Halogen bonding of N-bromosuccinimide by grinding†

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Two halogen bonded cocrystals of N-bromosuccinimide and 4,4′-bipyridine, with stoichiometric ratios 1:1 and 2:1, have been synthesized and characterized. We present the first mechanochemical cocrystallization of N-bromosuccinimide.

Halogen bonds are non-covalent interactions that form between polarisable and electron-deficient halogen atoms (Br, I) and electron-rich Lewis bases (nucleophilic atoms such as O, N, S, etc.). They have been utilized as alternatives to hydrogen bonds in crystal engineering of organic and metal-organic materials. An overview of the currently available literature reveals that studies of halogen bonding have largely focused on cocrystals of perfluorohalocarbons as classic halogen bond donors. Numerous reports have been published over the past decade dealing with crystal engineering and synthesis of multicomponent materials containing iodo-substituted perfluorinated donors. Recently, Rissanen and Raatikainen have introduced a new family of halogen bond donors. They have demonstrated how N-halosuccinimides can be used for constructing halogen bonded molecular crystals. Also, in a recent study, Fournigué and Espinosa described N-iodoimide derivatives as strong halogen bond donors. N-halosuccinimides are commercially available inexpensive compounds, commonly used as halogenation agents in organic synthesis. They possess an extremely polarised halogen atom but have not yet been fully recognized and studied as potential halogen bond donors. A cursory search of the Cambridge Structural Database (CSD) based on the ability of N-halosuccinimides to act as a halogen bond donor has resulted in 13 hits; 10 of those have been derived from N-iodosuccinimide (nis) and 3 from N-bromosuccinimide (nbs).

Herein we report a mechanochemical approach to the construction of halogen-bonded cocrystals of nbs. We have selected 4,4′-bipyridine (bpy) as a good counterpart to nbs, a molecule that is a commonly used ditopic hydrogen or halogen bond acceptor (Fig. 1). Compared to the traditional cocrystallization method, synthesis from a solution, mechanochemical synthesis has been proven superior and is preferable because it sidesteps finding an appropriate solvent or solvent mixture. In general it is much faster, therefore, unexpected products resulting from unwanted reactions such as oxidation, halogenation or solvolysis are avoided. To the best of our knowledge, this is the first mechanosynthesis of multicomponent halogen-bonded solids using nbs via neat grinding (NG) and liquid-assisted grinding (LAG). However, numerous reports have been published dealing with mechanochemical approaches to halogenation using nbs and nis compounds.

In order to explore the stoichiometric ratio of nbs and bpy as well as the reactivity in the solid state, we first attempted the mechanochemical synthesis of the cocrystals by NG and LAG of nbs with bpy in the stoichiometric ratios 1:1 and 2:1, respectively.

Fig. 1 Structures of reactants, N-bromosuccinimide (nbs) and 4,4′-bipyridine (bpy), and the cocrystal structures obtained as a result of the single crystal X-ray diffraction experiment.
respectively. Milling was conducted in a Retsch MM200 mill using a stainless steel milling assembly (see the ESI†) under normal laboratory conditions (temperature ca. 25 °C, 40–50% RH). To observe the grinding experiments, as well as to facilitate the characterisation of the new cocrystals by single-crystal X-ray diffraction, mechanochemical experiments were accompanied by crystallization from the solution. All reactants and products have been characterised by means of powder X-ray diffraction (PXRD) and differential scanning calorimetry (DSC). NG of bpy with nbs in a stoichiometric ratio of 1:1 for 15 min afforded a (nbs)(bpy) cocrystal, as proven by PXRD, but the NG experiment in a stoichiometric ratio of 2:1 for 15 min afforded a mixture of products, where the (nbs)(bpy) cocrystal was present as the major product and the (nbs)2(bpy) cocrystal only in traces (see the ESI†). When the same NG experiment was carried out for 80 min, the reaction was still not quantitatively complete – this time the experiment resulted in a mixture with the (nbs)2(bpy) cocrystal as the major product (see the ESI†). Considering that a liquid phase can significantly enhance the scope and rate of the observation indicating the presence of a strong halogen bond with a partially covalent character and with charge transfer between the N–Br bond to the Br molecule is extended compared to that observed for the pure nbs crystal16 (0.03 Å for the 1:1 cocrystal; 0.07 Å and 0.08 Å for the 2:1 cocrystal). All these observations indicate the presence of a strong halogen bond with a partially covalent character and with charge transfer from the N–Br bond to the Br–N contact.17 Considering that an nbs molecule possesses carbonyl groups and is also a strong hydrogen bond acceptor, components in both cocrystals create a specific halogen and hydrogen bonding pattern. The crystal structure of (nbs)(bpy) can be described by the association of 1:1 halogen bonded complexes into a hydrogen bonded 3D network via the C–H...O hydrogen bonds between nbs and bpy (C9⋯O1 contact of 3.542(3) Å, C14⋯O2 contact of 3.342(3) Å and C11⋯O2 contact of 3.356(3) Å and via the C–H...O hydrogen bonds between nbs molecules (C3⋯O1 contact of 3.412(3) Å and C2⋯O1 contact of 3.384(3) Å). A notable detail of the cocrystal structure is with those calculated from the single crystal data (Fig. 2, see the ESI†). The (bpy)[nbs] and (nbs)2(bpy) cocrystals crystallize in the monoclinic system with four formula units per unit cell. The molecular structures of the cocrystals with the atom numbering scheme are given in Fig. S1 (see the ESI†). In the crystal structure of (bpy)[nbs], each nbs molecule is associated with a bpy molecule via an N⋯Br halogen bond. The Br⋯N distance of 2.386(2) Å is 23.9% less than the sum of the van der Waals (VDW) radii of nitrogen and bromine (3.4 Å) and almost linear, with an N–Br⋯N angle of 171.20(6)°. Only one of the two nitrogen atoms in each bpy molecule is involved in bonding with an nbs molecule (Fig. 3). Furthermore, crystalline structure determination of (nbs)2(bpy) revealed two pairs of symmetry equivalent (CO)2N–Br⋯N halogen bonds with contact distances of 2.358(3) and 2.374(2) Å, which are 30.6 and 30.2% less than the sum of the VDW radii, respectively. Both contacts are highly linear, with an N–Br⋯N angle of 176.5(1)° and 179.0(1)°. For both cocrystals, the N–Br bond in the nbs molecules is extended compared to that observed for the pure nbs crystal16 (0.03 Å for the 1:1 cocrystal; 0.07 Å and 0.08 Å for the 2:1 cocrystal). All these observations indicate the presence of a strong halogen bond with a partially covalent character and with charge transfer from the N–Br bond to the Br⋯N contact.17 Considering that an nbs molecule possesses carbonyl groups and is also a strong hydrogen bond acceptor, components in both cocrystals create a specific halogen and hydrogen bonding pattern. The crystal structure of (nbs)(bpy) can be described by the association of 1:1 halogen bonded complexes into a hydrogen bonded 3D network via the C–H...O hydrogen bonds between nbs and bpy (C9⋯O1 contact of 3.542(3) Å, C14⋯O2 contact of 3.342(3) Å and C11⋯O2 contact of 3.356(3) Å and via the C–H...O hydrogen bonds between nbs molecules (C3⋯O1 contact of 3.412(3) Å and C2⋯O1 contact of 3.384(3) Å). A notable detail of the cocrystal structure is
the fact that each nbs molecule is connected to seven neighbouring molecules (three nbs and four bpy) as a halogen bond donor, ditopic hydrogen bond donor and ditopic hydrogen bond acceptor (Fig. 3). As in the 1 : 1 cocrystal, the crystal structure of \((\text{nbs})_2(\text{bpy})\) can be described by the association of 2 : 1 halogen bonded complexes into a hydrogen bonded 3D network via the C–H⋯O hydrogen bonds. The crystal structure contains two symmetrically nonequivalent nbs and bpy molecules (Fig. 4). One bpy molecule is connected with four neighbouring nbs molecules via the described halogen bond and bifurcated C–H⋯O hydrogen bonds (C15⋯O3 contact of 3.477(4) Å and C17⋯O3 contact of 3.341(5) Å). The second bpy molecule is connected with six neighbouring nbs molecules via the described halogen bond, C–H⋯O hydrogen bonds (C9⋯O4 contact of 3.515(4) Å), and via the bifurcated C–H⋯O hydrogen bonds (C10⋯O2 contact of 3.403(4) Å and C12⋯O2 contact of 3.434(4) Å). The overall structure results from the combination of such interactions and it is additionally stabilized by the C–H⋯O hydrogen bonds between nbs molecules (C3⋯O4 contact of 3.406(5) Å). Furthermore, the described interactions in both cocrystals can be observed by close analysis of the two-dimensional fingerprint plot derived from the Hirshfeld surface\(^1\) of the cocrystal components (see the ESI†). Thermal analysis of the prepared cocrystals by DSC reveals that both compounds are of similar thermal stability. The DSC curves of both cocrystals have the same profile and show one weak endothermic change peak and one strong exothermic change peak. They occur for (nbs)(bpy) at 116 °C (3.5 kJ mol\(^{-1}\)) and 131 °C (58.4 kJ mol\(^{-1}\)), and for (nbs)_2(bpy) at 103 °C (3.7 kJ mol\(^{-1}\)) and 136 °C (114.6 kJ mol\(^{-1}\)) (see the ESI†). For both cocrystals, the exothermic change peak corresponds to the cocrystal decomposition and is at a lower temperature than that of the pure nbs reactant (208 °C, 42.9 kJ mol\(^{-1}\)).

Conclusions

In summary, we have demonstrated how nbs, a commercially available compound commonly used as a brominating agent in organic synthesis, can be used in cocrystal synthesis. To the best of our knowledge, this is the first demonstration of mechanochemistry of multicomponent halogen-bonded solids using nbs as a halogen bond donor. The mechanochemical approach exhibits notable advantages over conventional solution-based synthesis of such compounds, not only through reducing the time of preparation and isolation, but also by evading the formation of potentially unwanted halogenation products. Crystal structure analysis of (nbs)(bpy) and (nbs)_2(bpy) demonstrated that the presence of a polarised halogen atom and two carbonyl groups on the nbs molecule leads to a specific halogen and hydrogen bonding pattern in their crystal structure. The high stability of both materials, as a result of halogen and hydrogen bonding patterns, was proven by a relatively high decomposition temperature. Also, crystal structure analysis revealed indications of a very strong halogen bond with a partially covalent character and with charge transfer from the N–Br bond to the Br⋯N contact. We believe that the described results will have significant impact on the design and synthesis of future halogen-bonded materials containing N-haloimides as well as on halogen bond-driven crystal engineering in general.

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Notes and references


15 The amount of liquid was 40 μL per 200 mg of reactants, corresponding to a liquid-to-solid ratio of η = 0.2 μL mg⁻¹, see: T. Friščič, S. L. Childs, S. A. A. Rizvi and W. Jones, CrystEngComm, 2009, 11, 418.

