

Practical synthesis of polysubstituted naphthalene derivatives *via* HNTf₂-catalyzed benzannulation reaction

Sudipta Ponra, Maxime R. Vitale,* Véronique Michelet,*
and Virginie Ratovelomanana-Vidal*

PSL Research University, Chimie ParisTech - CNRS, Institut de Recherche de Chimie Paris,
75005, Paris, France

E-mail: maxime.vitale@chimie-paristech.fr, veronique.michelet@chimie-paristech.fr,
virginie.vidal@chimie-paristech.fr

Dedicated to Professor J. S. Yadav on the occasion of his 65th birthday

DOI: <http://dx.doi.org/10.3998/ark.5550190.p009.219>

Abstract

The synthesis of polysubstituted naphthalenes using the triflimide-catalyzed benzannulation of arylacetaldehydes with alkynes at room temperature is described. This method demonstrates a high functional group tolerance and, in the case of halogen substituted naphthalenes, opens the route for further functionalization by palladium-catalyzed cross-coupling reactions. With an analogous organocatalytic strategy, arylepoxides or 2-arylacetal derivatives are also suitable partners in the related benzannulation reactions with aryl-alkynes.

Keywords: Organocatalysis, naphthalene, benzannulation, triflimide

Introduction

Substituted naphthalene derivatives are important pharmacophores in many biologically active compounds which possess anti-inflammatory, antibacterial, antimicrobial and anticancer activities.¹⁻⁹ Additionally, this particular aromatic structure can be found in numerous optical and electronic materials¹⁰⁻¹² and constitutes the backbone of many chiral ligands.¹³ Nafacillin,¹⁴ suramin,^{15,16} which play a vital role in the control of microbial infection, are typical examples of drugs that present a naphthalene moiety (Figure 1). Therefore, the development of new and efficient methods for the synthesis of naphthalene skeletons has been a subject of great interest in recent years.¹⁷⁻⁴² Most of the traditional approaches toward naphthalene derivatives involve the stepwise introduction of a substituent through electrophilic substitution or coupling reactions.^{17,18} Construction of the second aromatic ring of the naphthalene core *via* a formal [4+2] process

under various catalytic conditions is definitely one of the most direct and efficient methods.¹⁹⁻⁴² In particular, TiCl_4 or FeCl_3 in stoichiometric quantities as well as catalytic GaCl_3 or $\text{AuCl}_3/\text{AgSbF}_6$ systems were found to promote the benzannulation reaction of arylacetaldehydes with alkynes.²⁹⁻³² Boron trifluoride etherate complex was also described as an appropriate catalyst for this transformation, in the specific case of terminal arylacetylenes.³⁴

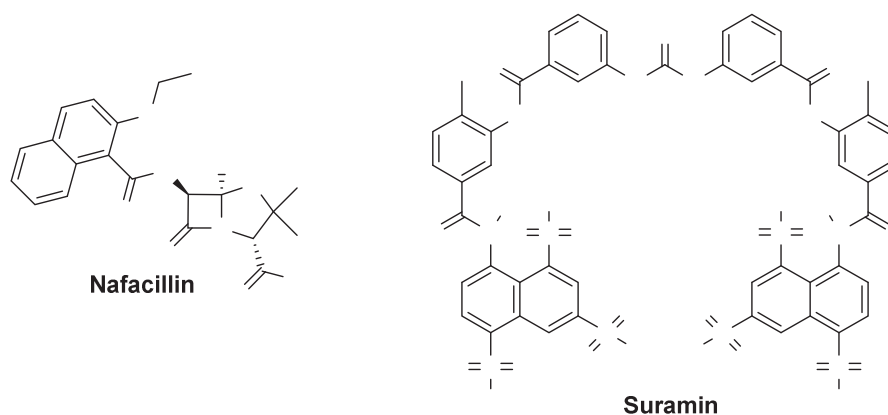


Figure 1. Two representative naphthalene containing drugs.

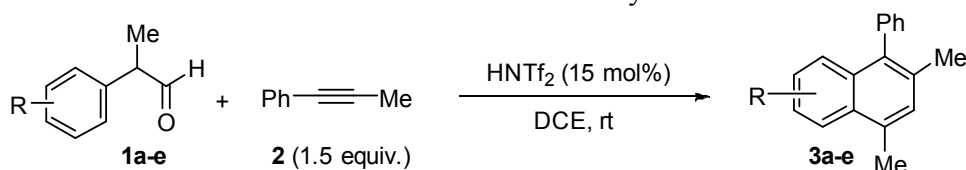
In this field, we recently discovered that triflimide (HNTf_2)⁴³⁻⁴⁷ is also an efficient catalyst for the benzannulation of arylacetaldehyde derivatives with alkynes.⁴⁸ This organocatalytic metal-free reaction proceeds under mild reaction conditions at room temperature and leads to a variety of substituted naphthalene compounds with, in the vast majority of cases, perfect regioselectivity. We report herein a more detailed study of this benzannulation reaction for the synthesis of a larger number of naphthalene derivatives displaying various useful functionalities. Additionally, this strategy could be applied to the analogous benzannulation of aryloxydes and arylacetals. The further functionalization of these aromatic scaffolds by palladium-catalyzed cross-coupling reactions is also presented.

Results and Discussion

Initial optimization studies allowed us to determine that the benzannulation reactions of phenylacetaldehydes with 1.5 equivalents of alkynes in DCE at room temperature were efficiently catalyzed by 15 mol % HNTf_2 .⁴⁸ Accordingly, these reaction conditions were applied for the synthesis of a wide variety of naphthalenes in order to evaluate the scope and limitations of this novel protocol. At first, various 2-methyl-phenylacetaldehydes **1a-h** were reacted with 1-phenyl-1-propyne **2** to assess the influence of the aromatic substitution of the aldehyde partner over benzannulation efficiency (Table 1 & Scheme 1). This reaction gave almost similar yields (62-70%) when the aryl group was not substituted **1a** ($\text{R} = \text{H}$) or when it bears an electron-

donating substituent **1b-c** (R = Me, OMe) on 4-position of the aromatic ring (Table 1, entries 1-3). In the specific case of 2-(3,5-dimethoxyphenyl)-propionaldehyde **1d**, a degradation of the reaction mixture was observed so that naphthalene **3d** was isolated in a low yield of 8% as a probable consequence of the increased electron-richness of this substrate (Table 1, entry 4). With **1e**, which presents a nitro group at the *para* position of its aromatic moiety, a slightly diminished yield of 46% was obtained which is in good correlation with the deactivating effect of such group in electrophilic aromatic substitutions (Table 1, entry 5).

Table 1. Influence of electronic factors induced by the substitution of the aldehyde aromatic ring

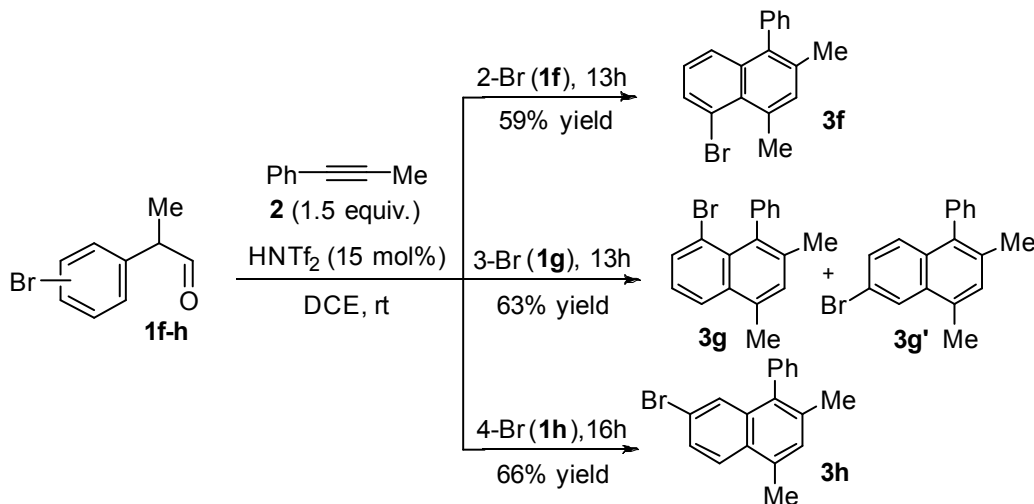


Entry	Aldehyde	Product	t (h)	Yield (%) ^a
1			13	70
2			14	70
3			20	62
4			13	8
5			20	46

^aIsolated yield.

We were pleased to observe a relatively small influence of the steric effect induced by the substitution of the aromatic ring. Indeed, the *ortho*-, *meta*- or *para*-bromo phenylacetaldehydes **1f-h** afforded the corresponding naphthalenes **3f-h** in almost identical yields (Scheme 1). Whereas **3f** and **3h** were obtained as single regioisomers in 59% and 66% yield respectively, the

benzannulation of 3-bromo phenylacetaldehyde **1g** led to a 1:1 regioisomeric mixture of naphthalenes **3g** and **3g'** in 63% yield.



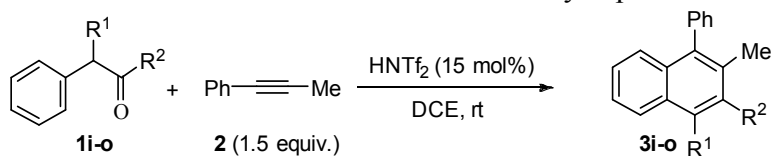
Scheme 1. Influence of steric factors induced by the substitution of the aldehyde aromatic ring.

We next moved our attention toward the influence of the α -substitution of the aldehyde partner and the evaluation of the reactivity of few ketones (Table 2). Switching from a methyl group to phenyl, ethyl or cyclohexyl α -substituents had a limited impact over the efficiency of the benzannulation process since compounds **3i-k** were obtained in good 69-78% yields (Table 2, entries 1-3). Unsubstituted phenylacetaldehyde **1l** reacted with somewhat lower efficiency to give the naphthalene adduct **3l** in 49% yield (Table 2, entry 4), which tended to support that α -substitution of the aldehyde group may prove itself beneficial to achieve better yields. On the other hand, ketones **1m-o** were tested. However, only 2-phenylcyclohexanone **1o** resulted in the formation of the desired benzannulation product **3o** in low 25% yield (Table 2, entry 7), thus suggesting that this HNTf₂-catalyzed benzannulation process is poorly efficient with ketones.

To further determine the scope of our organocatalyzed benzannulation reaction, we investigated the reactivity of different alkyne partners with 2-phenylpropionaldehyde **1a** (Table 3 & Table 4). We began our studies by performing some reactions with phenylacetylenes possessing various alkyl groups and we were pleased to observe that the reaction of alkynes **4a**, **4b** and **4c** ($R^1 = \text{Ph}$ and $R^2 = \text{Et}$, $n\text{Pr}$, $n\text{Bu}$) afforded the corresponding naphthalenes **5a-c** in 66-70% yields (Table 3, entries 1-3). We then decided to examine the influence of the aromatic substitution of the aryl-alkyne partner. Accordingly, the 4-chloro-, 4-bromo- and 4-methyl- substituted alkynes **4d-f** were submitted to reaction and gave access to desired products **5d-f** with comparable satisfactory results (Table 3, entries 4-6). In the case of alkynes **4g** and **4h**, which are bearing electron-withdrawing/donating 4-trifluoromethyl and 4-methoxy groups, a reduced reaction efficiency could however be noticed (Table 3, entries 7-8). Notably, the reaction of the electron rich alkyne **4h** was faster and the aldehyde **1a** was converted to **5h** in 44% yield along with some

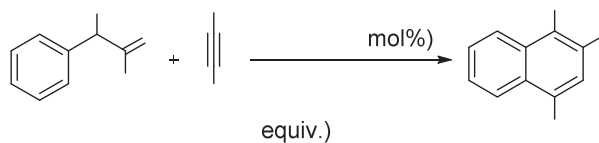
unidentified by-products (Table 3, entry 8). Diaryl-substituted alkynes were also prone to react under these catalytic conditions. The reaction of symmetrical alkynes **4i** and **4j** led to naphthalene compounds **5i** and **5j** in 46 and 49% yields respectively (Table 3, entries 9-10). On the other hand the unsymmetrical alkyne **4k** generated a 1:1 mixture of the two regioisomers **5k/5k'** in 39% yield (Table 3, entry 11). Trimethylsilyl-substituted phenylacetylene **4l** gave only degradation of starting materials (Table 3, entry 12).

Table 2. Influence of α -substitution of the aldehyde partner and evaluation of ketones



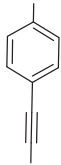
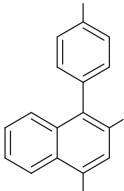
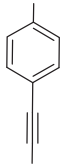
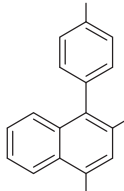

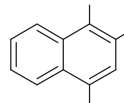

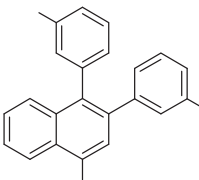
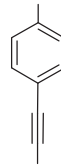
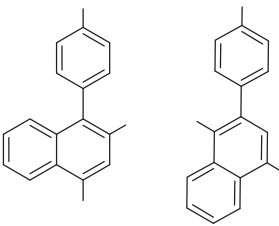

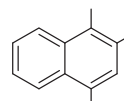
Entry	Aldehyde	Product	t (h)	Yield (%) ^a
1			16	69
2			13	78
3			14	69
4			13	49
5			60	-
6			13	-
7			60	25

^aIsolated yield.

Table 3. Benzannulation scope for di-substituted aryl-alkynes

Entry	Alkyne	Product	t (h)	Yield (%) ^a
1			15	69
2			15	66
3			15	70
4			15	69
5			15	68
6			15	62

Table 3 (continued)

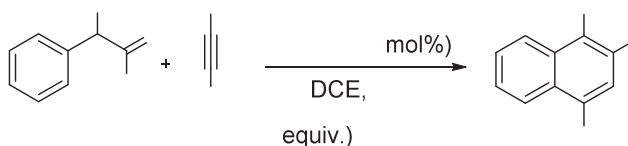
Entry	Alkyne	Product	t (h)	Yield (%) ^a
7			15	36
8			1	44
9 ^b			48	46
10 ^b			48	49
11 ^b			48	39 ^c
12			13	-

^a Isolated yield. ^b 1.5:1 aldehyde:alkyne ratio. ^c 1:1 mixture of regioisomers.

Afterwards, we turned our attention toward a wider range of aromatic alkynes and aliphatic alkynes (Table 4). In the case of terminal aromatic alkynes, the benzannulation process proved to be more challenging so that, for this particular class, the alkyne-loading was increased to 3 equivalents. Under the corresponding reaction conditions, phenylacetylene **4m** yielded the

desired 1,4-disubstituted naphthalene **5m** in 41% yield (Table 4, entry 1). The substitution of the phenylacetylene aromatic core with 4-fluoro or 4-bromo moiety did not affect the outcome of the benzannulation and produced the expected compounds **5n** and **5o** with similar results (Table 4, entries 2-3). Conversely, electron-rich 4-methyl and 4-methoxy terminal alkynes **4p** and **4q** provided complex mixtures of products from which only the naphthalene **5p** could be isolated in low 11% yield (Table 4, entries 4-5). Interestingly, the keto- and ester-substituted naphthalenes **5r** and **5s** were obtained in 46% and 35% yields respectively using alkynes **4r** and **4s** bearing electron-withdrawing groups (Table 4, entries 6-7). This method could also be extended to halogen-substituted phenylacetylenes. Indeed, the corresponding alkynes **4t-v** were converted to 2-chloro-, 2-bromo- substituted naphthalene compounds **5t** and **5u** with useful 43% and 42% yields respectively (Table 4, entries 8-9), and to the 2-iodonaphthalene **5v** with 34% yield (Table 4, entry 10). When opposed to diphenylacetaldehyde **1i**, bromo-alkyne **4u** led to naphthalene **6** in 37% yield (Table 4, entry 11). Then, terminal and internal aliphatic alkynes **4w** and **4x** were tested but reacted much more sluggishly than their aryl-alkyne counterparts resulting in the limited formation of the naphthalenes **5w** and **5x** in 21% and 11% respectively (Table 4, entries 12-13).

Table 4. Benzannulation scope for terminal aryl-alkynes, carbo-alkynes, halogeno alkynes and aliphatic alkynes



Entry	Alkyne	Product	t (h)	Yield (%) ^a
1 ^b			21	41
2 ^b			21	42

Table 4 (continued)

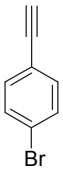
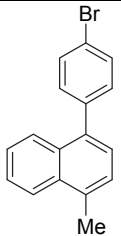
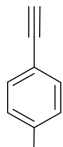
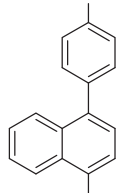
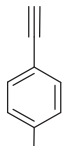
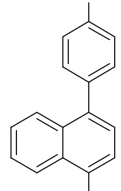

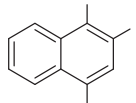

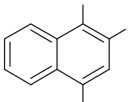

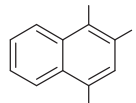

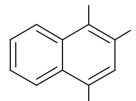

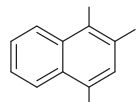

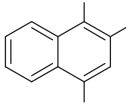

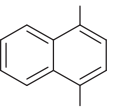

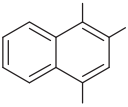
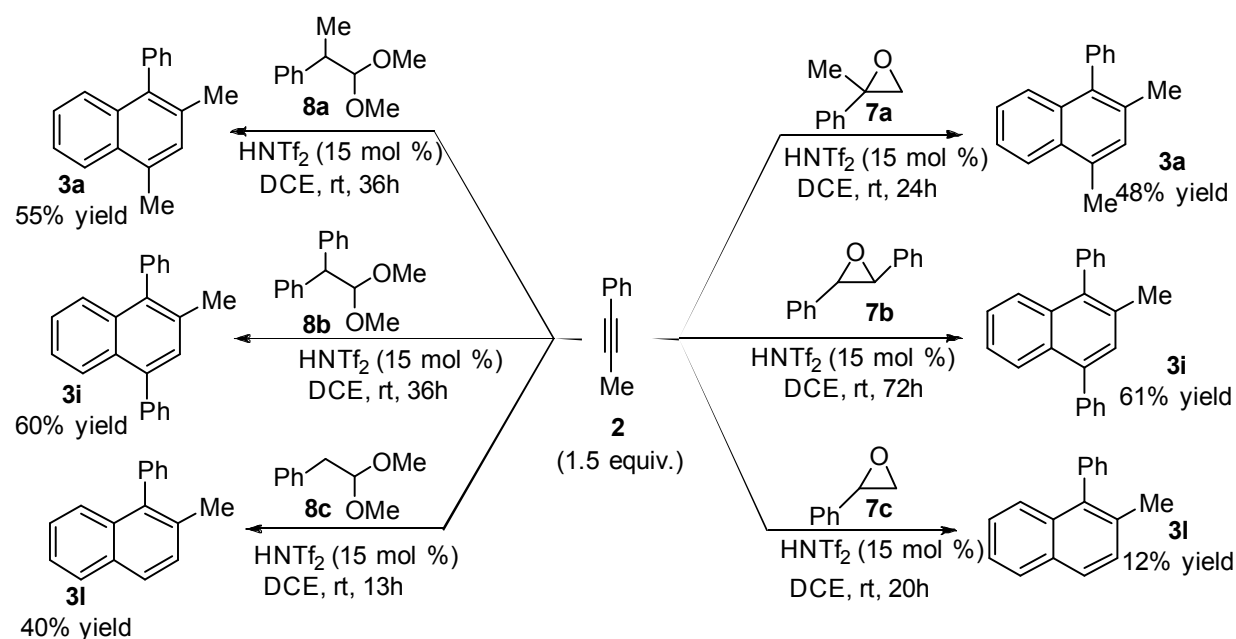
Entry	Alkyne	Product	t (h)	Yield (%) ^a
3 ^b	 4o	 5o	15	41
4 ^b			21	11
5 ^b			1	-
6			13	46
7			14	35
8			20	43
9			20	42
10			20	34

Table 4 (continued)

Entry	Alkyne	Product	t (h)	Yield (%) ^a
11			20	37
12			36	21
13			20	11

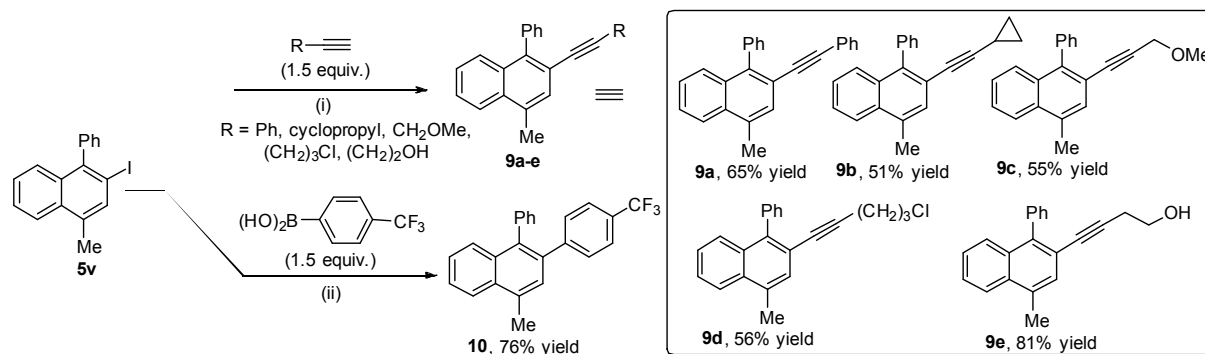
^a Isolated yield. ^b 3 equiv. of alkyne.

Considering literature precedents,³⁵⁻³⁷ we envisioned that this Brønsted acid catalyzed benzannulation reaction could be extended to the use of epoxides or acetals. Pleasingly, epoxides **7a-c** reacted with 1-phenyl-1-propyne **2** in the presence of 15 mol% HNTf₂ in DCE at room temperature to provide the corresponding naphthalenes **3a**, **3i** and **3l** in 12-61% yields. On the other hand, acetals **8a-c**, under the same reaction conditions, furnished the desired products **3a**, **3i** and **3l** with slightly better efficiency (Scheme 2).



Scheme 2. Scope of epoxides and acetals for the synthesis of naphthalene derivatives.

To further demonstrate the usefulness of this method of access to polyfunctionalized naphthalene derivatives, we then studied the reactivity of the iodonaphthalene **5v** in palladium catalyzed Sonogashira and Suzuki-Miyaura cross-couplings (Scheme 3).^{49,50} Satisfyingly, compound **5v** reacted well with various aryl and aliphatic alkynes to give the corresponding alkynyl-naphthalenes **9a-e** in 51-81% yields and when engaged with 4-(trifluoromethyl)phenylboronic acid allowed to obtain the polyaromatic **10** in good 76% yield.



Reagents and conditions: (i) Pd(PPh₃)₂Cl₂ (5 mol%), Cul (5 mol%), DMF/NEt₃ (7:3), rt, 13h. (ii) Pd(OAc)₂ (5 mol%), dppf (5 mol%), K₂CO₃ (3 equiv.), DMF, 80 °C, 13h.

Scheme 3. Pd-catalyzed cross-coupling reactions.

Conclusions

We have developed a practical and easy protocol to access naphthalene derivatives from readily available starting materials through the triflimide (HNTf₂) organocatalyzed benzannulation reactions of arylacetaldehyde, arylepoxide and arylacetal derivatives with alkynes. Attractive features of these organocatalytic transformations involve the mild reaction conditions and the wide substrate scope allowing the straightforward access to highly substituted naphthalenes with, in most cases, perfect regioselectivity. We could also demonstrate that this method may lead to valuable platforms such as iodonaphthalenes which can be further functionalized *via* palladium-catalyzed cross-coupling reactions.

Acknowledgements

This work was supported by the Ministère de l'Éducation Supérieure et de la Recherche (MESR) and the Centre National de la Recherche Scientifique (CNRS). S. Ponra is grateful to the "Fondation Pierre-Gilles de Gennes pour la Recherche" for a postdoctoral grant.

Experimental Section

General. All reactions were performed under argon atmosphere. 1,2-Dichloroethane was distilled from CaH₂. All products were purified by flash chromatography using silica gel (230-400 mesh). ¹H-NMR and ¹³C-NMR were recorded in CDCl₃ with chemical shifts reported relative to residual CHCl₃ peak for ¹H NMR (7.26 ppm) or the central peak of CDCl₃ for ¹³C NMR (77.16 ppm). HRMS data for new compounds were obtained using an atmospheric pressure photo ionization source (AAPI) coupled to a LTQ-Orbitrap high resolution detector. Unless otherwise noted, all the reagents were ordered and used without further purification. Starting materials were prepared according to literature (see supplementary information for more details). Naphthalene derivatives **3a-c**, **3e-f**, **3h-l** and **5a-f**, **5i-j**, **5m-o**, **5r**, **5t-u** have already been described in our previous report.⁴⁸

Procedure for the benzannulation reaction: In a screw cap vial under argon atmosphere were sequentially added the arylaldehyde **1** or arylepoxides **7** or 2-arylacetal **8** (1 mmol, 1 equiv.), the alkyne **2** or **4** (1.5 mmol, 1.5 equiv.), 1,2-dichloroethane (1 mL) and HNTf₂ (42 mg, 0.15 mmol, 0.15 equiv.). The resulting mixture was stirred at room temperature until TLC analysis showed completion of the reaction. The reaction mixture was then diluted with dichloromethane (5 mL) and water (15 mL) and transferred to a separating funnel. The aqueous phase was extracted with dichloromethane (3 x 15 mL) and the combined organic extracts washed by water (2 x 40 mL) and brine (40 mL) before being dried over MgSO₄. After filtration and evaporation of the solvents under reduced pressure, the crude material was purified by flash column chromatography on silica gel to afford the desired naphthalene **3** or **5**. In specific cases, this first purification step was followed by a bulb to bulb distillation under reduced pressure in order to remove residual alkyne.

6,8-Dimethoxy-2,4-dimethyl-1-phenylnaphthalene (3d). Starting from 2-(3,5-dimethoxyphenyl)-propanal **1d** (194 mg, 1.0 mmol) and 1-phenyl-1-propyne **2** (174 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (98:2) as eluent. The desired naphthalene **3d** was obtained as a pale yellow oil (22 mg, 8% yield). ¹H NMR (300 MHz, CDCl₃) δ 7.38 – 7.33 (m, 2H), 7.31 – 7.27 (m, 1H), 7.24 (s, 1H), 7.17 – 7.12 (m, 2H), 6.85 (d, *J* 2.4 Hz, 1H), 6.42 (d, *J* 2.4 Hz, 1H), 3.94 (s, 3H), 3.32 (s, 3H), 2.64 (d, *J* 0.7 Hz, 3H), 2.05 (s, 3H); ¹³C NMR (75 MHz, CDCl₃) δ 158.4, 157.3, 144.9, 135.3, 133.7, 131.9, 131.3, 130.8, 128.7, 127.2, 125.3, 120.6, 99.2, 95.5, 55.6, 55.4, 21.0, 20.4; HRMS (APPI) *m/z*: [M]⁺• calcd for C₂₀H₂₀O₂ 292.1458, found: 292.1461.

6-Bromo-2,4-dimethyl-1-phenylnaphthalene (3g) and 8-bromo-2,4-dimethyl-1-phenylnaphthalene (3g'). Starting from 2-(3-bromophenyl)-propanal **1g** (209 mg, 0.98 mmol) and 1-phenyl-1-propyne **2** (171 mg, 1.47 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane as eluent. The desired naphthalenes **3g** and **3g'** (1:1 mixture) were obtained as a colorless oil (192 mg, 63%

yield). ^1H NMR (300 MHz, CDCl_3) δ 8.21 (d, J 1.9 Hz, 1H), 8.08 (dd, J 8.4, 1.2 Hz, 1H), 7.83 (dd, J 7.4, 1.2 Hz, 1H), 7.61 – 7.43 (m, 7H), 7.36 (d, J 7.5 Hz, 3H), 7.33 – 7.24 (m, 5H), 2.77 (s, 3H), 2.74 (s, 3H), 2.25 (s, 3H), 2.19 (s, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 141.6, 139.5, 136.7, 136.4, 136.2, 134.2, 134.0, 133.7, 133.4, 132.6, 132.5, 131.7, 131.3, 130.6, 130.5, 130.4, 130.3, 128.8 (2C), 128.6, 127.9, 127.3, 127.0, 126.4, 125.0, 124.6, 120.5, 119.2, 22.1, 20.8, 20.4, 19.4; HRMS (APPI) m/z : $[\text{M}]^+$ calcd for $\text{C}_{18}\text{H}_{15}\text{Br}$ 310.0352, found: 310.0357.

10-Methyl-9-phenyl-1,2,3,4-tetrahydrophenanthrene (3o). Starting from 2-phenylcyclohexanone **1o** (174 mg, 1.0 mmol) and 1-phenyl-1-propyne **2** (174 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane as eluent. The desired naphthalene **3o** was obtained as a white low melting solid (68 mg, 25% yield). ^1H NMR (300 MHz, CDCl_3) δ 8.04 (d, J 8.5 Hz, 1H), 7.55 – 7.40 (m, 4H), 7.37 – 7.24 (m, 4H), 3.23 (t, J 5.2 Hz, 2H), 2.85 (t, J 5.2 Hz, 2H), 2.12 (s, 3H), 2.05 – 1.90 (m, 4H); ^{13}C NMR (75 MHz, CDCl_3) δ 141.0, 136.6, 134.1, 133.0, 131.4, 131.3, 131.0, 130.6, 128.4, 127.1, 126.9, 124.9, 124.8, 122.7, 28.4, 26.7, 23.4, 23.0, 17.6. These analytical data are in accordance with literature.³²

2,4-Dimethyl-1-(4-(trifluoromethyl)-phenyl)-naphthalene (5g). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and 1-(prop-1-ynyl)-4-(trifluoromethyl)benzene **4g** (276 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent followed by bulb to bulb distillation of residual alkyne at 100°C/0.3 Torr during 1 hour. The desired naphthalene **5g** was obtained as a white solid (108 mg, 36% yield). mp 121-123 °C; ^1H NMR (300 MHz, CDCl_3) δ 8.07 (d, J 8.4, 1H), 7.85 – 7.78 (m, 2H), 7.57 – 7.49 (m, 1H), 7.48 – 7.38 (m, 4H), 7.34 (s, 1H), 2.78 (d, J 1.0 Hz, 3H), 2.24 (s, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 144.1, 135.1, 134.2, 132.8, 132.8, 131.2, 131.0, 129.5, 129.4 (q, $^3J_{\text{C-F}}$ 32.2 Hz), 126.4, 126.0, 125.5 (q, $^2J_{\text{C-F}}$ 3.8 Hz), 125.0, 124.5 (q, $^1J_{\text{C-F}}$ 271.5 Hz), 124.2, 20.7, 19.5; ^{19}F NMR (282 MHz, CDCl_3) δ -63.2 (s); HRMS (APPI) m/z : $[\text{M}]^+$ calcd for $\text{C}_{19}\text{H}_{15}\text{F}_3$ 300.1120, found: 300.1124.

1-(4-Methoxyphenyl)-2,4-dimethylnaphthalene (5h). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and 1-methoxy-4-(prop-1-ynyl)benzene **4h** (219 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent followed by bulb to bulb distillation of residual alkyne at 100°C/0.3 Torr during 1 hour. The desired naphthalene **5h** was obtained as a white solid (114 mg, 44% yield). mp 103-105 °C; ^1H NMR (300 MHz, CDCl_3) δ 7.99 (dt, J 8.1, 1.0 Hz, 1H), 7.50 – 7.40 (m, 2H), 7.38 – 7.30 (m, 1H), 7.28 (bs, 1H), 7.22 – 7.14 (m, 2H), 7.07 – 7.00 (m, 2H), 3.91 (s, 3H), 2.72 (d, J 1.0 Hz, 3H), 2.22 (s, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 158.7, 136.3, 133.6, 133.3, 133.2, 132.3, 131.5, 131.2, 129.6, 126.9, 125.6, 124.7, 124.0, 113.9, 55.4, 20.9, 19.5; HRMS (APPI) m/z : $[\text{M}]^+$ calcd for $\text{C}_{19}\text{H}_{18}\text{O}$ 262.1352, found: 262.1354.

1-(4-Chlorophenyl)-4-methyl-2-phenylnaphthalene (5k) and 2-(4-chlorophenyl)-4-methyl-1-phenylnaphthalene (5k'). Starting from 2-phenylpropionaldehyde **1a** (201 mg, 1.5 mmol) and 1-chloro-4-(phenylethynyl)benzene **4k** (212 mg, 1.0 mmol) and following the general procedure,

the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent. The desired naphthalenes **5k** and **5k'** (1:1 mixture) were obtained as a white low melting solid (128 mg, 39% yield). ^1H NMR (300 MHz, CDCl_3) δ 8.14 – 8.03 (m, 2H), 7.71 – 7.60 (m, 2H), 7.59 – 7.51 (m, 2H), 7.47 – 7.37 (m, 4H), 7.33 – 7.24 (m, 6H), 7.22 – 7.04 (m, 12H), 2.79 (s, 6H); ^{13}C NMR (75 MHz, CDCl_3) δ 141.9, 140.7, 139.0, 138.3, 137.9, 136.8, 136.2, 134.7, 134.3, 134.1, 133.1, 132.9, 132.7, 132.6, 132.3, 132.1, 132.0, 131.7, 131.5, 130.2, 129.2, 128.8, 128.6, 128.2, 128.1, 127.9, 127.6, 127.2, 126.9, 126.5, 126.2 (2C), 125.9, 125.8, 124.3, 124.2, 19.7 (2C); HRMS (APPI) m/z : $[\text{M}]^{+\bullet}$ calcd for $\text{C}_{23}\text{H}_{17}\text{Cl}$ 328.1013, found: 328.1013.

1-Methyl-4-(*p*-tolyl)-naphthalene (5p). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and *p*-tolylacetylene **4p** (348 mg, 3.0 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent. The desired naphthalene **5p** was obtained as a white solid (25 mg, 11% yield). mp 95–97 °C; ^1H NMR (300 MHz, CDCl_3) δ 8.11 (dd, J 8.4, 0.7 Hz, 1H), 8.01 (dd, J 8.4, 0.7 Hz, 1H), 7.59 (ddd, J 8.4, 6.8, 1.4 Hz, 1H), 7.54 – 7.31 (m, 7H), 2.80 (d, J 0.7 Hz, 3H), 2.51 (s, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 138.8, 138.2, 136.9, 133.7, 132.9, 131.9, 130.2, 129.1, 126.9, 126.7, 126.3, 125.7 (2C), 124.5, 21.4, 19.7; MS (EI) for $\text{C}_{18}\text{H}_{16}$ m/z 232 (100) $[\text{M}]^{+\bullet}$, 217 (22) $[\text{M-Me}]^{+\bullet}$.

2-Carbomethoxy-4-methyl-1-phenylnaphthalene (5s). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and methyl 3-phenylpropionate **4s** (240 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (98:2) as eluent followed by bulb to bulb distillation of residual alkyne at 115°C/0.3 Torr during 1 hour. The desired naphthalene **5s** was obtained as a pale yellow low melting solid (96 mg, 35% yield). ^1H NMR (300 MHz, CDCl_3) δ 8.10 – 8.03 (m, 1H), 7.79 (d, J 0.9 Hz, 1H), 7.65 – 7.56 (m, 2H), 7.49 – 7.39 (m, 4H), 7.33 – 7.28 (m, 2H), 3.61 (s, 3H), 2.77 (d, J 0.9 Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 169.0, 140.0, 139.3, 134.3, 134.1, 132.8, 129.9, 128.6, 127.9, 127.7, 127.4, 127.3, 126.3, 126.0, 124.2, 52.0, 19.6; HRMS (APPI) m/z : $[\text{M}]^{+\bullet}$ calcd for $\text{C}_{19}\text{H}_{16}\text{O}_2$ 276.1145, found: 276.1150.

2-Iodo-4-methyl-1-phenylnaphthalene (5v). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and 2-iodoethynylbenzene **4v** (342 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent. The desired naphthalene **5v** was obtained as a pale orange solid (118 mg, 34% yield). mp 108–110 °C; ^1H NMR (300 MHz, CDCl_3) δ 8.02 (dt, J 8.5, 1.0 Hz, 1H), 7.91 (d, J 1.0 Hz, 1H), 7.64 – 7.45 (m, 5H), 7.42 – 7.34 (m, 1H), 7.33 – 7.26 (m, 2H), 2.72 (d, J 1.0 Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 143.6, 142.9, 135.9, 135.7, 133.4, 132.2, 130.3, 128.5, 127.9, 127.8, 126.5, 126.1, 124.3, 98.5, 19.1; HRMS (APPI) m/z : $[\text{M}]^{+\bullet}$ calcd for $\text{C}_{17}\text{H}_{13}\text{I}$ 344.0056, found: 344.0058. These analytical data are in accordance with literature.³²

2-Bromo-1,4-diphenylnaphthalene (6). Starting from diphenylacetaldehyde **1i** (196 mg, 1.0 mmol) and 2-bromoethynylbenzene **4u** (272 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum

ether as eluent. The desired naphthalene **6** was obtained as a white solid (132 mg, 37% yield). mp 134-136 °C; ¹H NMR (300 MHz, CDCl₃) δ 8.01 – 7.93 (m, 1H), 7.78 (s, 1H), 7.64 – 7.39 (m, 13H); ¹³C NMR (75 MHz, CDCl₃) δ 141.5, 139.8, 139.4, 139.3, 134.3, 130.8, 130.6, 130.3, 130.1, 128.5 (2C), 127.9 (2C), 127.20, 126.8, 126.4, 126.1, 121.2; HRMS (APPI) *m/z*: [M]⁺ calcd for C₂₂H₁₅Br 358.0357, found: 358.0354.

1-(*n*-Butyl)-4-methylnaphthalene (5w). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1.0 mmol) and 1-hexyne **4w** (123 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent. The desired naphthalene **5w** was obtained as a white low melting solid (42 mg, 21% yield). ¹H NMR (300 MHz, CDCl₃) δ 8.18 – 8.04 (m, 2H), 7.62 – 7.55 (m, 2H), 7.33 – 7.26 (m, 2H), 3.17 – 3.07 (m, 2H), 2.74 (s, 3H), 1.87 – 1.72 (m, 2H), 1.61 – 1.45 (m, 2H), 1.04 (t, *J* 7.3 Hz, 3H); ¹³C NMR (75 MHz, CDCl₃) δ 137.3, 133.1, 132.4, 132.1, 126.4, 125.7, 125.4, 125.3, 125.0, 124.6, 33.3, 32.9, 23.0, 19.6, 14.2. These analytical data are in accordance with literature.²⁹

4-Methyl-1,2-di-(*n*-propyl-naphthalene (5x). Starting from 2-phenylpropionaldehyde **1a** (134 mg, 1 mmol) and 4-octyne **4x** (165 mg, 1.5 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using petroleum ether as eluent. The desired naphthalene **5x** was obtained as colorless oil (25 mg, 11% yield). ¹H NMR (300 MHz, CDCl₃) δ 8.09 – 8.03 (m, 1H), 8.02 – 7.95 (m, 1H), 7.55 – 7.42 (m, 2H), 7.18 (s, 1H), 3.10 – 2.99 (m, 2H), 2.85 – 2.70 (m, 2H), 2.67 (s, 3H), 1.78 – 1.61 (m, 4H), 1.12 (t, *J* 7.3 Hz, 3H), 1.05 (t, *J* 7.3 Hz, 3H); ¹³C NMR (75 MHz, CDCl₃) δ 137.3, 133.7, 132.6, 131.9, 131.8, 129.5, 125.5, 124.7 (2C), 124.5, 35.9, 30.5, 25.0, 24.7, 19.6, 14.9, 14.6. These analytical data are in accordance with literature.²⁹

General procedure for the Sonogashira reaction: Iodonaphthalene derivative **5v** (50 mg, 0.145 mmol), Pd(PPh₃)₂Cl₂ (5 mg, 0.007 mmol, 5 mol%) and CuI (1.4 mg, 0.007 mmol, 5 mol%) were weighed out to a round bottom flask equipped with a magnetic stir bar and fitted with a rubber septa. The flask was purged with argon and DMF (0.5 mL) followed by triethylamine (0.25 mL) and the corresponding alkyne (0.217 mmol, 1.5 eq.) were added to the mixture and the reaction was stirred overnight at room temperature. After completion and hydrolysis (H₂O, 10 mL) the reaction mixture was transferred to a separating funnel. The aqueous phase was extracted with ethyl acetate (3 x 15 mL) and the combined organic extracts washed by water (4 x 40 mL) and brine (40 mL) before being dried over MgSO₄. After filtration and evaporation of the solvents under reduced pressure, the crude material was purified by flash column chromatography on silica gel to afford the desired naphthalene **9**. In specific cases, this first purification step was followed by a bulb to bulb distillation under reduced pressure in order to remove residual alkyne.

4-Methyl-1-phenyl-2-(phenylethynyl)-naphthalene (9a). Starting from iodonaphthalene derivative **5v** (50 mg, 0.145 mmol) and phenylacetylene (22.2 mg, 0.217 mmol) and following

the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (500:1) as eluent. The desired naphthalene **9a** was obtained as a white low melting solid (30 mg, 65% yield). mp 96-98 °C; ¹H NMR (300 MHz, CDCl₃) δ 7.95 (d, *J* 8.0 Hz, 1H), 7.61 (dd, *J* 8.4, 0.8 Hz, 1H), 7.53 – 7.32 (m, 8H), 7.22 – 7.03 (m, 5H), 2.66 (d, *J* 0.8 Hz, 3H); ¹³C NMR (75 MHz, CDCl₃) δ 141.7, 139.3, 133.9, 132.6, 132.5, 131.5, 131.0, 128.9, 128.3, 128.1 (2C), 127.5 (2C), 126.4, 126.2, 124.3, 123.7, 119.9, 93.1, 90.3, 19.5; MS (EI) for C₂₅H₁₈ *m/z* 318.19 (100) [M]⁺.

2-(Cyclopropylethynyl)-4-methyl-1-phenylnaphthalene (9b). Starting from iodonaphthalene derivative **5v** (50 mg, 0.145 mmol) and ethynylcyclopropane (14.3 mg, 0.217 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (500:1) as eluent. The desired naphthalene **9b** was obtained as a yellow colored low melting solid (21 mg, 51% yield). ¹H NMR (300 MHz, CDCl₃) δ 8.02 – 7.90 (m, 1H), 7.55 – 7.65 (m, 1H), 7.54 – 7.32 (m, 8H), 2.70 (d, *J* 0.9 Hz, 3H), 1.31 – 1.21 (m, 1H), 0.82 – 0.61 (m, 2H), 0.50 – 0.35 (m, 2H); ¹³C NMR (75 MHz, CDCl₃) δ 141.1, 139.6, 133.6, 132.5, 132.1, 130.8, 129.3, 127.9, 127.2 (2C), 126.0, 125.9, 124.2, 120.5, 97.5, 75.9, 19.4, 8.8, 0.4; MS (EI) for C₂₂H₁₈ *m/z* 282.17 (92) [M]⁺.

2-(3-Methoxyprop-1-ynyl)-4-methyl-1-phenylnaphthalene (9c). Starting from iodonaphthalene derivative **5v** (50 mg, 0.145 mmol) and 3-methoxyprop-1-yne (15.2 mg, 0.217 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (99.5:0.5) as eluent. The desired naphthalene **9c** was obtained as a yellow colored solid (22.8 mg, 55% yield). mp 64-66 °C; ¹H NMR (300 MHz, CDCl₃) δ 8.02 (d, *J* 8.4 Hz, 1H), 7.61 (d, *J* 8.4 Hz, 1H), 7.57 – 7.35 (m, 8H), 4.14 (s, 2H), 3.14 (s, 3H), 2.72 (s, 3H); ¹³C NMR (75 MHz, CDCl₃) δ 141.6, 139.2, 133.9, 132.6, 132.4, 130.7, 129.2, 128.1, 127.5 (2C), 126.4, 126.2, 124.2, 119.2, 88.4, 86.9, 60.4, 57.2, 19.4; MS (EI) for C₂₁H₁₈O *m/z* 286.15 (85) [M]⁺.

2-(5-Chloropent-1-ynyl)-4-methyl-1-phenylnaphthalene (9d). Starting from iodonaphthalene derivative **5v** (50 mg, 0.145 mmol) and 5-chloropent-1-yne (22.2 mg, 0.217 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (250:1) as eluent followed by bulb to bulb distillation of residual alkyne at 115°C/0.3 Torr during 1 hour. The desired naphthalene **9d** was obtained as a yellow oil (26 mg, 56% yield). ¹H NMR (300 MHz, CDCl₃) δ 8.08 – 7.88 (m, 1H), 7.58 (ddd, *J* 8.4, 1.3, 0.6 Hz, 1H), 7.55 – 7.34 (m, 8H), 3.27 (t, *J* 6.5 Hz, 2H), 2.71 (d, *J* 0.9 Hz, 3H), 2.43 (t, *J* 6.5 Hz, 2H), 1.76 (p, *J* 6.5 Hz, 2H); ¹³C NMR (75 MHz, CDCl₃) δ 141.3, 139.6, 133.8, 132.5, 132.2, 130.7, 129.2, 128.1, 127.4, 127.3, 126.1 (2C), 124.2, 120.1, 91.7, 82.0, 43.5, 31.4, 19.4, 17.0; MS (EI) for C₂₂H₁₉Cl *m/z* 318.10 (100) [M]⁺.

4-(4-Methyl-1-phenylnaphthalen-2-yl)-but-3-yn-1-ol (9e). Starting from iodonaphthalene derivative **5v** (50 mg, 0.145 mmol) and but-3-yn-1-ol (15.2 mg, 0.217 mmol) and following the general procedure, the crude material was purified by flash column chromatography on silica gel using cyclohexane:ethyl acetate (250:1) as eluent. The desired naphthalene **9e** was obtained as yellow oil (33.5 mg, 81% yield). ¹H NMR (300 MHz, CDCl₃) δ 8.01 (dd, *J* 8.3, 0.7 Hz, 1H), 7.66

– 7.34 (m, 9H), 3.48 (q, J 5.9 Hz, 2H), 2.72 (d, J 0.9 Hz, 3H), 2.49 (t, J 5.9 Hz, 2H), 1.24 (t, J 5.9 Hz, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 141.4, 139.8, 133.9, 132.4, 132.3, 130.6, 128.9, 128.2 (2C), 127.6, 127.3, 126.2, 124.3, 119.8, 90.2, 83.2, 61.0, 24.1, 19.4; MS (EI) for $\text{C}_{21}\text{H}_{18}\text{O}$ m/z 286.12 (69) $[\text{M}]^+$.

4-Methyl-1-phenyl-2-(4-(trifluoromethyl)-phenyl)-naphthalene (10). In a screw cap vial under argon atmosphere were sequentially added the idonaphthalene derivative **5v** (42 mg, 0.121 mmol), $\text{Pd}(\text{OAc})_2$ (1.3 mg, 0.006 mmol, 5 mol%), K_2CO_3 (50.5 mg, 0.365 mmol, 3 equiv.), dppf (3 mg, 0.006 mmol, 5 mol%), 4-(trifluoromethyl)-phenylboronic acid (46 mg, 0.243 mmol, 2 equiv.) and DMF (0.5 mL). The reaction was stirred overnight at 80 °C. After completion and hydrolysis (H_2O , 10 mL) the reaction mixture was transferred to a separating funnel. The aqueous phase was extracted with ethyl acetate (3 x 15 mL) and the combined organic extracts washed by water (4 x 40 mL) and brine (40 mL) before being dried over MgSO_4 . After filtration and evaporation of the solvents under reduced pressure, the crude material was purified by flash column chromatography on silica gel using cyclohexane as eluent to afford the desired naphthalene **10** as a white solid (33.7 mg, 76% yield). mp 126-128 °C; ^1H NMR (300 MHz, CDCl_3) δ 8.11 (d, J 8.4 Hz, 1H), 7.75 – 7.67 (m, 1H), 7.61-7.55 (m, 1H), 7.49 – 7.39 (m, 4H), 7.35 – 7.24 (m, 5H), 7.22 – 7.14 (m, 2H), 2.81 (d, J 0.9 Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 146.0, 138.8, 136.6, 136.5, 134.3, 132.9, 132.3, 131.7, 130.5, 128.6, 128.1, 127.7, 127.1 (2C), 126.3, 126.1, 124.7, 124.6, 124.2, 19.7; ^{19}F NMR (282 MHz, CDCl_3) δ - 63.34. MS (EI) for $\text{C}_{24}\text{H}_{17}\text{F}_3$ m/z 362.15 (100) $[\text{M}]^+$.

References

1. Wang, Y. X.; Zhang, X. B.; Zhao, J.; Xie, S. Q.; Wang, C. J. *J. Med. Chem.* **2012**, *55*, 3502.
<http://dx.doi.org/10.1021/jm300168w>
2. Upadhayaya, R. S.; Vandavasi, J. K.; Kardile, R. A.; Lahore, S. V.; Dixit, S. S.; Deokar, H. S.; Shinde, P. D.; Sarmah, M. P.; Chattopadhyaya, J. *Eur. J. Med. Chem.* **2010**, *45*, 1854.
<http://dx.doi.org/10.1021/jm300168w>
3. Humljan, J.; Kotnik, M.; Contreras-Martel, C.; Blanot, D.; Urleb, U.; Dessen, A.; Sol-majer, T.; Gobec, S. *J. Med. Chem.* **2008**, *51*, 7486.
<http://dx.doi.org/10.1021/jm800762u>
4. Krohn, K.; Kounam, S. F.; Cludius, B. S.; Draeger, S.; Schulz, B. *Eur. J. Org. Chem.* **2008**, 3615.
<http://dx.doi.org/10.1002/ejoc.200800255>
5. Lowell, A. N.; Fennie, M. W.; Kozlowski, M. C. *J. Org. Chem.* **2008**, *73*, 1911.
<http://dx.doi.org/10.1021/jo7024114>
6. Dai, J.; Liu, Y.; Zhou, Y. D.; Nagle, D. G. *J. Nat. Prod.* **2007**, *70*, 1824.
<http://dx.doi.org/10.1021/np070337f>

7. Wang, Z.; Elokda, H.; McFarlane, G.; Pan, S.; Antane, M. *Tetrahedron Lett.* **2006**, *47*, 3365.
<http://dx.doi.org/10.1016/j.tetlet.2006.03.090>
8. Silva, O.; Gomes, E. T. *J. Nat. Prod.* **2003**, *66*, 447.
<http://dx.doi.org/10.1021/np0204904>
9. Rokade, Y. B.; Sayyed, R. Z. *Rasayan J. Chem.* **2009**, *2*, 972.
10. Pan, M.; Lin, X. M.; Li, G. B.; Su, C. Y. *Coord. Chem. Rev.* **2011**, *255*, 1921.
<http://dx.doi.org/10.1016/j.ccr.2011.03.013>
11. Anthony, J. E. *Angew. Chem.* **2008**, *120*, 460; *Angew. Chem. Int. Ed.* **2008**, *47*, 452.
<http://dx.doi.org/10.1002/anie.200604045>
12. Watson, M. D.; Fethtenkötter, A.; Müllen, K. *Chem. Rev.* **2001**, *101*, 1267.
<http://dx.doi.org/10.1021/cr990322p>
13. Shibasaki, M.; Matsunaga, S. in: *Privileged Chiral Ligands and Catalysts*; Q. L. Zhou (Ed.), Wiley, New York, 2011, pp 295–332 and references cited therein.
<http://dx.doi.org/10.1002/9783527635207.ch8>
14. Tan, A. K.; Fink, A. L. *Biochem. J.* **1992**, *281*, 191.
15. Kopp, R.; Pfeiffer, A. *Cancer Res.* **1990**, *50*, 6490.
16. Ullmann, H.; Meis, S.; Hongwiset, D.; Marzian, C.; Wiese, M.; Nickel, P.; Communi, D.; Boeynaems, J.-M.; Wolf, C.; Hausmann, R.; Schmalzing, G.; Kassack, M. U. *J. Med. Chem.* **2005**, *48*, 7040.
<http://dx.doi.org/10.1021/jm050301p>
17. Smith, M. B.; March, J. *Advanced Organic Chemistry*, 6th edn. Wiley, New York, 2007, Chapter 11, p 657.
18. Suzuki, A. in: *Modern Arene Chemistry*; D. Astruc (Ed.), Wiley-VCH, Weinheim, 2002.
19. Larock, R. C.; Doty, M. J.; Tian, Q.; Zenner, J. M. *J. Org. Chem.* **1997**, *62*, 7536.
<http://dx.doi.org/10.1021/jo9712557>
20. Larock, R. C.; Tian, Q. *J. Org. Chem.* **1998**, *63*, 2002.
<http://dx.doi.org/10.1021/jo972154b>
21. Larock, R. C. *J. Organomet. Chem.* **1999**, *576*, 111.
[http://dx.doi.org/10.1016/S0022-328X\(98\)01053-5](http://dx.doi.org/10.1016/S0022-328X(98)01053-5)
22. Ohno, H.; Yamamoto, M.; Iuchi, M.; Tanaka, T. *Angew. Chem. Int. Ed.* **2005**, *44*, 5103.
<http://dx.doi.org/10.1002/anie.200500159>
23. Ohno, H.; Yamamoto, M.; Iuchi, M.; Fujii, N.; Tanaka, T. *Synthesis* **2011**, 2567.
<http://dx.doi.org/10.1055/s-0030-1260098>
24. Asao, N.; Takahashi, K.; Lee, S.; Kasahara, T.; Yamamoto, Y. *J. Am. Chem. Soc.* **2002**, *124*, 12650.
<http://dx.doi.org/10.1021/ja028128z>
25. Asao, N.; Nogami, T.; Lee, S.; Yamamoto, Y. *J. Am. Chem. Soc.* **2003**, *125*, 10921.
<http://dx.doi.org/10.1021/ja036927r>

26. Asao, N.; Sato, K.; Menggenbateer; Yamamoto, Y. *J. Org. Chem.* **2005**, *70*, 3682.
<http://dx.doi.org/10.1021/jo0500434>
27. Asao, N. *Synlett* **2006**, 1645 and references cited therein.
<http://dx.doi.org/10.1021/jo0500434>
28. Isogai, Y.; Nawaz Khan, F.; Asao, N. *Tetrahedron* **2009**, *65*, 9575.
<http://dx.doi.org/10.1016/j.tet.2009.09.061>
29. Kabalka, G. W.; Ju, Y.; Wu, Z. *J. Org. Chem.* **2003**, *68*, 7915.
<http://dx.doi.org/10.1021/jo034330o>
30. Bu, X.; Hong, L.; Liu, R.; Hong, J.; Zhang, Z.; Zhou, X. *Tetrahedron* **2012**, *68*, 7960.
<http://dx.doi.org/10.1016/j.tet.2012.07.007>
31. Viswanathan, G. S.; Wang, M.; Li, C.-J. *Angew. Chem. Int. Ed.* **2002**, *41*, 2138.
[http://dx.doi.org/10.1002/1521-3773\(20020617\)41:12<2138::AID-ANIE2138>3.0.CO;2-T](http://dx.doi.org/10.1002/1521-3773(20020617)41:12<2138::AID-ANIE2138>3.0.CO;2-T)
32. Balamurugan, R.; Gudla, V. *Org. Lett.* **2009**, *11*, 3116.
<http://dx.doi.org/10.1021/ol900863d>
33. Gulda, V.; Balamurugan, R. *J. Org. Chem.* **2011**, *76*, 9919.
<http://dx.doi.org/10.1021/jo201918d>
34. Xiang, S.; Hu, H.; Ma, J.; Li, Y.; Wang, B.; Feng, C.; Zhao, K.; Hu, P.; Chen, X. *Sci. China Chem.* **2013**, *56*, 945.
<http://dx.doi.org/10.1007/s11426-013-4843-7>
35. Viswanathan, G. S.; Li, C.-J. *Synlett* **2002**, 1553.
36. Gulda, V.; Balamurugan, R. *Chem. Asian J.* **2013**, *8*, 414.
<http://dx.doi.org/10.1002/asia.201200817>
37. Umeda, R.; Nishi, S.; Kojima, A.; Kaiba, K.; Nishiyama, Y. *Tetrahedron Lett.* **2013**, *54*, 179.
<http://dx.doi.org/10.1016/j.tetlet.2012.10.123>
38. Wang, Z.-Q.; Liang, Y.; Lei, Y.; Zhou, M.-B.; Li, J.-H. *Chem. Commun.* **2009**, 5242.
<http://dx.doi.org/10.1039/b911669a>
39. Zhu, S.; Xiao, Y.; Guo, Z.; Jiang, H. *Org. Lett.* **2013**, *15*, 898.
<http://dx.doi.org/10.1021/ol4000394>
40. Boominathan, S. S. K.; Senadi, G. C.; Vandavasi, J. K.; Chen, J. Y.; Wang, J.-J. *Chem. Eur. J.* **2015**, *21*, 3193.
<http://dx.doi.org/10.1002/chem.201405695>
41. Manojveer, S.; Balamurugan, R. *Chem. Commun.* **2014**, *50*, 9925.
<http://dx.doi.org/10.1002/chem.201405695>
42. Manojveer, S.; Balamurugan, R. *Org. Lett.* **2014**, *16*, 1712.
<http://dx.doi.org/10.1021/ol5003835>
43. Inanaga, K.; Takasu, K.; Ihara, M. *J. Am. Chem. Soc.* **2005**, *127*, 3668.
<http://dx.doi.org/10.1021/ja042661s>
44. Boxer, M. B.; Yamamoto, H. *J. Am. Chem. Soc.* **2006**, *128*, 48.
<http://dx.doi.org/10.1021/ja054725k>

45. Boxer, M. B.; Yamamoto, H. *J. Am. Chem. Soc.* **2007**, *129*, 2762.
<http://dx.doi.org/10.1021/ja0693542>
46. Mundal, D. A.; Avetta, C. T.; Thomson, R. J. *Nat. Chem.* **2010**, *2*, 294.
<http://dx.doi.org/10.1038/nchem.576>
47. Ding, F.; William, R.; Wang, F.; Liu, X.-W. *Chem Commun.* **2012**, *48*, 8709.
<http://dx.doi.org/10.1039/c2cc33641c>
48. Ponra, S.; Vitale, M. R.; Michelet, V.; Ratovelomanana-Vidal, V. *J. Org. Chem.* **2015**, *80*, 3250.
<http://dx.doi.org/10.1021/acs.joc.5b00353>
49. Chinchilla, R.; Najera, C. *Chem. Rev.* **2007**, *107*, 874 and references cited therein.
<http://dx.doi.org/10.1021/cr050992x>
50. Lennox, A. J. J.; Lloyd-Jones, G. C. *Chem. Soc. Rev.* **2014**, *43*, 412 and references cited therein.
<http://dx.doi.org/10.1039/C3CS60197H>