to a simple procedure, but still a light is necessary. Apart from the fact that the current methods require an activator, they are also not entirely satisfactory in terms of yields and substrates scope; moreover, side-products, notably the 2:1 adduct and the *Z* stereoisomer, cause purification difficulties. Interestingly, Tlili noticed that SF₅Cl in hexane can react with **1a** in absence of BEt₃ albeit in lower yield (58% versus 73% with BEt₃); however, this remarkable insight has not been further explored.²⁷ Similar observations were made in triethylborane-free reactions of SF₅Cl in CH₂Cl₂ (<2% yield)²⁸ and SF₅Cl in Et₂O (13% yield).²⁹ In the same vein, (het)ArSF₄Cl reagents reacted without any activation in cyclopentyl methyl ether or in Et₂O.^{30,31} This literature survey shows that the current mainstream in chloropentafluorosulfanylation of alkynes is to look for alternative radical initiators or new methods by light activation. This prompted us to search for a safe, simple and efficient protocol to generate an extensive portfolio of novel SF₅ molecules. We hypothesized that SF₅Cl could react in high yields under reagent-free and light-free conditions by exploiting the intrinsic susceptibility of ethereal solvents to form hydroperoxides that could thereafter generate the SF₅ radical. We herein report a paradigm shift in which simplicity goes hand-to-hand with efficiency in the regio- and stereoselective chloropentafluorosulfanylation of alkynes. In addition, the structural determination of Cl/SF₅-alkenes and the secondary products was carried out.

Our study began by assessing the stability of SF₅Cl solutions. SF₅Cl gas was condensed in a sealed flask at $-40\text{ }^{\circ}\text{C}$, and the solvent was added (see Supporting Information (SI) for detailed procedures). Since Dolbier's work with the aid of triethylborane,¹³ hexane has remained the most widely used solvent, even when SF₅Cl is homemade, and solutions of SF₅Cl in hexane are reported as bench-stable.^{32–34} The situation is different for THF solutions in which SF₅Cl is responsible for the polymerization of THF rapidly forming a gel at rt and

within 12 h at $-20\text{ }^{\circ}\text{C}$. This catalyzed polymerization slows down sufficiently at $-40\text{ }^{\circ}\text{C}$ and stops at $-78\text{ }^{\circ}\text{C}$. Our model reaction used phenylacetylene **1a** and SF₅Cl to allow comparison with all of the reported syntheses. The temperature was set at $-40\text{ }^{\circ}\text{C}$ to minimize THF polymerization. Besides the desired Cl/SF₅-alkene **2a**, we detected 2:1 adduct **3a** (Table 1). Our hypothesis turned out to be correct, as the Cl/SF₅-alkene **2a** was obtained in 92% yield after purification (entry 1). Other ethereal solvents, Et₂O and tetrahydropyran (THP), delivered **2a** in more than 72% calculated yields, but the reactions were not so clean since **3a** was detected by GC-MS and ¹⁹F NMR (entries 2, 3). In hexane at $-40\text{ }^{\circ}\text{C}$, substrate **1a** was fully recovered after 2 h, while dichloromethane, toluene, and acetonitrile gave conversions of 30–54% but not into fluorinated products (entries 4–7). Note that, in the dark, the reaction still proceeded in full conversion to end up with 86% of **2a** (entry 8). Degassing the solution of **1a** in THF by the freeze–pump–thaw method allowed a very clean reaction leading to 90% of **2a** as single product (entry 9). This result could mean that dissolved oxygen plays no role, but this may be a hasty conclusion, given that SF₅Cl added after degassing may contain dissolved oxygen. An experiment carried out in Nalgene instead of glass gave almost identical product distribution and yield indicating that glass has no contribution to the reaction mechanism (entry 10). Finally, using nondistilled THF appeared detrimental to the selectivity, with an increased amount of **3a** (entry 11). This case study with phenylacetylene allowed a clear comparison with the methods currently available (Scheme 1A). A maximum yield of 49% was recorded with the previously reported methods, while we significantly improved the isolated yield to 92% (Scheme 1A). During this first phase of our investigation, 2:1 adduct **3a** was crystallized and its single-crystal X-ray diffraction (SCXRD) analysis confirmed the structure proposed by Dolbier, also enabling us to prove the *trans* stereochemistry of each vinylic radical (Figure 2).¹³ With these conditions in hand, we applied our simple protocol in THF to aromatic and aliphatic alkynes already reported in the literature for chloropentafluorosulfanylation (Scheme 1B). The present method compared favorably with the literature, affording higher yields of Cl/SF₅-alkenes. Starting from 4-octyne, we obtained an excellent yield by GC-MS but due to high volatility of **2j** we only obtained it in 74% isolated yield. Radical addition of SF₅Cl to alkynes was supposed to produce the *E* stereoisomers, although this was never firmly established by earlier contributors in the field. A large-scale experiment was carried out that yielded 1.9 g of product *E*-**2e** as a solid suitable for SCXRD to definitively ascertain the *E* configuration of this family of compounds. We could photoisomerized *E*-**2e** into *Z*-**2e** at least partially (see SI). Having access to both *E*- and *Z*-stereoisomers of **2e**, we have deduced by ¹⁹F NMR for the characteristic AB₄ pair of quintet and doublet signals that the doublet is 3.1 ppm upper field for the *Z* stereoisomer in comparison with the *E* stereoisomer (65.9 ppm for *Z*-**2e** versus 69.0 ppm for *E*-**2e**). Clarification of the configuration of Cl/SF₅-alkenes was necessary because some misidentifications were noted in the literature. For example, **2l** was isolated as the *E*-isomer and not the *Z*-isomer as it was reported (see SCXRD of **2l** in Figure 2 and NMRs in the SI).²⁵ In addition, *E*-**2m** is the product of the chloropentafluorosulfanylation of 5-phenyl-1-pentyne **1m** whatever the method (C or the present one). The reaction with **1m** does not give a mixture of *Z* + *E*-**2m** (75:25)²⁵ but rather *E*-**2m** along with the

Scheme 1. Chloropentafluorosulfanylation of Alkynes^a

A) Phenylacetylene as model alkyne for comparison with all existing methods.

 2a	Method A (Et ₃ B/O ₂ , hexane, -30 °C, 0.5 h)	2a	3a	Dolbier 2002/2015 Tlili 2022 Borggraeve 2023 Paquin 2020 Paquin 2021 Paquin 2023 Tlili 2022 Bizet / Cahard 2023
	Method A' (Et ₃ B/O ₂ , hexane, rt, 0.5 h)	49%	27%	
	Method A'' (Et ₃ B/O ₂ , hexane, -94 °C, 3 h)	30% (73%)	not mentioned	
	Method B (Amine-borane/O ₂)	15% (18%)	26% (32%)	
	Method C (EDA complex/light)	(23%)	5%	
	Method D (blacklight irradiation in AcOEt)	33% (51%)	0%	
Method E (without activation in hexane)	25% (E/Z >97:3)	not mentioned	not mentioned	
Method E (with activation in hexane)	(58%)	not mentioned	not mentioned	
This work (THF only, -40 °C)	92% (99%)	0%	0%	Bizet / Cahard 2023

B) Comparison with reported aromatic and aliphatic alkynes.

 2b	 2c	 2d	 2e	 2f	 2g	 2h
Method A 50%	Method A 27% ^b	Method A 35% ^c	Method A 38–88% ^c	Method A 49% ^c	Method A 44%	Method A 61–76%
Method E (41%)	This work 95% (100%)	This work 88% (100%)	This work 99% (100%)	This work 77% (92%)	Method E (25%)	This work 68% (95%)
This work 88% (100%)			(1.9 g of 2e)		This work 83% (98%)	
 2i	 2j	 2k	 4m	 2l	 4m	
Method A 49%	Method A 93%	Method D 66%	Method A 91%	Method A 91%	Method A (68:0:32 E-2m/Z-2m/4m) ^c	
Method E (40%)	This work 74% (95%)	This work 72% (94%)	Method C 86% ^d	Method C 86% ^d	Method C 84% (75:25 postulated E-2m/Z-2m)	
This work 51% (94%)			This work 94% (100% E)	This work 94% (100% E)	This work 59% (86:0:14 E-2m/Z-2m/4m)	

C) New alkyne substrates engaged in the chloropentafluorosulfanylation reaction. 2n–af are new compounds.^e

 2n	 2o	 2p	 2q	 2r	 2s	 2t
74% (95%)	54% (93%) + (7%) 3o	92% (100%)	100%	48% (60%) + (19%) 3r	47% (97%)	31% (38%) + (31%) 3t
	(74%) + (13%) 3o with BEt ₃ in hexane	73% with BEt ₃ in DCM ^f	54% with BEt ₃ in hexane	(60%) + (20%) 3r with BEt ₃ in hexane		messy with BEt ₃ in hexane
 2u	 2v	 2w	 2x	 2y	 2z	 2aa
63% (75%) + (12%) 3u	31% (34%) + (33%) 3v	56% (83%)	77% (97%)	73% (95%)	13% (15%)	100%
messy with BEt ₃ in hexane	0% with BEt ₃ in DCM ^f	0% with BEt ₃ in DCM ^f	0% with BEt ₃ in DCM ^f		0% with BEt ₃ in DCM ^f	53% with BEt ₃ in DCM ^f
 2ab	 2ac	 2ad	 2ae	 2af	 2ag	 2ah
36% (54%)	86% (96%) ^f	95% (97%)	71% (95%)	70% (79%)	0% with BEt ₃ in DCM ^f	24% (40%) with BEt ₃ in AcOEt
		(96%) with BEt ₃ in hexane	(25%) with BEt ₃ in hexane			

^aSee the SI for detailed experimental procedures. Yield of isolated product (the number in parentheses refers to calculated yield by ¹⁹F NMR using PhSCF₃ as internal standard). References of methods: A,^{13,21,16} A',²⁷ A'',³⁵ B,^{23,24} C,²⁵ D,²⁶ E.²⁷ ^bOverall yield of the SF₅-alkyne over two steps after HCl elimination. ^cReaction performed in DCM from -40 °C to rt. ^dThe product was misidentified as Z. ^eUnless otherwise noted, reactions have been performed in THF at -40 °C for 2 h. ^fReaction performed at -78 °C.

cyclic nonchlorinated compound **4m** (68:32 in DCM with catalytic Et₃B or 86:14 in THF without initiator). To further demonstrate the greater reactivity of alkynes in THF versus Dolbier's method, we evaluated both conditions with new alkynes (Scheme 1C). Again, the superiority of our method has been clearly illustrated with variously decorated aryl alkynes featuring alkyl, ester, ether, or nitro groups (**2n–v**) as well as with (het)aryl alkynes (**2w–x**). For instance, aromatics

featuring electron-withdrawing groups do not react well using Dolbier's conditions while the corresponding Cl/SF₅-addition products were obtained using the present method (e.g., **2t–v**); idem for 3-ethynylpyridine and thiophene derivatives **2w** and **2x**. Bis-alkyne **1y** afforded **2y** in a 73% isolated yield. Addition of SF₅Cl onto silylated terminal alkynes was unknown, and the only reported method to produce a silylated SF₅-alkyne involved SF₅Br, which is not commercially

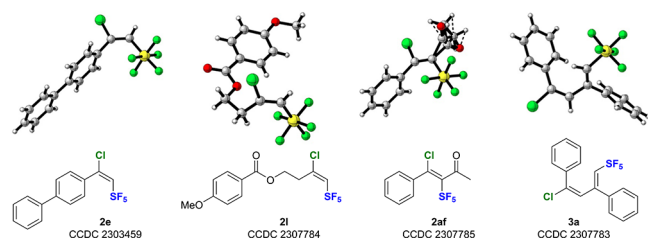


Figure 2. SCXRD of Cl/SF₅-alkenes **2e**, **2l**, and **2af** and 2:1 adduct **3a**.

available and difficult to prepare.³⁶ With our present method, triisopropylsilyl-acetylene **1z** delivered **2z** in a modest 13% isolated yield, while we obtained a quantitative yield of **2aa** featuring a triethylsilyl group. The latter was convenient to prepare, was not so volatile, and was easily purified by distillation. Addition of SF₅Cl to a boron-substituted alkyne (BMIDA, **1ab**) is described for the first time with up to 54% conversion and 36% isolated yield. The reaction also performed well with functionalized alkyl alkynes **2ac–ad** and with triple bonds embedded in ynoate (product **2ae**) and ynone (product **2af**) leading to unprecedented motifs in 71% and 70% yields, respectively. With these conjugated substrates, the SF₅ was selectively introduced in the α -position of the carbonyl as demonstrated by SCXRD of **2af** (Figure 2). Overall, the present conditions allowed improvement of the yields of Cl/SF₅-alkenes, reduction of the amounts of 2:1 adducts, and access to completely new structures.

Additional substrates were evaluated, for instance, some featuring an acid or an alcohol function, but reactions were messy and GC-yields did not exceed 30% (see SI for reluctant substrates). Extension of our conditions to alkenes as substrates ended up with no product formation or low reaction yields (see SI for examples and a possible explanation). The complete synthetic sequence toward SF₅-alkynes by HCl elimination with the aid of a base is a straightforward step except for boron-substituted alkene **2ab** (see SI for examples). From a mechanistic viewpoint, the reaction proceeds through free radical chain addition of SF₅Cl to alkynes that is inhibited in the presence of TEMPO (see SI). Since the reaction also takes place in the dark, we assumed that the proclivity of THF to autoxidation³⁷ induces formation of hydroperoxides, as evidenced in our distilled THF by the Creary's method,³⁸ that are responsible for the initiation step (see a proposed mechanism in SI). However, the amount of peroxide is a crucial factor, as too much is impaired with the reaction (Table 1, entry 11).

In conclusion, we have demonstrated that the right choice of solvent and only that solvent is the key for streamlined chloropentafluorosulfanylation of alkynes in the absence of added reagent or light irradiation. We are convinced that this simple and economical approach is of immediate benefit to the chemists eager to access SF₅-containing building blocks for further elaboration of complex fluorinated molecules. We hope that this work will be seen as a major step forward, two decades after Dolbier's seminal paper.

■ ASSOCIATED CONTENT

Data Availability Statement

The data underlying this study are available in the published article and its Supporting Information.

SI Supporting Information

The Supporting Information is available free of charge at <https://pubs.acs.org/doi/10.1021/acs.orglett.3c04043>.

Experimental procedures, characterization data for all new compounds, along with copies of spectra (PDF)

Accession Codes

CCDC 2303459–2303460 and 2307783–2307785 contain the supplementary crystallographic data for this paper. These data can be obtained free of charge via www.ccdc.cam.ac.uk/data_request/cif, or by emailing data_request@ccdc.cam.ac.uk, or by contacting The Cambridge Crystallographic Data Centre, 12 Union Road, Cambridge CB2 1EZ, UK; fax: +44 1223 336033.

■ AUTHOR INFORMATION

Corresponding Authors

Vincent Bizet – Université de Haute-Alsace, Université de Strasbourg, CNRS, LIMA, UMR 7042, 68000 Mulhouse, France; orcid.org/0000-0002-0870-1747;

Email: vbizet@unistra.fr

Dominique Cahard – CNRS, UMR 6014 COBRA, Univ Rouen Normandie, INSA Rouen Normandie, Normandie Univ, F-76000 Rouen, France; orcid.org/0000-0002-8510-1315; Email: dominique.cahard@univ-rouen.fr

Authors

Thi Mo Nguyen – CNRS, UMR 6014 COBRA, Univ Rouen Normandie, INSA Rouen Normandie, Normandie Univ, F-76000 Rouen, France

Lucas Popek – Université de Haute-Alsace, Université de Strasbourg, CNRS, LIMA, UMR 7042, 68000 Mulhouse, France

David Matchavariani – Université de Haute-Alsace, Université de Strasbourg, CNRS, LIMA, UMR 7042, 68000 Mulhouse, France

Nicolas Blanchard – Université de Haute-Alsace, Université de Strasbourg, CNRS, LIMA, UMR 7042, 68000 Mulhouse, France; orcid.org/0000-0002-3097-0548

Complete contact information is available at: <https://pubs.acs.org/doi/10.1021/acs.orglett.3c04043>

Funding

This work was supported by the French National Research Agency – ANR SulFive project (Grant No. ANR-21-CE07–0042) and the CNRS (Centre National de la Recherche Scientifique). This work has also been partially supported by University of Haute-Alsace, University of Strasbourg, University of Rouen Normandie, INSA Rouen Normandie, European Regional Development Fund (ERDF), Labex SynOrg (ANR-11-LABX-0029), Carnot Institute I2C, the graduate school for research XL-Chem (ANR-18-EURE-0020XL CHEM). The authors acknowledge Didier Le Nouën (LIMA UMR7042 CNRS-Unistra-UHA), Corinne Bailly and Nathalie Gruber (Service de radiocristallographie de la Fédération de Chimie Le Bel FR 2010) for their technical and scientific support in NMR and SCXRD analyses.

Notes

The authors declare no competing financial interest.

REFERENCES

- (1) Savoie, P. R.; Welch, J. T. Preparation and utility of organic pentafluorosulfanyl-containing compounds. *Chem. Rev.* **2015**, *115*, 1130–1190.
- (2) Kordnezhadian, R.; Li, B. Y.; Zogu, A.; Demaerel, J.; De Borggraeve, W. M.; Ismalaj, E. Chemistry of Pentafluorosulfanyl Derivatives and Related Analogs: From Synthesis to Applications. *Chem.—Eur. J.* **2022**, *28*, No. e202201491.
- (3) Haufe, G. Synthesis and application of pentafluorosulfanylation reagents and derived aliphatic SF₅-containing building blocks. *Tetrahedron* **2022**, *109*, No. 132656.
- (4) Sheppard, W. A. The Electrical Effect of the Sulfur Pentafluoride Group. *J. Am. Chem. Soc.* **1962**, *84*, 3072–3076.
- (5) Hansch, C.; Leo, A.; Unger, S. H.; Kim, K. H.; Nikaitani, D.; Lien, E. J. Aromatic substituent constants for structure-activity correlations. *J. Med. Chem.* **1973**, *16*, 1207–1216.
- (6) Bergmann, D.; Hinze, J. *Struct. Bonding (Berlin)*; Sen, K. D., Jørgensen, C. K., Eds.; Springer-Verlag: Berlin, 1987; Vol. 66, pp 145–190.
- (7) Nixon, P. G.; Winter, R.; Castner, D. G.; Holcomb, N. R.; Grainger, D. W.; Gard, G. L. Pentafluoro- λ^6 -sulfanyl-Terminated Chlorosilanes: New SF₅-Containing Films and Polysiloxane Materials. *Chem. Mater.* **2000**, *12*, 3108–3112.
- (8) Savoie, P. R.; von Hahmann, C. N.; Penger, A.; Wei, Z.; Welch, J. T. The control of stereochemistry by the pentafluorosulfanyl group. *Org. Biomol. Chem.* **2018**, *16*, 3151–3159.
- (9) Burton, D. J.; Wang, Y.; Bizet, V.; Cahard, D. Pentafluorosulfanyl Chloride. *Encyclopedia of Reagents for Organic Synthesis* **2020**, 1–7.
- (10) Beier, P., Pentafluorosulfanylation of Aromatics and Heteroaromatics. In *Emerging Fluorinated Motifs*; Ma, J.-A., Cahard, D., Eds. Wiley-VCH: Weinheim, Germany, 2020; Vol. 2, pp 551–570.
- (11) Popek, L.; Nguyen, T. M.; Blanchard, N.; Cahard, D.; Bizet, V. Synthesis and further use of SF₅-alkynes as platforms for the design of more complex SF₅-containing products. *Tetrahedron* **2022**, *117*–*118*, No. 132814.
- (12) Shou, J.-Y.; Xu, X.-H.; Qing, F.-L. The radical reaction of ethynylbenziodoxolone (EBX) reagents with pentafluorosulfanyl chloride: New approach to SF₅-substituted alkynes. *J. Fluorine Chem.* **2022**, *261*–*262*, No. 110018.
- (13) Ait-Mohand, S.; Dolbier, W. R. New and Convenient Method for Incorporation of Pentafluorosulfanyl (SF₅) Substituents into Aliphatic Organic Compounds. *Org. Lett.* **2002**, *4*, 3013–3015.
- (14) Cloutier, M.; Roudias, M.; Paquin, J.-F. Regioselective Gold-Catalyzed Hydration of CF₃- and SF₅-alkynes. *Org. Lett.* **2019**, *21*, 3866–3870.
- (15) Gauthier, R.; Mamone, M.; Paquin, J.-F. Gold-Catalyzed Hydrofluorination of Internal Alkynes Using Aqueous HF. *Org. Lett.* **2019**, *21*, 9024–9027.
- (16) Popek, L.; Cabrera-Trujillo, J. J.; Debrauwer, V.; Blanchard, N.; Miqueu, K.; Bizet, V. Regio- and Stereoselective Hydroelementation of SF₅-Alkynes and Further Functionalizations. *Angew. Chem., Int. Ed.* **2023**, *62*, No. e202300685.
- (17) Debrauwer, V.; Leito, I.; Lökov, M.; Tshepelevitsh, S.; Parmentier, M.; Blanchard, N.; Bizet, V. Synthesis and Physicochemical Properties of 2-SF₅-(Aza)Indoles, a New Family of SF₅ Heterocycles. *ACS Org. Inorg. Au* **2021**, *1*, 43–50.
- (18) Dolbier, W. R., Jr; Mitani, A.; Xu, W.; Ghiviriga, I. Synthesis of novel pentafluorosulfanylfurans. Two Retro-Diels-Alder approaches. *Org. Lett.* **2006**, *8*, 5573–5575.
- (19) Dolbier, W. R., Jr; Zheng, Z. Preparation of Pentafluorosulfanyl (SF₅) Pyrrole Carboxylic Acid Esters. *J. Org. Chem.* **2009**, *74*, 5626–5628.
- (20) Dolbier, W. R., Jr; Zheng, Z.-Y. Use of 1,3-dipolar reactions for the preparation of SF₅-substituted five-membered ring heterocycles. Pyrroles and thiophenes. *J. Fluorine Chem.* **2011**, *132*, 389–393.
- (21) Lopez, S. E.; Mitani, A.; Pena, P.; Ghiviriga, I.; Dolbier, W. R. Use of 1-pentafluorosulfanyl-phenylacetylenes for the preparation of SF₅-substituted five-membered ring heterocycles through 1,3-dipolar cycloadditions. Isoxazoles and isoxazolines. *J. Fluorine Chem.* **2015**, *176*, 121–126.
- (22) Emamian, S. Understanding the regioselectivity and molecular mechanism in the synthesis of isoxazoles containing pentafluorosulfanyl substitution via a [3 + 2] cycloaddition reaction: A DFT study. *J. Fluorine Chem.* **2015**, *178*, 165–172.
- (23) Gilbert, A.; Langowski, P.; Delgado, M.; Chabaud, L.; Pucheault, M.; Paquin, J.-F. Amine–borane complex-initiated SF₅Cl radical addition on alkenes and alkynes. *Beilstein J. Org. Chem.* **2020**, *16*, 3069–3077.
- (24) Correction ref 23: *Beilstein J. Org. Chem.* **2021**, *17*, 1725–1726.
- (25) Gilbert, A.; Birepinte, M.; Paquin, J.-F. Electron donor-acceptor (EDA)-complex enabled SF₅Cl addition on alkenes and alkynes. *J. Fluorine Chem.* **2021**, *243*, No. 109734.
- (26) Ouellet-Du Berger, M.-R.; Boucher, M.; Paquin, J.-F. Light-activation of SF₅Cl for the atom transfer radical addition onto alkenes and alkynes. *J. Fluorine Chem.* **2023**, *268*, No. 110131.
- (27) Taponard, A.; Jarrosson, T.; Khrouz, L.; Medebielle, M.; Broggi, J.; Tlili, A. Metal-Free SF₆ Activation: A New SF₅-Based Reagent Enables Deoxyfluorination and Pentafluorosulfanylation Reactions. *Angew. Chem., Int. Ed.* **2022**, *61*, No. e202204623.
- (28) Ponomarenko, M. V.; Kalinovich, N.; Serguchev, Y. A.; Bremer, M.; Röschenthaler, G.-V. Synthesis of pentafluoro- λ^6 -sulfanyl substituted acetylenes for novel liquid crystals. *J. Fluorine Chem.* **2012**, *135*, 68–74.
- (29) Birepinte, M.; Champagne, P. A.; Paquin, J. F. Photoinitiated anti-Hydro-pentafluorosulfanylation of Terminal Alkynes. *Angew. Chem., Int. Ed.* **2022**, *61*, No. e202112575.
- (30) Niina, K.; Tanagawa, K.; Sumii, Y.; Saito, N.; Shibata, N. Pyridine tetrafluoro- λ^6 -sulfanyl chlorides: spontaneous addition to alkynes and alkenes in the presence or absence of photo-irradiation. *Org. Chem. Front.* **2020**, *7*, 1276–1282.
- (31) Kraemer, Y.; Ghiazza, C.; Ragan, A. N.; Ni, S.; Lutz, S.; Neumann, E. K.; Fetting, J. C.; Nothling, N.; Goddard, R.; Cornella, J.; Pitts, C. R. Strain-Release Pentafluorosulfanylation and Tetrafluoro(aryl)sulfanylation of [1.1.1]Propellane: Reactivity and Structural Insight. *Angew. Chem., Int. Ed.* **2022**, *61*, No. e202211892.
- (32) Shou, J. Y.; Xu, X. H.; Qing, F. L. Chemoselective Hydro(Chloro)pentafluorosulfanylation of Diazo Compounds with Pentafluorosulfanyl Chloride. *Angew. Chem., Int. Ed.* **2021**, *60*, 15271–15275.
- (33) Pitts, C. R.; Santschi, N.; Togni, A. Method for preparing polyfluorinated compound. WO2019229103A1, 2019.
- (34) Gilbert, A.; Paquin, J.-F. Evaluation of the compatibility of pentafluorosulfanyl chloride with various solvents and additives. *J. Fluorine Chem.* **2019**, *221*, 70–74.
- (35) Kordnezhadian, R.; De Bels, T.; Su, K.; Van Meervelt, L.; Ismalaj, E.; Demaerel, J.; De Borggraeve, W. M. An Extrusion Strategy for On-demand SF₅Cl Gas Generation from a Commercial Disulfide. *Org. Lett.* **2023**, *25*, 8947–8951.
- (36) Lal, G. S.; Minnich, K. E., *US Patent* US6479645 B1, 2002.
- (37) Matsubara, H.; Suzuki, S.; Hirano, S. An ab initio and DFT study of the autoxidation of THF and THP. *Org. Biomol. Chem.* **2015**, *13*, 4686–4692.
- (38) Creary, X. No-Deuterium Proton (No-D) NMR as a Convenient Method for Analysis of Organic Solvents. *J. Org. Chem.* **2023**, *88*, 11545–11551.