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ARTICLE

Nitration Pattern of Energetic 3,6-Diamino-1, 2, 4, 5-tetrazine Derivatives Containing Azole Functional Groups

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One of the successful strategies for the design of promising new energetic materials is the incorporation of both fuel and oxidizer moieties into the same molecule. Therefore, during recent years, synthesis of various nitro-azole derivatives, as compounds with a more balanced oxygen content, became very popular. In the frame of this effort, we studied nitration of N^3, N^6 -bis(1H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (**BTATz**; **5**) and its alkylated derivative N^3, N^6 -bis(2-methyl-2H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **12**, using a ^{15}N -labeled nitration agent and monitoring and analyzing products of these reactions by ^{15}N NMR. It was shown that the nitration of both compounds takes place only on the exocyclic ("bridging") secondary amine groups. Possible tetranitro derivative isomers N, N' -(1,2,4,5-tetrazine-3,6-diyl)bis(N -(1-nitro-1H-tetrazol-5-yl)-nitramide) **6** and N, N' -(1,2,4,5-tetrazine-3,6-diyl)bis(N -(2-nitro-2H-tetrazol-5-yl)nitramide) **7**, both of which have OB = 0% and calculated VODs of 9,790 and 9,903 m/sec, respectively, could not be observed in the reaction mixtures, during the *in situ* ^{15}N NMR monitoring of nitration of **5**, using ^{15}N -labeled nitrating agents. Following a similar strategy, a new analog of **BTATz** – N^3, N^6 -Bis(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **15** was obtained and its nitration was studied. The reaction of **15** with a $\text{HNO}_3\text{-Ac}_2\text{O}$ nitration mixture resulted in the formation of a new N^3, N^6 -bis(3-nitro-1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine derivative **20** in a moderate yield. Structures and properties of **15** (in the form of its perchlorate salt, **16**) and **20** were measured by FTIR, multinuclear NMR, MS, DSC and x-ray crystallography. It is important to note that compound **20** exhibits exothermic decomposition at 302 °C (DSC) and >353 N (sensitivity to friction), making it a highly-promising thermally-insensitive energetic material for further development.

Introduction

The history of discovery and development of new energetic materials goes back to gunpowder, which was invented in Imperial China around the 9th century, in an attempt to create an immortality potion.¹ This ground-breaking discovery led to the invention of fireworks and a variety of weapons in China and, later, throughout the world.² Since the earliest findings, tremendous progress was made in the rational design and preparation of novel energetic materials and compositions.³ Important properties of such materials, which are commonly taken into account by the designers span heat of formation (HOF), density, melting and decomposition temperatures, sensitivity to friction and shock, velocity of detonation (VOD), synthetic complexity,⁴ as well as carbon content, nitrogen content and oxygen balance (OB).⁵

Energetic materials typically contain both oxidizing and reducing functional groups in their molecular structure (or in

the structure of their components). Under high temperature and pressure conditions, these materials would transform into more thermodynamically-stable products, including small molecules with low heats of formation, such as H_2O , N_2 , CO , CO_2 , SO_2 and metal oxides.⁶ The OB is a mathematical formula used to calculate the degree to which a given explosive or propellant could be oxidized. A "zero" OB value will be calculated when the chemical composition of the calculated energetic material will have the exact amount of oxygen atoms needed to convert all the carbon atoms to CO_2 , all hydrogen atoms to H_2O , all sulphur atoms to SO_2 and all metal atoms (if present in the material) to metal oxides. An energetic material would have a positive OB value if it contains more oxygen atoms than required for complete combustion and a negative OB value when the amount of oxygen atoms is insufficient for complete oxidation. The results of OB calculations were shown to have an excellent correlation with both sensitivity properties and performance of energetic compounds and their formulations, which have a tendency to reach their best values as their OB values are getting closer to "zero".⁷ When an energetic material has a negative OB value (an insufficient amount of oxygen for complete oxidation), it will typically exhibit an incomplete combustion, resulting in the formation of large amounts of toxic CO gas, smoke, soot and solid residues. Commonly, as the OB values for a certain explosive

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† Footnotes relating to the title and/or authors should appear here. Electronic Supplementary Information (ESI) available: [details of any supplementary information available should be included here]. See DOI: 10.1039/x0xx00000x

get lower, poorer performance for this explosive is observed, and the VOD and generated pressure for this explosive also become smaller.

In cases where an energetic material contains “too much oxygen” (has a positive OB value), the O₂ produced during explosion is absorbing a significant amount of energy, substantially reducing this explosive performance.⁸

There are several fascinating examples of energetic compounds possessing an OB value of 0%, such as the most potent chemical explosive known – octanitrocubane (ONC; R.E. factor = 2.38).⁹ Other examples include the recently prepared compound nitril cyanide¹⁰ and the still synthetically-elusive nitrogen-rich compounds – 3,6-dinitro-1,2,4,5-tetrazine,¹¹ 2,4,6-trinitro-1,3,5-triazine,¹² [1,2,3,4]-tetrazino-[5,6-e][1,2,3,4]-tetrazine-1,3,5,7-tetraoxide (TTTO),¹³ and (5-nitro-2H-tetrazol-2-yl)-methyl nitrate (Fig. 1).¹⁴

Due to their high density, highly positive enthalpy of formation, good detonation performances and excellent thermal stability in comparison to the conventional energetic materials, nitrogen-rich energetic compounds attract considerable scientific attention.¹⁵ The latter compounds were extensively explored during recent years for their great potential for both civilian and military applications such as gas generators, low-signature propellants, as well as additives to pyrotechnics and explosives.¹⁶ Among popular building blocks used in the construction of nitrogen-rich energetic materials are tetrazole¹⁷ and tetrazine¹⁸ functional groups. However, the majority of nitrogen-rich organic compounds are oxygen-deficient. Since one of the most popular strategies for the design of promising new energetic materials is incorporating both fuel and oxidizer properties into a single molecule, one of the important challenges in this field is the preparation of new nitrogen-rich compounds with a low OB value. Therefore, in recent years, the synthesis and evaluation of various nitroazole derivatives as compounds with a balanced oxygen content became highly-popular.¹⁹

Results and discussion

In our perspective, as part of the framework of these efforts, one of the interesting challenges was exploring the synthesis of unreported tetra-nitro derivatives of *N*³,*N*⁶-bis(1*H*-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (BTATz **5**; OB = -64.5%)²⁰ –

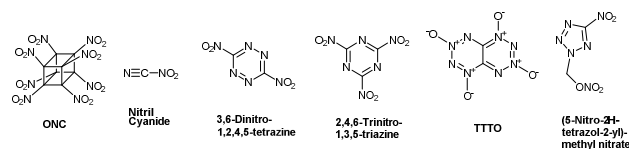


Fig. 1 Structures of ONC, nitril cyanide, 3,6-dinitro-1,2,4,5-tetrazine, 2,4,6-trinitro-1,3,5-triazine, TTTO and (5-nitro-2H-tetrazol-2-yl)-methyl nitrate.

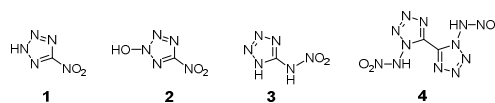


Fig. 2 Structures of compounds **1** (OB = -7.0%), **2** (OB = +6.1%), **3** (OB = -12.3%) and **4** (OB = -3.1%).

isomers *N,N'*-(1,2,4,5-tetrazine-3,6-diyl)bis(*N*-(1-nitro-1*H*-tetrazol-5-yl)-nitramide) **6** and *N,N'*-(1,2,4,5-tetrazine-3,6-diyl)bis(*N*-(2-nitro-2*H*-tetrazol-5-yl)nitramide) **7**, both of which have OB values of 0% and calculated VODs of 9,790 and 9,903 m/sec, respectively (Fig. 3, Table 1). Although it would seem that BTATz analogues exhibiting better OB values should be of significant interest, only a single example of such derivatives – 1,4-di-*N*-oxide **8** (OB = -45.7%) – was prepared by Chaves and coworkers.^{21a} Unfortunately, compound **8** displayed decomposition at 134 °C (*versus* decomposition of BTATz at 318 °C), indicating that other derivatisation strategies should be investigated.^{21b}

Approaches to prepare energetic molecules with improved OB values are frequently based on the conversion of NH groups in the starting materials into N-NO₂ groups in the corresponding, more energetic, derivatives.²² Since the BTATz molecule has two pairs of NH groups in its structure, we decided to probe whether the nitration of this molecule would lead to the development of more potent energetic compounds (ultimately, to compounds **6** and **7**). There are many methods for the conversion of amines to nitramines: using nitrating agents such as HNO₃, mixtures of HNO₃ and H₂SO₄, acetic anhydride and HNO₃, nitrated silica gel and many others.²³ Thus, our initial efforts were focused on a direct nitration of compound **5** under various reaction conditions. More specifically, we evaluated a series of nitrating conditions and temperature regimes which included the use of red fuming HNO₃ and mixtures of either HNO₃ and H₂SO₄ (1:1 v/v), HNO₃ and acetic anhydride (1:1 v/v) or HNO₃ and trifluoroacetic anhydride (1:1 v/v). However, all examined reaction conditions (and all examined temperature regimes) led to one of two results: recovery of only starting compound **5** at the end of the reaction or decomposition of **5**.

We further attempted to establish whether any nitration of compound **5** actually takes place, with the formed nitramines hydrolysing back to the starting material upon dilution of the

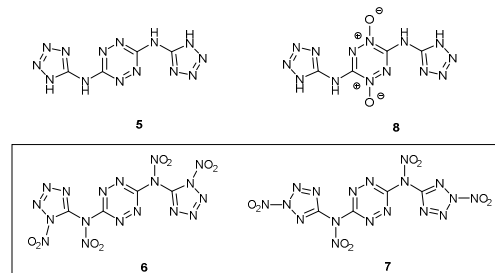


Fig. 3 Structures of compounds **5**, **6**, **7** and **8**.

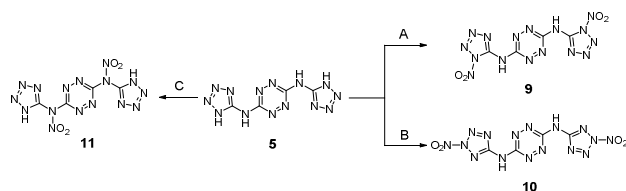


Fig. 4 Possible nitrated derivatives of compound **5**. Reaction conditions: $\text{Na}^{15}\text{NO}_3$, conc. H_2SO_4 .

purpose, we conducted *in situ* studies of the nitration of **5** by ^{15}N NMR using $\text{Na}^{15}\text{NO}_3$ in concentrated H_2SO_4 . A reference mixture of $\text{Na}^{15}\text{NO}_3$ in concentrated H_2SO_4 exhibited two ^{15}N NMR signals at 383 ppm and 248 ppm, indicating the presence of $^{15}\text{NO}_3^-$ and $^{15}\text{NO}_2^+$, respectively, and was consistent with previous reports.²⁴ Subsequent addition of BTATz to this nitration mixture at 0 °C resulted in the appearance of a new ^{15}N NMR signal at 336 ppm, which was assigned to the formation of a N- NO_2 adduct. Although it was obvious that, under explored conditions, no further nitration of **5** could be obtained, it was not clear which amine group in this compound underwent nitration (Fig. 4). Since the *in situ* ^{15}N NMR measurements did not help us determine whether endocyclic **9/10** (routes A and B, Fig. 4) or exocyclic **11** (route C, Fig. 4) nitramines were formed, and since the resulted product could not be isolated for further analysis due to its instability, other approaches were required. Therefore, we synthesized a model compound – N^3,N^6 -bis(2-methyl-2H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **12** – in which both endocyclic NH groups in the tetrazole rings were methylated (Fig. 5). The successful conversion of **5** to **12** was achieved by using dimethyl-sulfate in aqueous NaHCO_3 at room temperature. Comparison of ^1H and ^{13}C NMR data obtained for **12** to published reports for 2-methyl-2H-tetrazole derivatives (*versus* 1-methyl-1H-tetrazole derivatives) strongly suggested that the methylation of **5** took place on the second nitrogen atom of the tetrazole ring.

Further nitration of **12** with a mixture of concentrated HNO_3 and acetic anhydride (1:2 v/v) resulted in the formation of the new compound N,N' -(1,2,4,5-tetrazine-3,6-diyl)-bis(N -(2-methyl-2H-tetrazol-5-yl)-nitramide) **13** (Fig. 5). Unfortunately, nitramide **13** could not be fully characterized due to its very high sensitivity to impact and friction (*primary explosive!*). Also, based on ^1H NMR analysis in solution of DMSO-d_6 or CD_3CN , **13** underwent relatively quick hydrolysis back to the parent compound **12**. ^{13}C and ^{15}N NMR studies in DMSO-d_6 of the precipitate obtained in the nitration of **12** with a mixture of $\text{Na}^{15}\text{NO}_3/\text{HNO}_3$ (prepared separately) and acetic anhydride also showed only the presence of the starting material **12**. Yet, *in situ* ^{15}N NMR studies of the nitration of **12** with $\text{Na}^{15}\text{NO}_3$ in concentrated H_2SO_4 exhibited a new peak at 336 ppm (at the identical position of the nitramine's nitrate peak in compound **11**), strongly supporting our hypothesis that both compounds **5** and **12** undergo nitration on their exocyclic NH groups.

After realizing that the nitration of **5** (and its bis- N -methyl-tetrazole analog **12**) could not lead to the formation of stable nitramine products, we explored whether the nitration of the

unreported 1,2,4-triazole analog of compound **5** – N^3,N^6 -bis(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **15** would produce better results. The synthesis of compound **15** was achieved in an 81% yield, in a similar fashion to the synthesis of **5**, by reacting 3,6-bis(3,5-dimethyl-1H-pyrazol-1-yl)-1,2,4,5-tetrazine (BPT, **14**) with 1H-1,2,4-triazole-5-amine in sulfolane

at 135 °C (Fig. 6). The corresponding, more soluble, energetic perchlorate salt **16** and nitrate salt **17** were prepared by treating

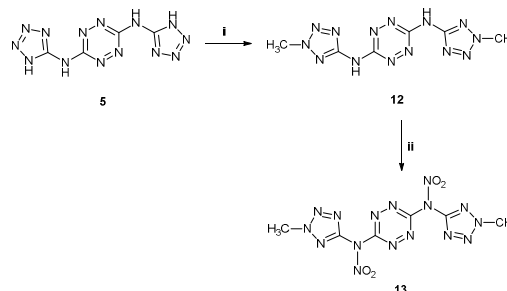


Fig. 5 Synthesis of bis- N -methylated compound **12**, with its subsequent nitration. Reaction conditions: (i) dimethylsulfate, NaHCO_3 , H_2O , r.t. (ii) conc. HNO_3 , Ac_2O .

15 with HClO_4 and HNO_3 , respectively (Fig. 6). The Structure of **16** was confirmed by X-ray crystallography (Fig. 9). Subsequently, a direct nitration of compound **15** was explored under various reaction conditions, which included either HNO_3 , a mixture of H_2SO_4 and HNO_3 (1:1; v/v), a mixture of $\text{CF}_3\text{CO}_2\text{H}$ and HNO_3 (1:1; v/v) or a mixture of NaNO_3 and H_2SO_4 . Under all tested temperature regimes, all aforementioned nitrating reagents led invariably to the formation of stable nitrate salt **17**.

In situ ^{15}N NMR studies of the nitration of **15** with $\text{Na}^{15}\text{NO}_3$ in concentrated H_2SO_4 (at 0 °C) displayed a peak at 336 ppm (at the identical position of the nitramine's nitrate peak in compound **13**), clearly indicating that the first nitration reaction takes place on the exocyclic NH groups of **15** and not on the CH or NH groups of its triazole rings (Fig. 7). This interesting observation demonstrates a similar nitration pattern for structurally related compounds **5**, **12** and **15**. We believe that similarly to **11** and **13**, attempts of isolation of compound **18** results in a quick hydrolysis of exocyclic nitramine group back to the original amine. In contrast to previous attempts, the nitration of **15** with a mixture of HNO_3 and acetic anhydride (1:1; v/v) at 0 °C resulted in the formation of a new nitramine – N,N' -(1,2,4,5-tetrazine-3,6-diyl)bis(N -(3-nitro-1H-1,2,4-triazol-5-yl)-nitramide) **19**, which was isolated by precipitation (Fig. 8). The latter compound could not be fully characterized due to its high sensitivity to impact and friction (*primary explosive!*). The

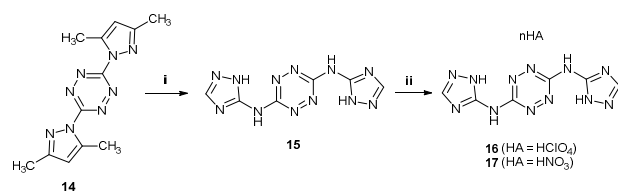


Fig. 6 Preparation of compound **15** and its perchlorate salt **16** and nitrate salt **17**. Reaction conditions: (i) 1*H*-1,2,4-triazole-5-amine, sulfolane, 135 °C; (ii) HClO₄ or HNO₃.

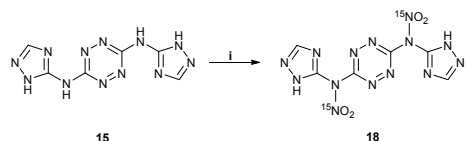


Fig. 7 Formation of the proposed compound **18**. Reaction conditions: (i) Na¹⁵NO₃, H₂SO₄.

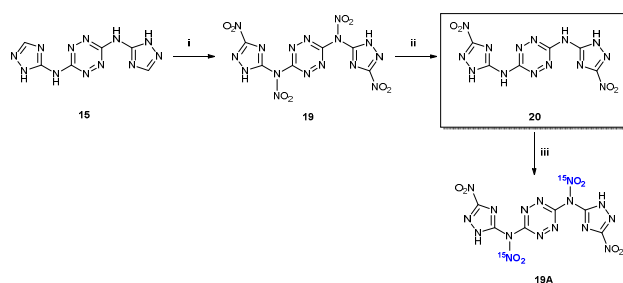


Fig. 8 Synthesis of compounds **19**, **20** and **19A**. Reaction conditions: (i) conc. HNO₃, Ac₂O; (ii) CH₃CN, 65 °C; (iii) Na¹⁵NO₃/HNO₃, Ac₂O.

solubility properties, sensitivity and thermal behaviour of **19** suggested its analogous structure to the exocyclic nitramine **13**. Controlled gradual heating of approximately 0.2 mg of wet **19** in a glass capillary (suitable for melting point measurements) resulted via its explosion at 110–112 °C, shuttering the capillary.

¹⁵N NMR studies of the precipitate **19** (dissolved in DMSO-*d*₆), obtained in nitration of **15** with a mixture of Na¹⁵NO₃/HNO₃ and acetic anhydride, showed the appearance of a new peak at 356.8 ppm (corresponding to C-¹⁵NO₂ nitrogen), indicating the formation of a new derivative with nitrated triazole rings *N*³,*N*⁶-bis(3-nitro-1*H*-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **20**. No exocyclic nitramine derivatives (N-¹⁵NO₂) of compound **15** were observed upon dissolution of **19** in DMSO-*d*₆, most reasonably due to the hydrolysis of **19** into the hydrolytically stable **20**. Also, the dissolution of **19** in hot CH₃CN (with further heating at 65 °C for 30 min) immediately after its separation from the nitration reaction mixture, resulted in the formation of a new compound – *N*³,*N*⁶-bis(3-nitro-1*H*-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **20** (Fig. 8). The structure of **20** was confirmed by X-ray crystallography (Fig. 10).

In order to check whether the structure of **19** was correct, *in situ* ¹⁵N NMR studies of the “back nitration” of **20** into exocyclic ¹⁵N nitramine **19A** were conducted. In these experiments, a solid **20** was slowly added to a mixture of

Na¹⁵NO₃ in concentrated H₂SO₄ at 0 °C and, after 30 min, the reaction mixture was analysed by ¹⁵N NMR. A new peak at 336.3 ppm (corresponding to the exocyclic N-¹⁵NO₂ nitrogen) appeared in the spectrum, indicating the formation of nitramine **19A** (Fig. 8), which perfectly matches our previous observation and conclusions.

Crystal Structures

X-ray measurements for single crystals of compounds **16** (perchlorate salt of compound **15**; CCDC 1041768) and **20** (CCDC 1041769) were performed. These data can be obtained free of charge from The Cambridge Crystallographic Data Centre via www.ccdc.cam.ac.uk/data_request/cif. The data for the crystals of **16** and **20** were collected using MoKα radiation (L = 0.71073 nm). An Oxford low-temperature device was used to keep the crystals at a constant temperature of 110K during the entire data collecting period. Details of the X-ray data collection and structure refinements are summarized in the Supporting Information. Detailed examination of the crystal structures obtained for **16** and **20** showed no significant differences in bond lengths and torsion angles reported for structures of other compounds containing 1,2,4-triazole and 1,2,4,5-tetrazine functional groups.²⁵ Both **16** and **20** molecules were found to be completely planar, due to an extensive delocalization of electrons in these molecules.

Compound **16** was crystallized as solvent-free crystals with the monoclinic space group P2(1)/c and a cell volume of 757.49 Å³. A crystal unit cell of **16** contains eight molecules of nitrogen-rich cations and four perchlorate anions. The measured density for **16** was found to be 1.960 g/cm³. The nitrogen-rich cation is protonated at nitrogen atom N6 in both triazole rings. Fig. 9 shows interactions between hydrogen atoms in compound **16** and oxygen atoms in perchlorate anions (each nitrogen-rich cation interacts with eight perchlorate anions). The hydrogen bonding parameters of these interactions are detailed in Table 1. The triazole moiety of **16** points towards nitrogen atom N6 and participates in an intramolecular hydrogen bond N1-H1...N4 with a D...A length of 2.732 Å and a D-H...A angle of 116°.

Compound **20** was crystallized as a monoclinic space group P2(1)/c and a cell volume of 1546.76 Å³ was calculated. Crystal unit cell of this compound contains eight molecules of **20** and ten molecules of DMF. The unit cell contains two additional disordered DMF molecules, which could not be reliably modelled by discrete atoms. Correspondingly, its contribution was subtracted from the diffraction pattern by the Squeeze

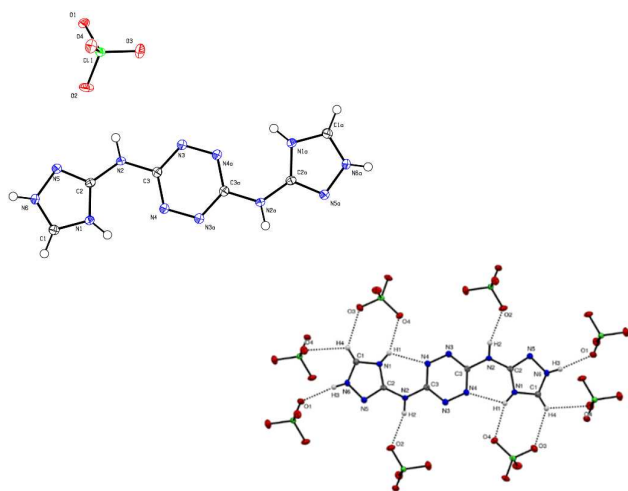


Fig. 9 (top): Molecular structure of compound **16**, (bottom): interactions between **16** and surrounding perchlorate anions.

Table 1 Hydrogen bonds present in crystal structure of **16**.

D-H...A	D-H/Å	H...A/Å	D...A/Å	D-H...A/°
N1-H1...O4	0.83	2.20	2.9669	153.7
N1-H1...O1	0.83	2.799	2.973	93.8
N2-H2...O2	0.87	2.10	2.9401	162.1
N6-H3...O1	0.80	2.06	2.8303	163
N1-H1...N4	0.83	2.265	2.732	116

technique, using the PLATON software.²⁶ As a result, the density measured was 1.35 g/cm³, which is significantly lower than 1.87 g/cm³ – the calculated density for this compound.²⁷ Fig. 10 shows interactions between nitrogen and hydrogen atoms of compound **20** and molecules of DMF (each molecule of **20** interacts with 10 molecules of DMF). The bond lengths of the bridge corresponds to a C-N single bond (C10-N9 1.359Å, N9-C7 1.370Å). The triazole moiety points towards atom N6 and participates in an intramolecular hydrogen bond – N6-H6...N12 – which has a D...A length of 2.7781Å and a D-H...A angle of 117° and an intermolecular hydrogen bond with the solvent (Table 2).

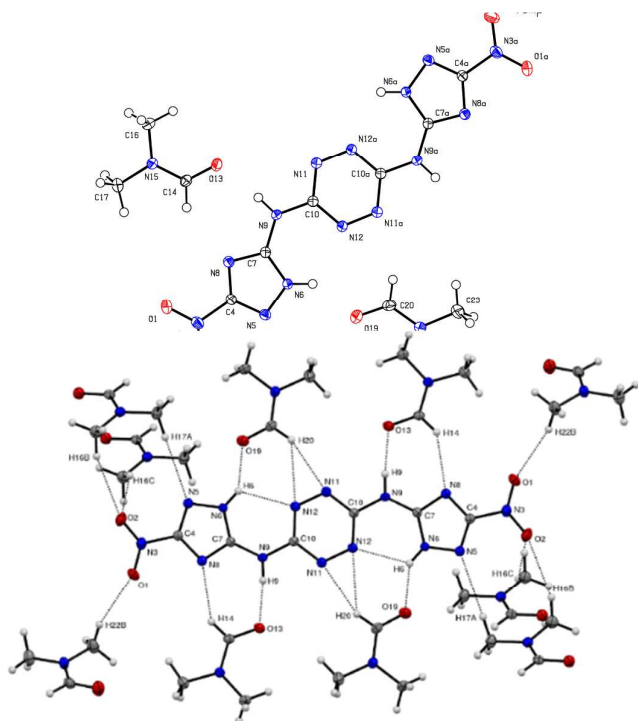


Fig. 10 (top): Molecular structure of compound **20**, (bottom): interactions between **20** and surrounding DMF molecules.

Table 2 Hydrogen bonds present in crystal structure of **20**.

D-H...A	D-H/Å	H...A/Å	D...A/Å	D-H...A/°
N6-H6...N12	0.88	2.27	2.7781	117
N6-H6...O19 ⁽ⁱ⁾	0.88	1.94	2.6939	143
N9-H9...O13 ⁽ⁱⁱ⁾	0.88	1.86	2.7333	175
C14-H14...N8 ⁽ⁱⁱⁱ⁾	0.95	2.43	3.2085	139
C16-H16A...O13	0.98	2.46	2.8005	100
C20-H20...N12 ^(iv)	0.95	2.58	3.2438	127
C20-H20...N11 ^(v)	0.95	2.43	3.3178	155=z

Symmetry codes: (i) 1-x, -1/2+y, 1/2-z (ii) 1+x, y, z (iii) -1+x, y, z (iv) 1-x, 1/2+y, 1/2-z (v) -1+x, 1/2-y, -1/2

Theoretical Calculations, Performance Characteristics and Thermal Stabilities

Calculations were performed using the Gaussian 09 software.²⁸ The geometric optimization of the structures and frequency analyses were carried out by using the B3LYP functional with the 6-311+G(d,p) basis set.²⁹ The optimized structures of all materials correspond to, at least, a local energy minimum on the potential energy surface. In order to predict energetic properties of new materials, the EXPLO5_v6.01³⁰ software was used. This software utilizes an algorithm based on the Becker-Kistiakowsky-Wilson equation of state (BKW EOS) for gaseous detonation products. Thermal stability is a very important property of energetic materials. The onset temperatures for thermal decomposition of compounds **5**, **12**, **15**, **16**, **17** and **20** were determined using differential scanning calorimeter (DSC). The results of these calculations and the experimental data are summarized in Table 3.

In contrast to compounds **15**, **16** and **17**, dinitro derivative **20** exhibited high thermal stability ($T_{\text{decomp.}}$ of 302°C), high detonation velocity (VOD of 8,903 m/s) and high detonation pressure, performing comparably to BTATz **5**. In addition, BAM measurements of dropping-hammer (7.67 Nm) and friction (> 353 N) for compound **20** indicate a relatively high stability of this energetic material under mechanical stress (for comparison, the equivalent values for RDX are 7.5 Nm and 120 N, respectively).

Conclusions

A methodical study of nitration patterns was conducted on a series of 3,6-diamino-1,2,4,5-tetrazine derivatives, revealing the specificity of the nitration in this type of nitrogen-rich compounds, as well as stability of the formed nitramines towards hydrolysis. The general motivation of this study was to evaluate whether energetic nitrogen-rich compounds, structurally-related to the N^3,N^6 -bis(1H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (BTATz; **5**), with improved oxygen balance characteristics, could be prepared. By ¹⁵N-labeling methodology, we found that although nitramine derivatives of **5**, as well as its dimethyl analogue N^3,N^6 -bis(2-methyl-2H-

tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (**12**), are formed in the reaction mixture, these nitramines could not be successfully isolated and quickly undergo hydrolysis back to the starting materials. Therefore, subsequent efforts were focused on more stable nitro derivatives of the newly synthesized compound N^3,N^6 -bis(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (**15**). The nitration of **15** was studied by ^{15}N NMR, showing a similar initial reactivity pattern of **5** and **12**. Yet, due to a presence of 1,2,4-triazole functional groups in

the structure of **15**, new C-NO₂ derivative N^3,N^6 -bis(3-nitro-1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (**20**) was obtained in a moderate yield. The structures of **16** (perchlorate salt of **15**) and **20** (with 4 molecules of DMF) were determined by x-ray crystallography. Importantly, it was found that compound **20** exhibited high stability towards friction (> 353 N) and very high thermal stability, decomposing at 302°C (by DSC). The standard

	5	6	7	9/10	11	12	13	15	16	17	18	19	20
Mw [g/mol]	248	428	428	338	338	276	366	246	447	372	336	426	336
%N [wt%] ^(a)	79	60	60	61.5	61.5	71	61	68	33.3	41.1	50	50	58
T _{Dec} [°C] ^(b)	309	-	-	-	-	306	-	350	246	167	N/A	-	302
ρ [g/cm ³] ^(c)	1.88	2.03	2.03	1.98	1.98	1.77	1.88	1.36*	1.96**	1.74*	1.87	1.95	1.87
Ω_{O_2} [%] ^(d)	-64.6	0	0	-23.6	-23.6	-92.6	-48	-97.5	-25	-43	-47.6	-18.7	-47.6
ΔH_f [kJ/mol] ^(e)	918	1148	1,334	113	964	1,022	996	831	-504	-655	965	864	1032
ΔU_f° [kJ/kg] ^(f)	3789	2756	3190	3436	2933	3801	2808	3468	-1056	-1667	2952	2103	3151
EXPLO5_v6.01 values:													
$-\Delta_e U^\circ$ [kJ/kg] ^(g)	4002	6,048	6400	5492	5032	4185	5097	3728	2947	1921	5212	5315	5398
T _E [K] ^(h)	2905	4632	4856	4101	3872	2763	3597	2805	2630	1873	3782	4091	3891
P _{C-J} [kbar] ⁽ⁱ⁾	301	443	456	415	394	265	333	124	251	161	329	390	339
V _{Det.} [m/s] ^(k)	8809	9790	9903	9604	9429	8492	8932	6240	711	6833	8830	9310	8903
Gas vol. [L/kg] ^(j)	758	759	762	794	789	743	778	734	751	792	753	777	753

(a) Nitrogen content; (b) measured by DSC start of decomposition temperature; (c) calculated density; (c*) density of dried powdered compounds was measured by gas pycnometry at 25 °C; (c**) density value that was obtained by a single-crystal x-ray diffraction; (d) oxygen balance (for C_aH_bN_cO_d $\Omega = (a - 2b - 0.5d) \times 1600 / \text{Mw}$); (e) calculated enthalpy of formation; (f) calculated energy of formation; (g) calculated heat of detonation; (h) calculated temperature of detonation; (i) calculated pressure of detonation; (k) calculated velocity of detonation; (j) calculated volume of gaseous products.

Table 3 Physico-chemical properties of compounds **5**, **6**, **7**, **9/10**, **11**, **12**, **13**, **15**, **16**, **17**, **18**, **19** and **20**.

heats of formation for all compounds in this study were calculated on the B3LYP 6311G+d,p level of theory, showing highly-positive values of 831.5 kJ/mol and 1032.9 kJ/mol for **15** and **20**, respectively. Detonation parameters for all compounds were calculated with the EXPLO5_v6.01 program. Theoretical density of 1.87 g/cm³ and velocity of detonation of 8,903 m/s were calculated for **20**, making it a highly promising new energetic compound.

Experimental Section

Caution! As certain triazole, tetrazole and tetrazine derivatives are unstable and can ignite or explode under the influence of impact, friction or heating, proper safety precautions should be taken in work with these materials. Lab personnel and equipment should be properly grounded and protective equipment including protective coat, Kevlar gloves, ear protection and face shield should be used.

General Information

All used chemical reagents and solvents were of analytical grade and were used as supplied, without further purification. ^1H , ^{13}C and ^{15}N NMR spectra were recorded on a 400 MHz NMR spectrometer at 25 °C. The chemical shifts are given relative to the residual DMSO-d₆ peaks or formamide (^{15}N) as external standards and coupling constants are given in Hertz (Hz). Infrared (IR) spectra were recorded on a FTIR spectrometer equipped with a diamond ATR unit. Mass spectra were recorded on a high-resolution ESI-Q-ToF machine. Elemental analyses (CHN) were performed at the service facility of the Hebrew University. Decomposition points were determined by differential scanning calorimetry (DSC). Measurements were performed at a heating rate of 5 °C/min in closed aluminium sample pans with a small hole in the lid under a nitrogen flow of 20 mL/min using an empty aluminium sample pan as a reference. Melting points were measured on a melting point apparatus in open glass capillaries. The impact sensitivities were tested according to STANAG 4489 using a BAM drop-hammer. The friction sensitivities were tested

according to STANAG 4487 using a BAM friction tester. Experimental densities were obtained by pycnometry measurements at ambient temperature.

General Procedures

N^3,N^6 -bis(2-methyl-2H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (12). A solid N^3,N^6 -di(1H-tetrazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (816 mg, 3.29 mmol) was added to a solution of NaHCO_3 (690 mg, 8.21 mmol) in H_2O (30 mL) and stirred at RT for 30 min, until dissolution. To the resulted solution, dimethylsulfate (1.04 g, 8.22 mmol) was added and the reaction mixture was stirred at RT for 3 days. After that time, the precipitate formed was collected by filtration, washed with a mixture of triethylamine in MeOH (1:20 v/v; 3×10 mL) and dried under vacuum to yield pure **12** (536 mg; 59%) as a red solid. DSC (5 °C/min) 306 °C (decomp.). ^1H NMR (400 MHz, DMSO-d_6): δ 4.32 (s, 6H). ^{13}C NMR (100 MHz, DMSO-d_6): δ 40.2, 159.0, 160.6. ^{13}C DEPT135 (100 MHz, DMSO-d_6): δ 40.2 (CH₃). HRMS (ESI^+): m/z = 277.1127 [$\text{M}+\text{H}^+$]. Elemental analysis: calcd. (%) for $\text{C}_6\text{H}_8\text{N}_{14}$: C 26.09, H 2.96; found: C 26.29, H 2.86. FTIR (ATR): ν 3235 (w), 3018 (w), 2865 (w), 1588 (s), 1538 (s), 1428 (s), 1262 (m), 1197 (m), 1049 (s), 950 (s), 898 (w), 815 (w), 744 (m), 672 (m), 547 (s) cm^{-1} .

N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (15). A solid 1H-1,2,4-triazol-5-amine (4.23 g, 50.31 mmol) was added to a solution of 3,6-bis(3,5-dimethyl-1H-pyrazol-1-yl)-1,2,4,5-tetrazine **14** (6.46 g, 23.90 mmol) in sulfolane (150 mL) and the reaction mixture was heated at 135 °C for 24 hr. After that time, the reaction mixture was cooled down to RT, DMF (150 mL) was added and the mixture was stirred for 1 hr. The precipitate formed was collected by filtration, washed with MeOH (3×80 mL) and dried under vacuum to yield **15** (4.77 g; 81%) as an orange solid. DSC (5 °C/min) 350 °C (decomp.). HRMS (ESI^-): m/z : 245.0697 [$\text{M}-\text{H}^-$]. Elemental Analysis: calcd. (%) for $\text{C}_6\text{H}_6\text{N}_{12}$: C 29.27, N 68.27, H 2.46; found: C 29.92, N 68.73, H 2.55. FTIR (ATR): ν 2807 (m), 2359 (w), 1597 (s), 1542 (s), 1486 (s), 1434 (s), 1312 (m), 1268 (s), 1207 (w), 1062 (s), 1047 (s), 1027 (s), 962 (s), 896 (s), 763 (m), 719 (s), 649 (m), 557 (s). Due to very low solubility of **15** in all tested organic solvents, the free base was converted into the corresponding nitrate and perchlorate salts for further analysis.

N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine perchlorate salt (16). A solid N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **15** (500 mg, 2.03 mmol) was added to aqueous HClO_4 (5 mL, 70%) and the mixture was stirred at RT for 15 min. To the resulted mixture, CH_3CN (5 mL) was added and the reaction mixture was stirred at RT for an additional 10 min. After that time, the precipitate formed was collected by filtration, washed with CH_3CN (3×3 mL) and dried under vacuum to yield pure **16** (653 mg; 72%) as an orange solid. DSC (5 °C/min) 246 °C (decomp.). ^1H NMR (400 MHz, DMSO-d_6): δ 8.56 (s), 9.49 (br s). ^{13}C NMR (100 MHz, DMSO-d_6): δ 144.7, 151.0, 158.6. FTIR (ATR): ν 2761 (w), 1602 (s), 1552 (s), 1499 (s), 1435 (s), 1312 (s), 1268 (s), 1207 (m), 1062 (s), 964 (s), 854 (m), 740 (s), 620 (s), 557 (s).

N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine nitrate salt (17). A solid N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **15** (500 mg, 2.03 mmol) was added to HNO_3 (70%; 5 mL) and the mixture was stirred at RT for 15 min. To the resulted solution, CH_3CN (10 mL) was added and the reaction mixture was stirred at RT for 15 min. After that time, the precipitate formed was collected by filtration, washed with CH_3CN (3×3 mL) and dried under vacuum to yield pure **17** (557 mg; 74%) as an orange solid. DSC (5 °C/min) 167 °C (decomp.). ^1H NMR (400 MHz, DMSO-d_6): δ 8.65 (s), 11.35 (br s). ^{13}C NMR (100 MHz, DMSO-d_6): δ 143.4, 150.1, 158.6. FTIR (ATR): ν 2943 (w), 1697 (s), 1621 (s), 1551 (s), 1407 (s), 1303 (s), 1060 (s), 1041 (s), 999 (s), 937 (s), 855 (s), 713 (s), 557 (s).

N^3,N^6 -bis(3-nitro-1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine (20). HNO_3 (99%; 2.5 mL, 60.02 mmol) was added to acetic anhydride (7.5 mL, 79.49 mmol) at 0 °C and the mixture was stirred for 30 min. To this mixture, a solid N^3,N^6 -di(1H-1,2,4-triazol-5-yl)-1,2,4,5-tetrazine-3,6-diamine **15** (494 mg, 2.0 mmol) was slowly added at 0 °C and the reaction mixture was stirred at this temperature for 2 hr under anhydrous conditions. After that time, the reaction mixture was allowed to warm up to RT and the orange precipitate formed (*very sensitive explosive!*) was collected by filtration, quickly washed with $\text{CF}_3\text{CO}_2\text{H}$ (2×2 mL) and immediately re-dissolved in hot CH_3CN (30 mL). The resulted CH_3CN solution was heated at 65 °C for 30 min and then cooled to RT. The precipitate formed was collected by filtration, washed with CH_3CN (3×5 mL) and dried under vacuum to yield pure **20** (211 mg; 31%) as an orange solid. DSC (5 °C/min) 302 °C (decomp.). ^{13}C NMR (100 MHz, DMSO-d_6): δ 149.7, 158.2, 160.5. HRMS (ESI^-): m/z : 335.0492 [$\text{M}-\text{H}^-$]. Elemental Analysis: calcd. (%) for $\text{C}_6\text{H}_4\text{N}_{14}\text{O}_4$: C 21.44, N 58.33, H 1.20; found: C 21.75, N 57.56, H 1.30. FTIR (ATR): ν 3247 (w), 2996 (w), 1601(s), 1541 (s), 1494 (s), 1435 (s), 1417 (s), 1312 (s), 1296 (s), 1151 (m), 1072 (s), 1050 (s), 1018 (m), 950 (s), 891 (m), 835 (s), 748 (m), 692 (s), 570 (s).

Supporting Information: Contains copies of ^1H and ^{13}C NMR spectra for isolated compounds and ^{15}N NMR spectra for ^{15}N -labeled compounds formed *in situ* in the nitration experiments, and crystallographic data for compounds **16** and **20**.

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