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Halogen-bonded architectures of multivalent calix[4]arenes†

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A small family of novel halogen-bonded crystalline supramolecular architectures of calixarenes was obtained by the co-crystallization of cone (**1**) and 1,3-alternate tetrakis(3-iodopropargyloxy)calix[4]arene (**4**) as tetradentate halogen donors with different multidentate acceptors. Particularly interesting is the interpenetrated diamondoid network of **4** with DABCO, which represents the first example of a 2D network of calixarene macrocycles where halogen bonding is the key interaction for self-organization.

Introduction

Crystalline supramolecular infinite networks of calixarenes are attractive structures in the field of functional materials. Thanks to the combination of an aromatic cavity available for the complexation of small guests with the ease of functionalization of the phenol moieties, the calixarene macrocycle provides a versatile scaffold for the synthesis of hierarchically porous materials.¹ The insertion of appropriate ligating groups on the calixarene rims may result in macrocycle-containing tectons able to self-assemble in structures characterized by two levels of porosity, one associated with the macrocycle and one with the network.² The non-covalent bond most frequently exploited for the synthesis of this kind of architecture is the metal–ligand interaction, and a number of calixarene-based coordination polymers have been reported in recent decades,³ with promising adsorption,^{2,4,5} catalytic^{6–8} and nonlinear optical properties.⁹

On the other hand, for the formation of purely organic calixarene architectures, hydrogen bonding has been the interaction of choice for several years.¹⁰ Thanks to its strength and directionality, several hydrogen bonded 1D,^{11–13} 2D^{14,15} or 3D^{13,16} infinite networks of calixarenes have been reported. These are based either on the self-assembly of calixarenes functionalized with self-complementary hydrogen bonding groups, such as amides¹⁷ or carboxylic acids,¹³ or take advantage of an external multidentate hydrogen bonding donor or acceptor moiety to bridge complementarily functionalized macrocycles.^{12,15}

In recent years, also halogen bonds (XBs) have been frequently exploited in crystal engineering,¹⁸ thanks to some

favorable properties,¹⁹ such as their high directionality, strength tunability and resistance to solvent polarity.²⁰ Despite these advantages, however, examples of supramolecular networks of calixarenes assembled through XBs are extremely scarce. A few reports describe the crystal structures of upper rim iodo- and bromocalix[4]arenes²¹ and thiacalix[4]arenes,^{22–24} where the halogen atoms exchange XBs with the aromatic rings (π acceptors) or halogen and S atoms of neighboring calixarenes. Only one example takes advantage of an external bifunctional XB donor (a 1,4-diiodotetrafluorobenzene) to co-crystallize a calixarene functionalized at the lower rim with two pyridine units, working as a bifunctional acceptor.²⁵ In this case, due to the presence of additional π – π interactions between the external faces of two phenol rings of the macrocycle and the aromatic ring of the XB donor, a 2D network of calixarenes is formed.

With the aim of obtaining robust and possibly porous supramolecular networks of calixarene macrocycles assembled through XBs, we have recently reported the synthesis of tetrakis(3-iodopropargyloxy)calix[4]arene (**1**, Fig. 1) in the cone geometry as a tetravalent XB donor.²⁶ The iodoalkyne group is well-known to give strong XB interactions^{27,28} thanks to the increased electronegative character of the sp hybridized C atom linked to the iodine.²⁹ As expected, also when linked to the calixarene, this moiety confirmed its excellent XB donor ability. When crystallized in the absence of external XB acceptors, **1** self-assembles in a compact structure where all the iodine atoms are engaged in XBs with the alkyne C $\equiv$ C triple bonds of adjacent molecules (Fig. 2, top). On the contrary, in presence of 4,4'-bipyridine (BIPY), a divalent XB acceptor, the four iodine atoms form XBs with the BIPY nitrogen atoms giving rise to a 1D network of calixarenes alternated to couples of BIPYs (Fig. 2, bottom). To the best of our knowledge, this is the first example of a crystalline network of calixarene macrocycles where XB is the key interaction for self-assembly.³⁰

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quenched with 1 M HCl (15 mL). The resulting white precipitate was collected by filtration and recrystallized from CH₂Cl₂/methanol obtaining a mixture of 1,3-alternate derivative **3** and its partial cone isomer, from which **3** was isolated by column chromatography (CH₂Cl₂/hexane 75:25, v/v) in 32% yield (214 mg, 0.37 mmol). M.p. 183–184 °C. ¹H NMR (300 MHz, CDCl₃) δ (ppm): 7.16 (d, J = 7.5 Hz, 8H, ArH); 6.84 (t, J = 7.5 Hz, 4H, ArH); 4.13 (d, J = 2.4 Hz, 8H, OCH₂-C $\equiv$ CH); 3.80 (s, 8H, ArCH₂Ar); 2.47 (t, J = 2.4 Hz, 4H, OCH₂-C $\equiv$ CH). ¹³C NMR (100 MHz, CDCl₃) δ (ppm): 155.3 (OC_{Ar}); 134.2 (C_{Ar}); 130.4 (C_{Ar}); 123.1 (C_{Ar}); 80.4 (OCH₂C $\equiv$ CH); 74.6 (OCH₂C $\equiv$ CH); 58.9 (OCH₂C $\equiv$ CH); 37.0 (ArCH₂Ar). ESI-MS: m/z calcd for C₄₀H₃₂O₄Na [(3 + Na)⁺] 599.2, found 599.4; calcd for C₄₀H₃₂O₄K [(3 + K)⁺] 615.2, found 615.4.

1,3-Alternate 25,26,27,28-tetrakis(3-iodopropargyloxy) calix[4]arene (4). CuI (3.6 mg, 0.02 mmol) was added to a stirred solution of calixarene **3** (220 mg, 0.38 mmol) in dry THF (15 mL). After 15 min, *N*-iodomorpholine-HI (1.04 g, 3.05 mmol) was added and the mixture was stirred in the dark overnight. After TLC analysis (CH₂Cl₂/hexane 6:4, v/v) had revealed the complete conversion of calixarene **3**, 5 mL of a saturated solution of NH₄Cl and 10 mL of CH₂Cl₂ were added. The target compound precipitated as a whitish solid, which was isolated by filtration in 38% yield (155 mg, 0.14 mmol). M.p. 210 °C (dec.). ¹H NMR (400 MHz, DMSO-*d*₆) δ (ppm): 7.04 (d, J = 7.5 Hz, 8H, ArH); 6.76 (t, J = 7.5 Hz, 4H, ArH); 4.28 (s, 8H, ArOCH₂C $\equiv$ CI); 3.76 (s, 8H, ArCH₂Ar). ¹³C NMR (100 MHz, DMSO-*d*₆) δ (ppm): 156.1 (OC_{Ar}); 134.7 (C_{Ar}); 130.3 (C_{Ar}); 123.2 (C_{Ar}); 90.9 (OCH₂C $\equiv$ CI); 60.7 (OCH₂C $\equiv$ CI); 37.0 (ArCH₂Ar); 13.8 (OCH₂C $\equiv$ CI).

Single-crystal structure determination

Single crystal data were collected with a Bruker D8 Photon II diffractometer, with Mo (λ = 0.71073 Å) or Cu (λ = 1.54178 Å) X-ray source. The intensity data were integrated from several series of exposure frames covering the sphere of reciprocal space.³⁸ Absorption correction was applied using the program SADABS³⁹ for 4·4THF, 4·4DABCO·THF, and 1·2HMTA·3/2CH₂Cl₂. Crystals of **4** were non-merohedral twins and the absorption correction was applied with the TWINABS program.⁴⁰ The structures were solved with the ShelXT⁴¹ structure solution program using intrinsic phasing and refined with the ShelXL⁴² refinement package using least squares minimisation. Graphical material was prepared with the Mercury program.⁴³ A summary of data collection and structure refinement for **4**, 4·4THF, 4·4DABCO·THF, and 1·2HMTA·3/2CH₂Cl₂ is reported in Table 1. CCDC 2143352–2143355 contains the ESI† crystallographic data for this paper.

Results and discussion

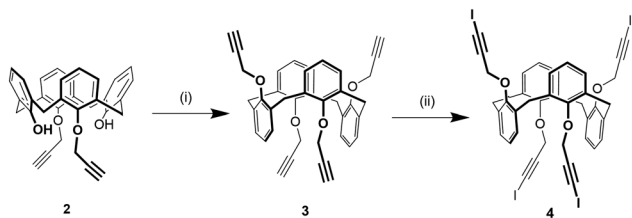
Synthesis

1,3-Alternate tetrakis(3-iodopropargyloxy)calix[4]arene **4** was synthesized in two steps from 1,3-di(propargyloxy)calix[4]arene **2**³⁶ (Scheme 1). First, compound **2** was alkylated with propargylbromide in presence of Cs₂CO₃ as a base obtaining a mixture of the 1,3-alternate and partial cone isomers of tetrakis(propargyloxy)calix[4]arene from which the desired compound **3** was isolated in 32% yield by column chromatography. This strategy was preferred to the direct alkylation of parent calix[4]arene⁴⁴ with excess

Table 1 Crystal data and structure refinement for **4**, 4·4THF, 4·4DABCO·THF, and 1·2HMTA·3/2CH₂Cl₂

	4	4·4DABCO·THF	4·4THF	1·2HMTA·3/2CH ₂ Cl ₂
Empirical formula	C ₄₀ H ₂₈ I ₄ O ₄	C ₅₆ H ₆₀ I ₄ N ₄ O ₅	C ₅₆ H ₆₀ I ₄ O ₈	C ₅₄ H ₅₆ Cl ₄ I ₄ N ₈ O ₄
Formula weight	1080.22	1376.68	1368.64	1530.46
Temperature/K	200	200	200	100
Crystal system	Monoclinic	Orthorhombic	Triclinic	Triclinic
Space group	<i>C2/c</i>	<i>Fddd</i>	<i>P1</i>	<i>P1</i>
<i>a</i> /Å	12.8824(8)	12.9348(2)	13.2357(16)	13.1998(18)
<i>b</i> /Å	19.0447(11)	22.8426(4)	15.3241(18)	14.997(2)
<i>c</i> /Å	15.1918(11)	39.6053(6)	15.3955(19)	30.958(4)
α /°	90	90	112.406(2)	100.754(4)
β /°	98.548(5)	90	101.238(2)	90.152(4)
γ /°	90	90	99.482(2)	105.402(4)
Volume/Å ³	3685.8(4)	11702.0(3)	2731.4(6)	5795.8(14)
<i>Z</i>	4	8	2	4
ρ_{calc} g cm ⁻³	1.947	1.563	1.664	1.754
μ /mm ⁻¹	26.874	2.177	2.333	2.386
<i>F</i> (000)	2048.0	5408.0	1344.0	2992.0
Crystal size/mm ³	0.09 × 0.06 × 0.05	0.19 × 0.15 × 0.15	0.28 × 0.23 × 0.16	0.16 × 0.14 × 0.08
Radiation (Å)	CuK α (λ = 1.54178)	MoK α (λ = 0.71073)	MoK α (λ = 0.71073)	MoK α (λ = 0.71073)
2 θ range (°)	8.35 to 133.16	4.756 to 51.454	2.98 to 53.632	4.024 to 51.542
Reflections collected	2999	17 013	34 973	137 466
<i>N</i> _{indep} (<i>R</i> _{int})	2999 (0.1373)	2779 (0.0281)	11 670 (0.0328)	22 020 (0.0762)
Data/restraints/parameters	2999/67/206	2779/0/157	11 670/110/705	22 020/19/1343
Goodness-of-fit on <i>F</i> ²	1.072	1.067	1.048	1.077
<i>R</i> ₁ [<i>I</i> ≥ 2 σ (<i>I</i>)]	0.0990	0.0246	0.0402	0.0743
<i>wR</i> ₂ [<i>I</i> ≥ 2 σ (<i>I</i>)]	0.2093	0.0543	0.0922	0.1778
$\Delta\rho_{\text{min/max}}$ (e Å ⁻³)	1.11/−0.74	0.51/−0.47	1.76/−1.55	3.58/−1.88





Scheme 1 Synthesis of 1,3-alternate tetrakis(3-iodopropargyloxy)calix[4]arene **4**: (i) propargyl bromide, Cs₂CO₃, dry acetone, reflux, 12 h, 32%; (ii) *N*-iodomorpholine hydrogen iodide, CuI, dry THF, 12 h, 38%.

propargylbromide due to an easier purification of the product. Subsequently, the iodoalkyne derivative **4** was obtained by reaction of **3** with *N*-iodomorpholine hydrogen iodide in 38% yield.

Solid state self-assembly

1,3-Alternate iodoalkenyl calixarene **4** resulted insoluble in all the common organic solvents except THF, dioxane, pyridine and DMSO. The reason for this insolubility, which was not observed for the cone isomer **1**, was attributed to multiple intermolecular XB interactions that can possibly form between the eight potential XB acceptors (the electron-rich π clouds of the $\text{C}\equiv\text{C}$ triple bonds and of the aromatic rings) and the four excellent XB donor groups (the iodoalkyne moieties). Moreover, the iodine atoms are positioned at the end of linear struts that are connected to the oxygen atoms of the calixarene through a rotationally unhindered methylene group and are therefore able to explore an ample area to install XB interactions. Only solvents that are good XB acceptors are therefore able to break the intermolecular XBs between the **4** molecules and dissolve the compound.

Two batches of single crystals of calixarene **4** have been obtained, one from a DMSO solution that was left standing at room temperature for a couple of months and the other by slow evaporation of a THF solution. X-ray diffraction analysis of a crystal from each batch revealed two different structures, the first consisting of the sole calixarene **4**, the other of a THF solvate.

Structure of calix[4]arene 4. It is instructive to compare the X-ray crystal structure of **4** (Fig. 3) and that of **1**²⁶ (Fig. 2, top). Even though the 3-iodopropargyloxy groups of **4** are linked to the 1,3-alternate calixarene scaffold, the iodoalkyne struts are not perpendicular to the mean plane passing through the methylene bridges, but are nearly parallel to each other and to this plane. Consequently, in this situation, **4** behaves as a linear XB donor synthon. This peculiar conformation allows both the iodine atoms I1 and I2 of two proximal iodopropargyloxy chains to asymmetrically interact with two carbon atoms of the phenyl rings of adjacent molecules, with the shortest distances I1...C5 (3.34 Å) and I2...C9 (3.27 Å). The first hierarchical structural motif is, therefore, a linear supramolecular chain of calixarenes. At a second level, the chains are interacting through offset $\pi\cdots\pi$

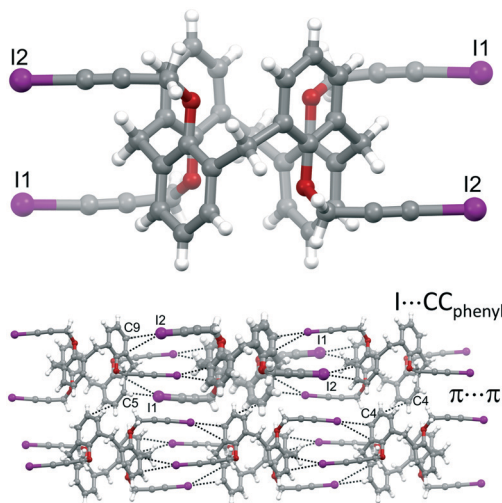


Fig. 3 Molecular structure of **4** (top), together with portion of the crystal packing highlighting intermolecular interactions (bottom).

stacking with symmetry related C4 and C10 atoms (C4 $\cdots$ C4, 3.39 Å, and C4 $\cdots$ C10, 3.42 Å, Fig. S1†). In a direction perpendicular to the $\pi\cdots\pi$ stacking, the 3-iodopropargyloxy moieties are exchanging CH $\cdots$ C_{alkyno} interactions (C16 $\cdots$ C20, 3.68 Å, and C17 $\cdots$ C19, 3.77 Å, Fig. S1†). On the opposite, in **1**, the four iodine atoms act as XB donors towards four alkyno groups of four different calixarenes. According to the location of the halogen atoms on the same side of the calixarene scaffold, the result is the formation of alternate layers dominated by the presence of iodine atoms or phenyl rings, respectively (Fig. 2, top).

Structure of the THF solvate of calix[4]arene 4. A different situation is encountered upon crystallization of **4** in THF, since the solvent molecules act as XBs acceptors towards the iodine atoms of the calixarene (Fig. 4). Hence, the presence of the THF molecules hinders the formation of the XBs between the iodine and the phenyl rings of adjacent macrocycles. The distance between iodine and oxygen is in the 2.72–2.89 Å range, which is significantly shorter than the van der Waals radii sum (approximately 3.50 Å). Despite the presence of THF molecules, which might have prevented the expansion of the chain-like motif found in **4**, the repetitive

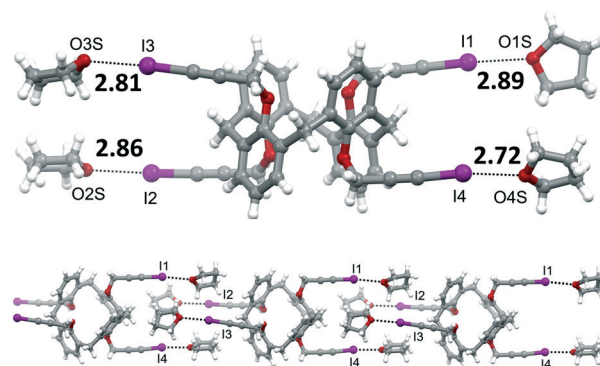


Fig. 4 Molecular structure of 4·4THF, with XB distances reported in Å (top). Portion of the supramolecular chain formed by 4·4THF (bottom).

unit 4·4THF expands linearly in a fashion surprisingly similar to that of **4**. The four 3-iodopropargyloxy·THF struts are then wedged in with those of symmetry related molecules (Fig. 4, bottom). In this system, the most acidic hydrogen of THF interacts with the carbon atoms of the alkyno groups (C16S···C40, 3.52 Å, Fig. S2†).

We then explored the ability of iodopropargylcalixarenes **1** and **4** to form self-assembled networks in presence of suitable multidentate XB acceptors by attempting to grow co-crystals of both calixarenes with DABCO and HMTA and of **4** with BIPY (see Table 2). While the high solubility of **1** in most organic solvents allowed us to try several crystallization solvent mixtures, for **4** we were limited to the use of THF, a very competitive XB acceptor. Co-crystals suitable for X-ray diffraction studies were only obtained for **1** with HMTA and for **4** with DABCO. When trying to grow crystals from a mixture of **1** and DABCO in CH₂Cl₂/hexane, we only obtained crystals of **1** having a structure identical to the one previously reported (Fig. 2, top),²⁶ and no crystalline material was collected from a mixture of **4** and HMTA in THF. The crystals grown from the sample containing **4** and BIPY, instead, consisted of the 4·4THF solvate, with no inclusion of BIPY in the structure. This can be viewed as a confirmation of the lower XB acceptor ability of the sp² N atoms of bipyridine with respect to the sp³ nitrogens of DABCO. Only the latter, in fact, are able to successfully compete with the THF solvent for halogen bonding with the iodine atoms.

Co-crystal structure of 1-HMTA. Co-crystals of **1** with HMTA were obtained by slow evaporation of a solution of **1** and HMTA (1 : 1, mol mol⁻¹) in CH₂Cl₂/hexane (1 : 1, v/v). The asymmetric unit comprises two calixarene moieties, four HMTA and three disordered CH₂Cl₂ molecules of crystallization, 1·2HMTA·3/2CH₂Cl₂. All of the iodine atoms of **1** are involved in XBs with the nitrogen atoms of HMTA. On the opposite, HMTA employs only two out of four nitrogen atoms to interact with the calixarenes. Interestingly, adjacent calixarenes engage XBs using two iodopropargyloxy moieties on the same couple of HMTA. Consequently, and according to the arrangement of the nitrogen atoms in HMTA at the vertices of an ideal tetrahedron, supramolecular zig-

zag chains can be identified, when viewing the calixarenes from the top (Fig. 5). The iodine···nitrogen distances are in the 2.78–2.92 Å range, which is significantly shorter than the van der Waals radii sum (approximately 3.53 Å). Furthermore, it is instructive to analyze the geometric parameters pertaining to the XBs within a single chain. Indeed, the two HMTA molecules interact differently with the 3-iodopropargyloxy systems. On one side, the XBs are shorter (~2.78 Å), the angle I···HMTA_{centroid}···I is significantly smaller than 109°, and the C–I···N angle deviates from the linearity (~171°). The other almost parallel motif is characterized by a longer XB distance (~2.92 Å), by a greater I···HMTA_{centroid}···I angle (~118°) and by a more linear XB geometry, with C–I···N of ~178°. The crystal packing is reported in Fig. S3,† showing a relatively compact structure, in which the dichloromethane molecules of crystallization occupy 13% of the unit cell volume.

Co-crystal structure of 4-DABCO. Co-crystals of **4** with DABCO were obtained by slow evaporation of a solution of **1** and DABCO (1 : 2, mol mol⁻¹) in THF. The asymmetric unit is represented by 1/4 of the calixarene, a DABCO molecule and 1/4 of disordered THF, with an overall molecular stoichiometry of 4·4DABCO·THF (Fig. 6).

The main structural difference, which is then reflected in the supramolecular arrangement, between **4** and 4·4THF with 4·4DABCO·THF is in the peripheral orientation of the iodopropargyl groups with respect to the corresponding phenyl rings. In 4·4DABCO·THF the four iodopropargyloxy struts point in four divergent directions in a conformation suitable to expand the interactions above and below the plane defined by the C atoms of the four methylene bridges. The torsion angle associated to this arrangement is close to 180° in 4·4DABCO·THF, whereas it varies in the 65–87° range for the other compounds. Moreover, in 4·4DABCO·THF the angle formed by two alternate linear C–I vectors is approximately 25° (Fig. 6, top). This overall calixarene arrangement, together with the divergent orientation of the N lone pairs of DABCO, results in the formation of a 2D diamondoid network. The N1···I1 distance is 2.74 Å, pointing to the presence of a strong XB. The crystal packing shows then the presence of a structural layer

Table 2 Summary of the crystallization outcomes and geometric parameters for the XB interactions

Entry	XB donor	External XB acceptor	Solvent	Crystal composition	XBs formed	XB length ^a	R ^b	C–I···A
1 ^c	1	—	CH ₂ Cl ₂	1	I···π cloud of C≡C	3.40–3.60 ^d	0.92–0.98	164.1–171.4
2 ^c	1	BIPY	CHCl ₃	1·3BIPY·CHCl ₃	I···N of BIPY	2.82–2.85	0.80–0.81	171.3–177.2
3	1	DABCO	CH ₂ Cl ₂	1	I···π cloud of C≡C (same as 1)			
4	1	HMTA	CH ₂ Cl ₂ /hexane	1·2HMTA·3/2CH ₂ Cl ₂	I···N of HMTA	2.78–2.92	0.79–0.83	170.7–179.2
5	4	—	DMSO	4	I···π cloud of Ar	3.27–3.34 ^d	0.89–0.91	165.5–168.9
6	4	—	THF	4·4THF	I···O of THF	2.72–2.89	0.78	170.7–176.6
7	4	BIPY	THF	4·4THF	I···O of THF (same as 6)			
8	4	DABCO	THF	4·4DABCO·THF	I···N of DABCO	2.74	0.78	178.8
9	4	HMTA	THF	No crystals	—			

^a Ranges are indicated when multiple interactions are present. ^b R is the normalized interaction distance, defined, according to Lommerse *et al.*,⁴⁷ as $R = d(I···A)/(r_I + r_A)$ where A represents the XB acceptor (C, N or O). ^c Reported in ref. 26. ^d Calculated considering the shortest I···C distance for each XB.



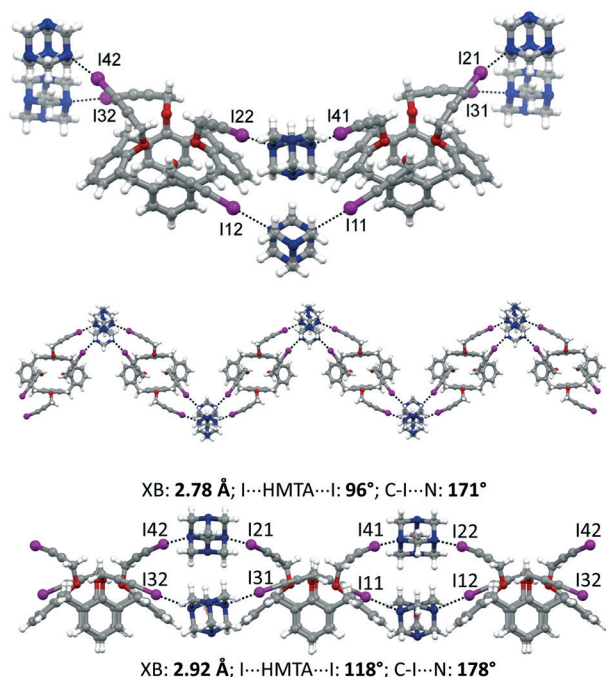


Fig. 5 Molecular structure of 1-2HMTA-3/2CH₂Cl₂ (top). Zig-zag supramolecular chain (middle and bottom), with relevant geometric parameters.

formed by *n*-interpenetrated networks, which are parallel to the *ab* crystallographic plane (Fig. S4†). Embedded between the calixarene frameworks is located the THF of crystallization. Interestingly, even after applying moderate vacuum ($P \sim 1$ mbar) the THF could still be identified by the difference Fourier map, suggesting a relative capability of the framework in trapping the solvent molecule. The volume occupied by THF is approximately 16% of the unit cell volume.

Conclusions

Together with the previously reported structures of cone tetraiodopropargyloxy calix[4]arene **1** and of the co-crystal **1-BIPY**, these results confirm the excellent XB ability of the iodoalkyne strut.²⁹ Systematically, in all the structures, all the iodine atoms are engaged in XB interactions with the same type of acceptor, even when more than one nucleophilic sites are present in the crystallization environment. However, as seen from Table 2, the outcome of the crystallization experiments, when two or three different XB acceptors are available, was in some cases surprising. DMSO, which was reported to be an effective XB acceptor for iodoethynyl moieties,^{45,46} when used as crystallization solvent, was not halogen-bonded to calixarene **4**, which, instead, self-assembles at the solid state through $I \cdots \pi$ XBs. Calixarene **1** forms XBs with BIPY and HMTA, but not with DABCO, which is the most Lewis basic of the three (Table 2, entries 2–4). On the contrary, 1,3-alternate **4** produces a halogen-bonded network only with DABCO (Table 2, entries 7–9). These observations confirm that, due to the moderate strength of XB interactions, the resulting architectures are driven by a

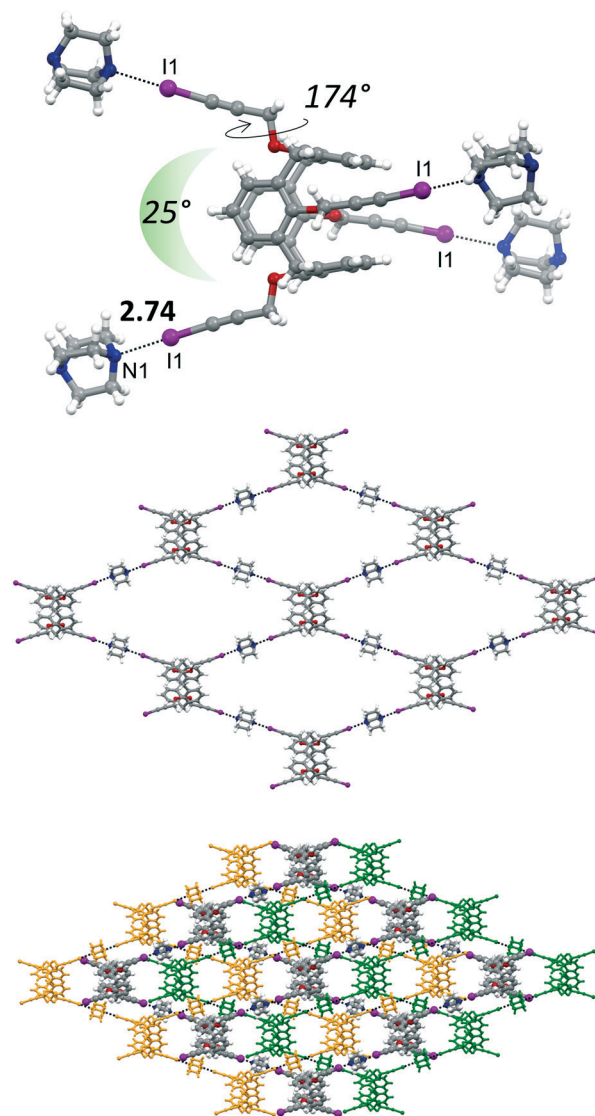


Fig. 6 Molecular structure of 4-4DABCO-THF, with XB distance reported in Å (top), single diamondoid network (middle), interpenetrated diamondoid networks (bottom). Solvent of crystallization was omitted for clarity.

subtle interplay of other different factors, which include the spatial orientation and the rotational freedom of the XB donors, the magnitude of secondary supramolecular interactions and the competition by a nucleophilic solvent.³⁴ Nonetheless, the reported crystal structures demonstrate the effectiveness of XBs as key interactions for the self-assembly of supramolecular networks of calixarenes. The dimensionality of these architectures, however, proved hard to predict, and despite the multivalency of both the calixarenes and the XB acceptors, in all but one co-crystals, the main self-assembled structures are 1D chains of alternating calixarenes and XB acceptors. Only for **4-DABCO**, the combination of the 1,3-alternate conformation of the macrocycle with the valence and geometry of DABCO allowed to obtain the first example of a 2D halogen-bonded calix[4]arene architecture. Yet, we believe our results provide



additional insights into the design of crystalline supramolecular networks of calixarene macrocycles, and further studies are currently underway in our laboratories towards the synthesis of robust 3D supramolecular networks where the macrocycle aromatic cavity could be exploited for host-guest properties. In particular, we plan to extend this approach also to the calix[6]- and calix[8]arene scaffolds, whose higher valency and larger size could facilitate the synthesis of halogen-bonded architectures of higher dimensionality and improved complexation ability.

Author contributions

M. C. Gullo: methodology and investigation; L. Marchiò: methodology, visualization, writing – original draft; A. Casnati: supervision, writing – review & editing, funding acquisition; L. Baldini: conceptualization, supervision, writing – original draft.

Conflicts of interest

There are no conflicts of interest to declare.

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Notes and references

- H. Zhang, R. Zou and Y. Zhao, *Coord. Chem. Rev.*, 2015, **292**, 74–90.
- S. P. Bew, A. D. Burrows, T. Düren, M. F. Mahon, P. Z. Moghadam, V. M. Sebestyen and S. Thurston, *Chem. Commun.*, 2012, **48**, 4824–4826.
- A. Ovsyannikov, S. Solovieva, I. Antipin and S. Ferlay, *Coord. Chem. Rev.*, 2017, **352**, 151–186.
- Z. Zhang, A. Drapailo, Y. Matvieiev, L. Wojtas and M. J. Zaworotko, *Chem. Commun.*, 2013, **49**, 8353–8355.
- E. Lee, Y. Kim, J. Heo and K. M. Park, *Cryst. Growth Des.*, 2015, **15**, 3556–3560.
- L. L. Liu, J. Chen, C. X. Yu, W. X. Lv, H. Y. Yu, X. Q. Cui and L. Liu, *Dalton Trans.*, 2017, **46**, 178–185.
- C.-X. Yu, F.-L. Hu, M.-Y. Liu, C.-W. Zhang, Y.-H. Lv, S.-K. Mao and L.-L. Liu, *Cryst. Growth Des.*, 2017, **17**, 5441–5448.
- L.-J. Yue, Y.-Y. Liu, G.-H. Xu and J.-F. Ma, *New J. Chem.*, 2019, **43**, 15871–15878.
- Y. J. Liu, J. S. Huang, S. S. Y. Chui, C. H. Li, J. L. Zuo, N. Zhu and C. M. Che, *Inorg. Chem.*, 2008, **47**, 11514–11518.
- M. J. Hardie, *Hydrogen Bonded Network Structures Constructed from Molecular Hosts*, Springer, Berlin, Heidelberg, 2004, pp. 139–174.
- W. Wang, S. Gong, Y. Chen and J. Ma, *New J. Chem.*, 2005, **29**, 1390–1392.
- W. Jaunky, M. W. Hosseini, J. M. Planeix, A. De Cian, N. Kyritsakas and J. Fischer, *Chem. Commun.*, 1999, 2313–2314.
- Z. Xiao, W. Yang, F. Yan, L. Ji, W. Li and W. Wang, *CrystEngComm*, 2019, **21**, 439–448.
- L. Baldini, F. Sansone, C. Massera, A. Casnati, F. Ugozzoli and R. Ungaro, *Inorg. Chim. Acta*, 2007, **360**, 970–980.
- Y. Liu and M. D. Ward, *Cryst. Growth Des.*, 2009, **9**, 3859–3861.
- Y. Li, W. Yang, Y. Chen and S. Gong, *CrystEngComm*, 2011, **13**, 259–268.
- L. Baldini, F. Sansone, A. Casnati, F. Ugozzoli and R. Ungaro, *J. Supramol. Chem.*, 2002, **2**, 219–226.
- G. R. Desiraju, *J. Am. Chem. Soc.*, 2013, **135**, 9952–9967.
- G. Cavallo, P. Metrangolo, R. Milani, T. Pilati, A. Priimagi, G. Resnati and G. Terraneo, *Chem. Rev.*, 2016, **116**, 2478–2601.
- C. C. Robertson, R. N. Perutz, L. Brammer and C. A. Hunter, *Chem. Sci.*, 2014, **5**, 4179–4183.
- S. R. Kennedy, M. U. Main, C. R. Pulham, I. Ling and S. J. Dalgarno, *CrystEngComm*, 2019, **21**, 786–790.
- M. Yamada, Y. Ootashiro, Y. Kondo and F. Hamada, *Tetrahedron Lett.*, 2013, **54**, 1510–1514.
- M. Yamada, R. Kanazawa and F. Hamada, *CrystEngComm*, 2014, **16**, 2605–2614.
- M. Yamada and F. Hamada, *Cryst. Growth Des.*, 2015, **15**, 1889–1897.
- M. T. Messina, P. Metrangolo, S. Pappalardo, M. F. Parisi, T. Pilati and G. Resnati, *Chem. – Eur. J.*, 2000, **6**, 3495–3500.
- M. C. Gullo, L. Baldini, A. Casnati and L. Marchiò, *Cryst. Growth Des.*, 2020, **20**, 3611–3616.
- L. Turunen, N. K. Beyeh, F. Pan, A. Valkonen and K. Rissanen, *Chem. Commun.*, 2014, **50**, 15920–15923.
- C. B. Aakeröy, T. K. Wijethunga, J. Desper and M. Đaković, *Cryst. Growth Des.*, 2015, **15**, 3853–3861.
- C. B. Aakeröy, M. Baldrighi, J. Desper, P. Metrangolo and G. Resnati, *Chem. – Eur. J.*, 2013, **19**, 16240–16247.
- On the contrary, few examples of halogen bonded networks of resorcinarene cavitands have been reported, see for example: C. B. Aakeröy, A. Rajbanshi, P. Metrangolo, G. Resnati, M. F. Parisi, J. Desper and T. Pilati, *CrystEngComm*, 2012, **14**, 6366–6368 and ref. 27 and 34.
- K. Raatikainen and K. Rissanen, *CrystEngComm*, 2011, **13**, 6972–6977.
- C. Perkins, S. Libri, H. Adams and L. Brammer, *CrystEngComm*, 2012, **14**, 3033–3038.
- L. Catalano, S. Pérez-Estrada, G. Terraneo, T. Pilati, G. Resnati, P. Metrangolo and M. A. Garcia-Garibay, *J. Am. Chem. Soc.*, 2015, **137**, 15386–15389.
- L. Turunen, F. Pan, N. K. Beyeh, J. F. Trant, R. H. A. Ras and K. Rissanen, *Cryst. Growth Des.*, 2018, **18**, 513–520.
- G. Anyfanti, A. Bauzá, L. Gentiluomo, J. Rodrigues, G. Portalone, A. Frontera, K. Rissanen and R. Puttreddy, *Front. Chem.*, 2021, 165.
- W. Xu, J. J. Vittal and R. J. Puddephatt, *Can. J. Chem.*, 2011, **74**, 766–774.



- 37 J. E. Hein, J. C. Tripp, L. B. Krasnova, K. B. Sharpless and V. V. Fokin, *Angew. Chem., Int. Ed.*, 2009, **48**, 8018–8021.
- 38 Bruker, *SMART*, Bruker AXS Inc., Madison, Wisconsin, USA, 2012.
- 39 L. Krause, R. Herbst-Irmer, G. M. Sheldrick and D. Stalke, *J. Appl. Crystallogr.*, 2015, **48**, 3–10.
- 40 G. M. Sheldrick, *TWINABS_2012/1*, University of Göttingen, Germany, 1996.
- 41 G. M. Sheldrick, *Acta Crystallogr., Sect. A: Found. Crystallogr.*, 2015, **71**, 3–8.
- 42 G. M. Sheldrick, *Acta Crystallogr., Sect. C: Struct. Chem.*, 2015, **71**, 3–8.
- 43 C. F. Macrae, P. R. Edgington, P. McCabe, E. Pidcock, G. P. Shields, R. Taylor, M. Towler and J. Van De Streek, *J. Appl. Crystallogr.*, 2006, **39**, 453–457.
- 44 W. Verboom, S. Datta, Z. Asfari, D. N. Reinhoudt and S. Harkema, *J. Org. Chem.*, 1992, **57**, 5394–5398.
- 45 J. Lieffrig, O. Jeannin and M. Fourmigué, *J. Am. Chem. Soc.*, 2013, **135**, 6200–6210.
- 46 L. Turunen, F. Pan, N. K. Beyeh, M. Cetina, J. F. Trant, R. H. A. Ras and K. Rissanen, *CrystEngComm*, 2017, **19**, 5223–5229.
- 47 J. P. M. Lommerse, A. J. Stone, R. Taylor and F. H. Allen, *J. Am. Chem. Soc.*, 1996, **118**, 3108–3116.

