

International Scientific Organization http://iscientific.org/ Chemistry International www.bosaljournals.com/chemint/



# Synthesis of different classes of five/six membered heterocyclic cyanine dyes: A review

## **Hassan Abazied Shindy**

Department of Chemistry, Faculty of Science, Aswan University, Aswan 81528, Egypt Corresponding author's E. mail: hashindy2@hotmail.com

#### ARTICLE INFO

Article type: Review article Article history: Received January 2019 Accepted July 2019 April 2020 Issue Keywords: Cyanine dyes Synthesis Heterocyclic dyes Classes of cyanine dyes Uses of cyanine dyes Applications of cyanine dyes

### ABSTRACT

In this paper, synthesis of different classes of five/six membered heterocyclic cyanine dyes has been reviewed. In this paper review detailed synthesis steps were represented via equations. The synthesis covers, monomethine cyanine dyes (simple cyanine dyes), dimethine cyanine dyes, trimethine cyanine dyes (carbocyanine dyes), styryl cyanine dyes (hemicyanine dyes), aza-styryl cyanine dyes (aza-hemicyanine dyes and/or aza-cyanine dyes), merocyanine dyes (acyclic merocyanine dyes and cyclic merocyanine dyes) and apocyanine dyes. Besides, in the introduction section of this review paper some light is focused on the uses, applications and properties of cyanine dyes. This review paper is informative, useful and very readable for synthetic dye chemists, researchers and students who look for the different methods in the synthesis and preparation of various classes of five/six membered heterocyclic cyanine dyes with special emphasize in the field of heterocyclic and/or cyanine dyes chemistry. This specific type of collective review in the synthesis of different classes of only five/six membered heterocyclic cyanine dyes has been paid little attention and has great importance in the chemistry literature.

© 2020 International Scientific Organization: All rights reserved.

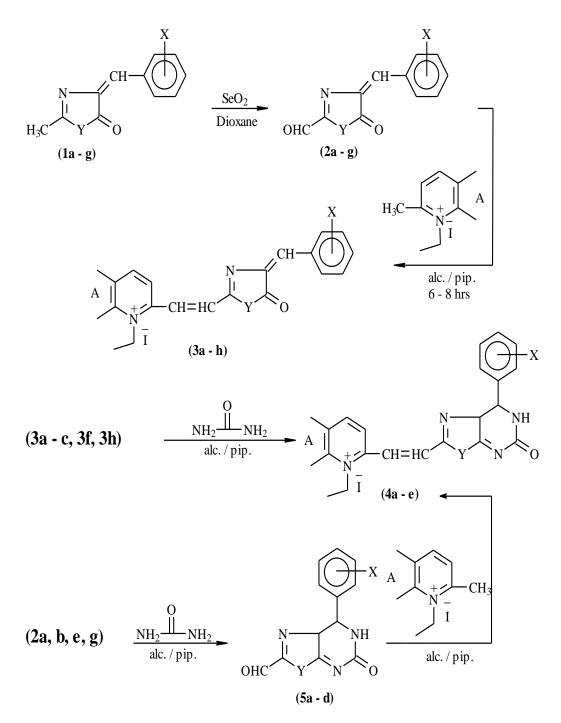
**Capsule Summary:** Synthesis of different classes of five/six membered heterocyclic cyanine dyes have been reviewed.

**Cite This Article As:** H. A. Shindy. Synthesis of different classes of five/six membered heterocyclic cyanine dyes: A review. Chemistry International 6(2) (2020) 56-74. https://doi.org/10.5281/zenodo.3361022

#### INTRODUCTION

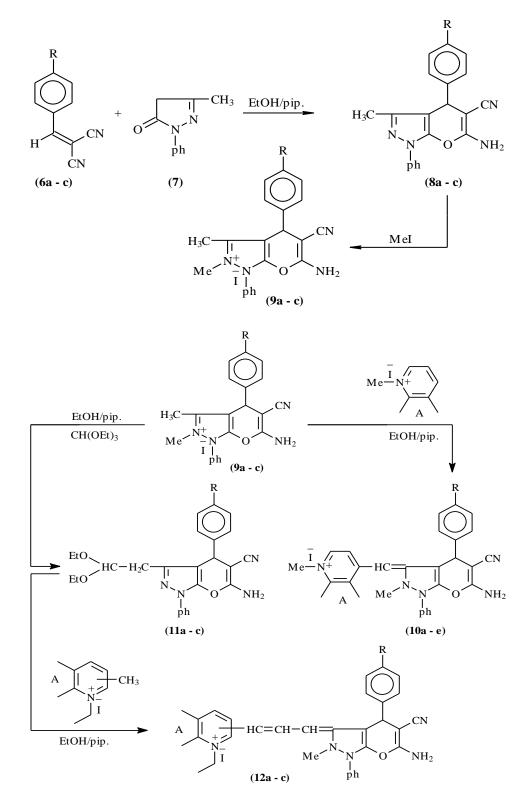
Much and much attention and have been paid to the chemistry of cyanine dyes (Shindy, 2014; Shindy, 2015; shindy, 2015a; Shindy, 2012; Yagupolski et al., 2008; Volkova et al., 2007; Xu et al., 2007; Yashchuk et al., 2007; Boto et al., 2008; Gadjev et al., 1999; Fasiulla et al., 2008; Deligeorgiev et al., 1998; Shindy, 2016; Shindy, 2017; Kovalska et al., 2010; Kabatc et al., 2010; Yakubovskyi et al., 2010; Klochko et al., 2010; Matsuoka, 1990). This is because the extraordinarily uses and

applications of these dyes in a broad and diverse area, such biotechnology, as biology, biochemistry, physics, engineering, pharmacology and medicine. In more than one century's time (In 1873) Vogel discovered that dye had spectrum sensitization added function (Hamer, 1964). From this time, the synthesis, development and application of the spectrum sensitization additive dyes (mainly refers to cyanine dyes) have made the important contribution to the photosensitize material industry. Now, the spectrum sensitization additive dyes was still one essential photography organic matter of kinds new variety photosensitive material.



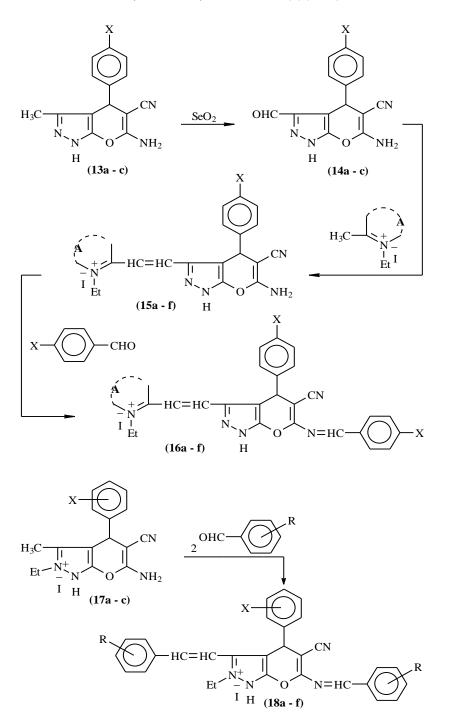
Scheme 1: Synthesis of dimethine cyanine dyes and their cyclocondensation reactions with urea.

Substituents in Scheme 1: {(1a-g): (2a-g): Y = 0; X = H(a),  $\rho$ .OCH<sub>3</sub> (b),  $\rho$ .NO<sub>2</sub> (c),  $\rho$ .NO<sub>2</sub> (d), m.NO<sub>2</sub> (e),  $\rho$ .OH (f); Y = NH; X = H (g). (3a-h): Y = 0; X = H, A = H-2yl salt (a); X = H, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (b); X =  $\rho$ .OCH<sub>3</sub>, A = C<sub>6</sub>H<sub>4</sub>-2yl salt(c); X =  $\rho$ .NO<sub>2</sub>, A-C<sub>6</sub>H<sub>4</sub>-2-yl salt(d); X =  $\rho$ .NO<sub>2</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (e); X = m.NO<sub>2</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (f); X =  $\rho$ .OH, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (g); Y = NH; X = H, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (h). (5a-d): Y=0; X = H (a),  $\rho$ .OCH<sub>3</sub> (b), m.NO<sub>2</sub> (c); Y = NH; X = H (d). (4a-e): Y = 0; X = H, A = H-2yl salt (a), X = H, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (b), X =  $\rho$ .OCH<sub>3</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (c), X = m.NO<sub>2</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (d); Y = NH; X = H (d). (4a-e): Y = 0; X = H, A = H-2yl salt (a), X = H, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (b), X =  $\rho$ .OCH<sub>3</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (c), X = m.NO<sub>2</sub>, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (d); Y = NH, X = H, A = C<sub>6</sub>H<sub>4</sub>-2-yl salt (e)}



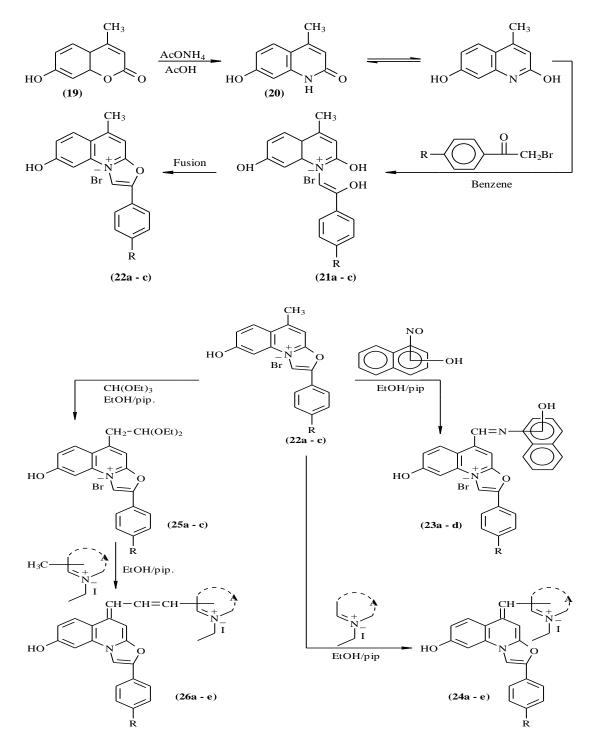
**Scheme 2:** Synthesis of monomethine and trimethine cyanine dyes.

Substituents in Scheme 2: (6a-c); (8a-c); (9a-c); (11a-c): R = H (a),  $OCH_3$  (b),  $NO_2$  (c). (10a-e): R = H, A = H-4-yl (a); R = H,  $A = C_6H_4-4-yl$  (b); R = H,  $A = C_6H_4-1-yl$  (c);  $R = OCH_3$ ,  $A = C_6H_4-4-yl$  (d);  $R = NO_2$ ,  $A = C_6H_4-4-yl$  (e). (12a-e): R = H, A = H-2yl (a); R = H,  $A = C_6H_4-2-yl$  (b); R = H, A = H-4-yl (c);  $R = OCH_3$ ,  $A = C_6H_4-2-yl$  (d);  $R = NO_2$ ,  $A = C_6H_4-2-yl$  (e). (12a-e): R = H, A = H-2yl (a); R = H,  $A = C_6H_4-2-yl$  (b); R = H, A = H-4-yl (c);  $R = OCH_3$ ,  $A = C_6H_4-2-yl$  (d);  $R = NO_2$ ,  $A = C_6H_4-2-yl$  (e).



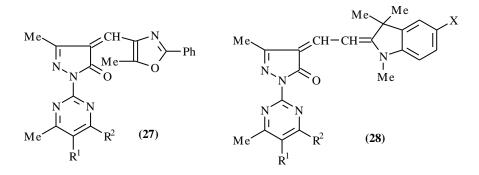
**Scheme 3:** Synthesis of dimethine, dimethine/aza-styryl mixed and styry/aza-styryl mixed cyanine dyes.

Substituents in Scheme 3: (13a - c), (14a - c), (17a - c): X =  $\rho$ .Cl (a),  $\rho$ .NO<sub>2</sub> (b),  $\rho$ .N(CH<sub>3</sub>)<sub>2</sub> (c). (15a - f): X =  $\rho$ .Cl, A = 1-ethyl quinolinium-2-yl salt (a); X =  $\rho$ .Cl, A = 1-ethyl pyridinium-2-yl salt (b); X =  $\rho$ .N(CH<sub>3</sub>)<sub>2</sub>, A = 1-ethyl quinolinium-2-yl salt (c); X =  $\rho$ .N(CH<sub>3</sub>)<sub>2</sub>, A = 1-ethyl pyridinium-2-yl salt (d); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl quinolinium-2-yl salt (e); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl pyridinium-2-yl salt (f); (16a - f): X =  $\rho$ .Cl, A = 1-ethyl quinolinium-2-yl salt (a); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl quinolinium-2-yl salt (b); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl quinolinium-2-yl salt (b); X =  $\rho$ .OH, A = 1-ethyl quinolinium-2-yl salt (c); X =  $\rho$ .OH, m.OCH<sub>3</sub>, A = 1-ethyl quinolinium-2-yl salt (d); X =  $\rho$ .OH, A = 1-ethyl quinolinium-2-yl salt (c); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl pyridinium-2-yl salt (f). (18a - f): X =  $\rho$ .OCH<sub>3</sub>, A = 1-ethyl quinolinium-2-yl salt (e); X =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl pyridinium-2-yl salt (f). (18a - f): X =  $\rho$ .NO<sub>2</sub> (a); X =  $\rho$ .Cl, R =  $\rho$ .N(CH<sub>3</sub>)<sub>2</sub> (b); X =  $\rho$ .NO<sub>2</sub>, R =  $\rho$ .N(CH<sub>3</sub>)<sub>2</sub> (d); X =  $\rho$ .NO<sub>2</sub> (e); X =  $\rho$ .NO<sub>2</sub> (e); X =  $\rho$ .NO<sub>2</sub> (f).

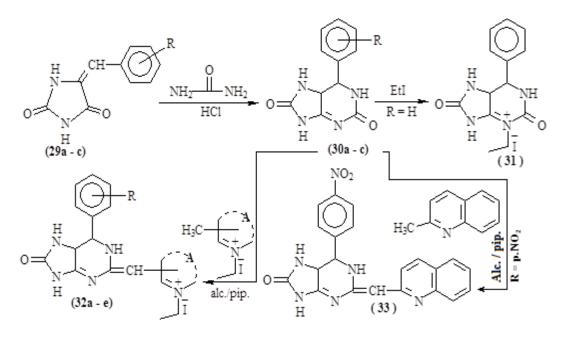


Scheme 4: Synthesis of monomethine, trimethine and aza-styryl cyanine dyes.

Substituents in Scheme 4: (21a-c), (22a-c), (25a-c): R = H (a);  $OCH_3$  (b);  $NO_2$  (c). (23a-d): R = H,  $\alpha$ -nitroso- $\beta$ -2327naphthol (a);  $R = OCH_3$ ,  $\alpha$ -nitroso- $\beta$ -naphthol (b);  $R = NO_2$ ,  $\alpha$ -nitroso- $\beta$ -naphthol (c);  $R = OCH_3$ ,  $\beta$ -nitroso- $\alpha$ -naphthol (d); (24a-e): R = H, A = N-ethyl quinolin-4-ium salt (a);  $R = OCH_3$ , A = N-ethyl pyridin-4-ium salt (b);  $R = OCH_3$ , A = N-ethyl quinolin-4-ium salt (c);  $R = OCH_3$ , A = N-ethyl pyridin-4-ium salt (d);  $R = NO_2$ , A = N-ethyl quinolin-4-ium salt (e). (26a-e): R = H, A = 1-ethyl quinolin-2-ium salt (a);  $R = OCH_3$ , A = 1-ethyl pyridin-2-ium salt (b);  $R = OCH_3$ , A = 1-ethyl quinolin-4-ium salt (c);  $R = OCH_3$ , A = 27 1-ethyl pyridin-4-ium salt (d);  $R = NO_2$ , A = 1-ethyl quinolin-2-ium salt (e).



Scheme 5: Cyclic merocyanine dyes useful for dyeing polyester fibers orange shades.

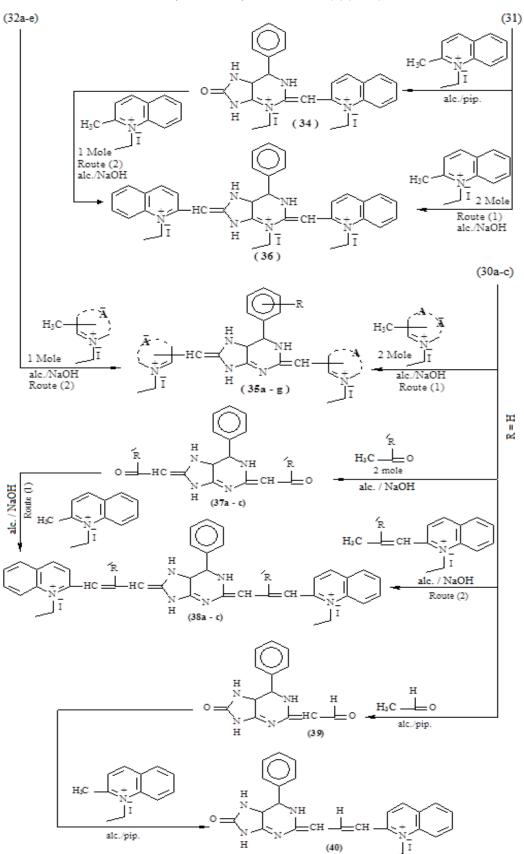


Scheme 6: Monomethine, bismonomethine, monomethine base, trimethine and bistrimethine cyanine dyes synthesis.

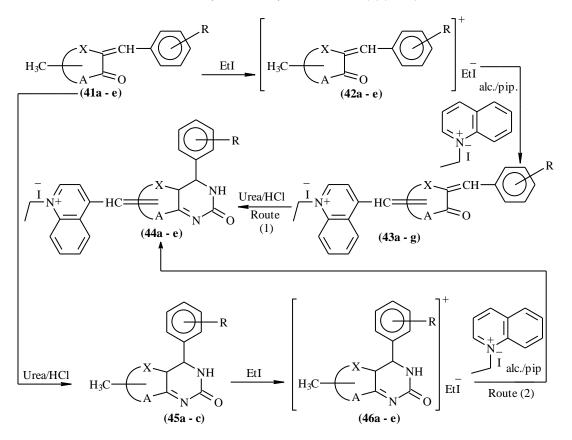
Substituents in Scheme 6: (29a - c); (30a - c): R = H (a),  $\rho$ .OCH<sub>3</sub> (b),  $\rho$ .NO<sub>2</sub> (c). (32a - e): R = H, A = 1-ethyl pyridinium-2yl salt (a); R = H, A = 1-ethyl quinolinium-2-yl salt (b); R = H, A = 1-ethyl pyridinium-4-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A = 1-ethyl quinolinium-2-yl salt (d); R =  $\rho$ .NO<sub>2</sub>, A = 1-ethyl quinolinium-2-yl salt (e). (35a - g): R = H, A =  $\overline{A}$  = 1-ethyl-pyridinium-2-yl salt (a); R = H, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (b); R = H, A =  $\overline{A}$  = 1-ethyl quinolinium-4-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-4-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\rho$ .OCH<sub>3</sub>, A =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R =  $\overline{A}$  = 1-ethyl quinolinium-2-yl salt (c); R = H, A(\overline{A}) = 1-ethyl quinolinium-2-yl salt (1-ethyl pyridinium-2-yl salt) (f); R = H, A(\overline{A}) = 1-ethyl quinolinium-2-yl salt (1-ethyl pyridinium-4-yl salt) (g). (37a - c), (38a - c): R = H (a), CH<sub>3</sub> (b), C<sub>6</sub>H<sub>5</sub> (c).

Therefore, it continuously received the attention in the research of the synthesis and applications both in domestic and in a broad. In addition, cyanine dyes have been used on new techniques in molecular imaging because of their usefulness in biological, medical and clinical research. Fluorescence imaging methods are generally superior in terms of sensitivity, selectivity and ease of use. Cyanine dyes have been employed as fluorescent labels in fluorescence imaging studies of biological mechanism, as

important signaling molecule involved in the regulation of a wide range of physiological and pathophysiological mechanisms and many disorders. Besides, polymethine cyanine dyes have many technology application areas such as forgery prevention, photoresists, spectrally sensitized photographic materials, thermal transfer printing and heat shielding materials. A number of publications published recently are also highlighting the importance of cyanine dyes.



Scheme 6: Continue...



Scheme 7: Synthesis of monomethine cyanine dyes.

Substituents in Scheme 7: (41a - e); (42a - e); (44a - e); (45a - e); (46a - e): X(A): N-ethyl-1-H-pyrazoline, R = H (a),  $\rho$ .OCH<sub>3</sub> (b),  $\rho$ .NO<sub>2</sub> (c), N-ethyl-1-phenyl-pyrazoline, R =  $\rho$ .OCH<sub>3</sub> (d); N-ethyl-oxazoline, R = H (e). (43a - g): X(A): N-ethyl-1-phenyl pyrazoline, R = H (a),  $\rho$ .OCH<sub>3</sub> (b),  $\rho$ .NO<sub>2</sub> (c); o.OH (d),  $\rho$ .OH (e); N-ethyl-1-H-pyrazoline, R = H (f); N-ethyl oxazoline, R = H (g).

The synthesis, properties and applications of cyanine dyes reflect the important and vital position of these dyes in the chemistry of dyes and pigments (Shindy et al., 2015; Soriano et al., 2016; Shindy 2017a; Shindy et al., 2017; Shindy 2016a; Shindy et al., 2016; Shindy et al., 2016a; Shindy et al., 2016b; Shindy et al., 2016c; Shindy et al., 2017a; Komljenovic et al., 2016; Zhang et al., 2016; Chen et al., 2016; Arjona et al., 2016; Shindy 2018a; Shindy et al., 2018; Shindy et al., 2018; Shindy et al., 2019; Shindy et al., 2019; Shindy et al., 2020).

# Synthesis of different classes of five/six membered heterocyclic cyanine dyes

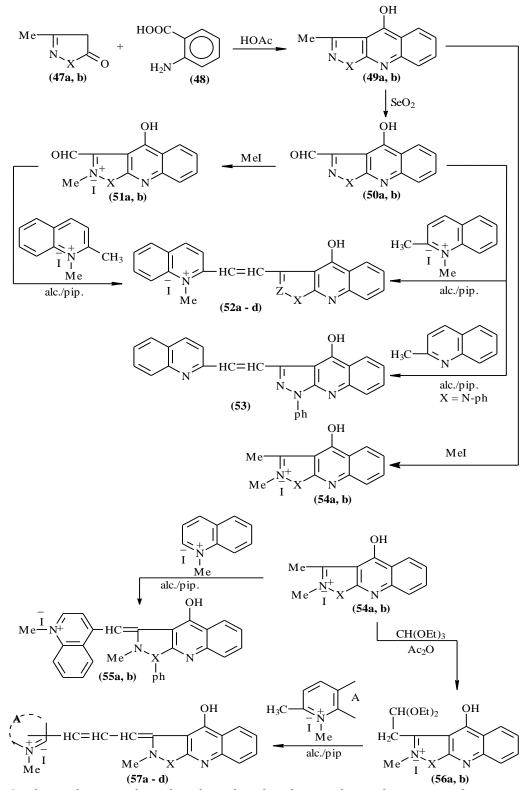
Dimethine cyanine dyes were prepared by the condensation of 4-benzylideno-oxazol- / imidazol-5-one-2-carboxaldehyde derivatives with 2-methyl pyridinium (quinolinium)-2-yl salts. Cyclocondensation reaction of these dyes with urea gave new asymmetrical oxazolo(imidazo)-[4,5-d]-pyrimidine-2(2)-dimethine cyanines (Koraiem, et al., 1991), Scheme (1).

Derivatives of 6-amino-5-cyano-3-methyl-1,4-diarylpyrano[2,3-c]pyrazole were prepared via the reaction of  $\alpha$ - $\beta$ -unsaturated nitriles with 3-methyl-1-phenyl-pyrazole-5one as starting materials in the synthesis of new asymmetrical mono-(tri-)-methine cyanine dyes (Abd El Aal et al., 1998), Scheme (2).

A number of photosensitizers asymmetrical dimethine and styryl cyanine dyes having pyrazolo pyran nucleus were synthesized (Hassan et al., 1991), Scheme (3).

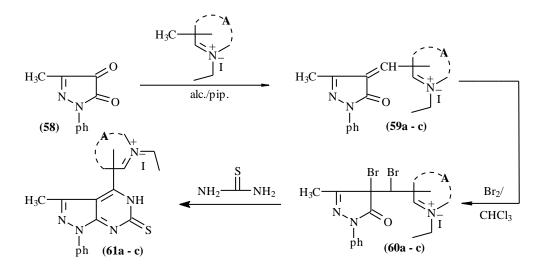
Oxazolo[1,2-a]quinoline derivatives were used to synthesis novel bridgehead cyanine dyes such as azamethine, monomethine and trimethine cyanines (Abd El Aal, 1999), Scheme (4).

A series of cyclic merocyanine dyes (27), (28) ( $R^1$  = H or lower alkyl, X = H, halo, OH, alkoxyl, alkyl, NO<sub>2</sub>,  $R^2$  = OH, alkoxyl, Cl), useful for dyeing polyester fibres orange shades, are manufactured by reaction of the corresponding 3-methyl-1-(6-methyl-2-pyrimidinyl)-5-pyrazolines with 5-methyl-2-phenyl-oxazole-4-carboxaldehyde or aldehydes (Ioan et al., 1998), Scheme (5).



Scheme 8: Synthesis of monomethine, dimethine, dimethine base, and trimethine cyanine dyes.

Substituents in Scheme 8: (47a, b); (49a, b); (50a, b); (51a, b); (55a, b); (55a, b); (56a, b): X = 0 (a); N-ph (b). (52a – d): X = 0, Z = N (a); X = N-ph, Z = N (b); X = 0,  $Z = N^+$ -MeĪ (c); X = N-ph,  $Z = N^+$ -MeĪ (d). (57a – d): X = 0, A = 1-methyl quinolinium-2-yl salt (a); X = N-ph, A = 1-methyl pyridinium-2-yl salt (b); X = N-ph, A = 1-methyl pyridinium-2-yl salt (c); X = N-ph, A = 1-methyl pyridinium-2-yl salt (d).



Scheme 9: Synthesis of apocyanine dyes (zero methine cyanine dyes).

Substituents in Scheme 9: (59a – c), (60a-c), (61a-c) : :A = 1-ethyl pyridinium-2-yl salt (a); 1-ethyl quinolinium-2-yl salt (b); 3-ethyl benzoxazolium-2-yl salt (c).

4-Aryl-1,2,3,4,5,6-hexahydroimidazo [4,5-d]pyrimidine-2,6diones were prepared and used to obtain (with quinaldine or picoline ethiodides) 6-monomethine, 2,6-bis monomethine, 6-trimethine and 2,6-bis trimethine cyanine dyes (Koraiem et al., 1998), Scheme (6).

A number of photosensitizers monomethine cyanine dyes incorporating pyrazolino and/or oxazolino nuclei were synthesized (Koraiem et al., 1990), Scheme (7). Different types of 2(3)-oxazolino(pyrazolino)-[4,3-d] quinoline-2(4)-methine cyanine dves including monomethine, dimethine and trimethine cyanines were prepared (El-Hamd et al., 1994), Scheme (8). 4,5-dioxo-3methyl-1-phenyl-pyrazoline is condensed with  $\alpha$ -picoline-EtI, quinaldine-EtI or 2-methyl benzoxazole-EtI to form monomethine derivatives, which is then brominated and finally cyclocondensed with hydrazines, hydroxylamine or with thiourea to give apocyanine dyes (Koraiem, 1984), Scheme (9).

Monomethine, B-substituted dimethine and styryl cyanine dyes were synthesized using oxonium salts (Abd El Aal, et al., 2005), Scheme (10).

A series of dimethine cyanine dyes, acyclic merocyanine dyes and cyclic merocyanine dyes containing pyrazine and/or oxazine nucleus were synthesized (Abu El hamd and Koraiem, 1990), Scheme (11). A series of different monomethine cyanine dyes containing thiazole and/or pyridine nucleus were prepared (Koraiem et al., 1995), Scheme (12).

Dimethine cyanine dyes (Koraiem et al., 1996) containing thiazole and/or pyridine nucleus were synthesized (Scheme 13). A series of monomethine cyanine dyes and trimethine cyanine dyes having pyrazolo/ oxadiazine nucleus were prepared (Eissa, 2009) (Scheme 14).

#### CONCLUSIONS

The structures of any cyanine dyes are characterized by the presence of at least two essential centers. The first one is the basic center which have the responsibility of pushing electrons across the conjugated chromophoric group system of the dyes molecules, and the second one is the acidic center which have the responsibility of pulling electrons across the conjugated chromophoric group system of the dyes structures, and vice versa, Scheme (15). These push-pull system in the dyes structures have the reason and/ or the responsibility for the intensity of the colour of cyanine dyes, where they leads to the formation of two mesomeric electronic transitions structures (two resonance forms) (A) and (B) inside the dyes molecules producing a delocalized positive charge over the conjugated chromophoric group system of the dyes structures, Scheme (15).

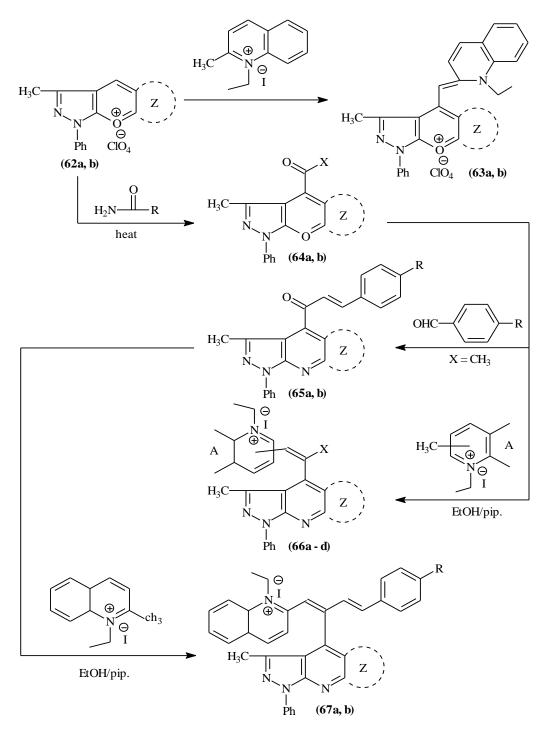
The synthesis and/or the preparation methods of cyanine dyes usually carried out through using a number of active heterocyclic quaternary salts residue, Scheme (16).

The most common active heterocyclic quaternary salts used in the preparation of cyanine dyes (Scheme16) are as follows:

a-N-iodomethane and/or N-iodoethane quaternary salts of 2–picoline, quinaldine, 4–picoline and/or lepidine,

b- N-iodomethane and/or N-iodoethane quaternary salts of 2-methyl thiazole, 2-methyl bcnzothiazole, 2-methyl oxazole, 2-methyl benzoxazole, 2-methyl selenazole, 2methyl benzoselenazole, 2-methyl imidazole and/or 2methyl benzoimidazole.

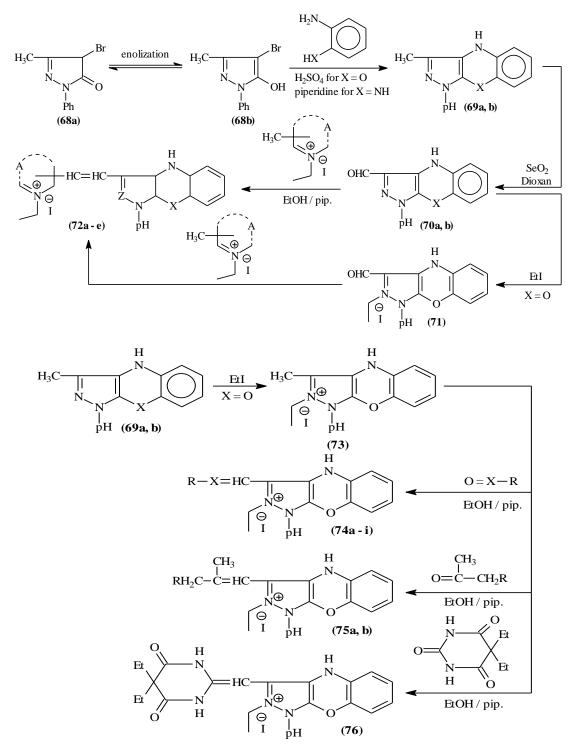
iscientic.org.



**Scheme 10:** Synthesis of monomethine, styryl and dimethine cyanine dyes.

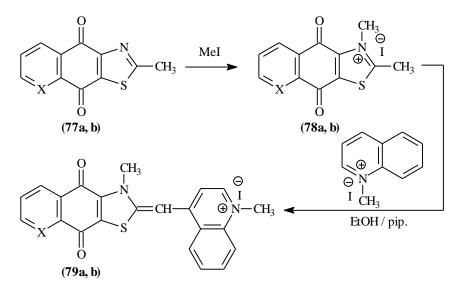
Substituents in Scheme 10: (62a, b); (63a, b): Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene (a); cyclohexene (b). (64a, b): Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene,  $X = CH_3$  (a); Z = cyclohexene,  $X = CH_3$  (b). (65a, b): Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene, R = OH (a); Z = cyclohexene, R = OH (b). (66a-d): Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene,  $X = CH_3$ , A = 1-ethylpyridine-2-ium (a); Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene,  $X = CH_3$ , A = 1-ethylpyridine-2-ium (b); Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene,  $X = CH_3$ , A = 1-ethylpyridine-2-ium (b); Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene,  $X = CH_3$ , A = 1-ethylpyridine-4-ium (c); Z = 5-cyclohexene,  $X = CH_3$ , A = 1-ethylpunolin-2-ium (d). (67a, b): Z = 1-methyl-4(2-propenyl)cyclohexen-l,5-diene, R = OH (a); Z = cyclohexene, R = OH (b).

#### Shindy / Chemistry International 6(2) (2020) 56-74

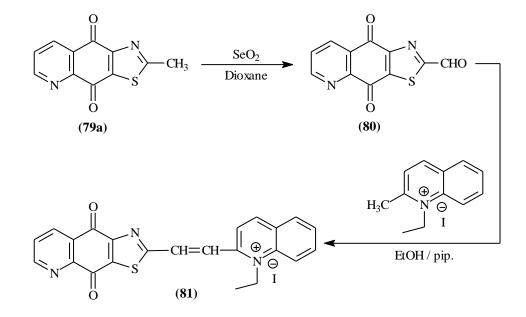


**Scheme 11:** Synthesis of dimethine, acyclic merocyanine and cyclic merocyanine dyes.

Substituents in Scheme 11: (69a, b); (70a, b): X = O (a); NH (b). (72a – e): X = O, Z = N, A = 1-ethyl pyridinium-2yl salt (a); X = O, Z = N, A = 1-ethyl quinolinium-2yl salt (b); X = O, Z = N, A = 1-ethyl pyridinium-4yl salt (c); X = NH, Z = N, A = 1-ethyl quinolinium-2yl salt (d); X = O,  $Z = N^+$ -Etl-, A = 1-ethyl quinolinium-2yl salt (e). (74a – i):  $X = CH_3$ ,  $R = C_6H_5$  (a); X = CH,  $R = C_6H_4$ -p.OCH<sub>3</sub> (b); X = CH,  $R = C_6H_4$ -p.OH (c); X = CH,  $R = C_6H_4$ -p.OH<sub>3</sub> (d); X = CH,  $R = C_6H_4$ -o.OH (e); X = CH,  $R = C_6H_3$ -p.OH. m.OCH<sub>3</sub> (f); X = CH,  $R = C_4H_3O$  (g); X = N,  $R = C_6H_4$ -p.OH (h); X = N,  $R = C_6H_4$ -p.N(CH<sub>3</sub>)<sub>2</sub> (i). (75a, b):  $R = COCH_3$  (a); COOEt (b).



# Scheme 12: Synthesis of different monomethine cyanine dyes.Substituents in Scheme 12: (77a, b); (78a, b); (79a, b): X = N (a);



Scheme 13: Synthesis of dimethine cyanine dyes.

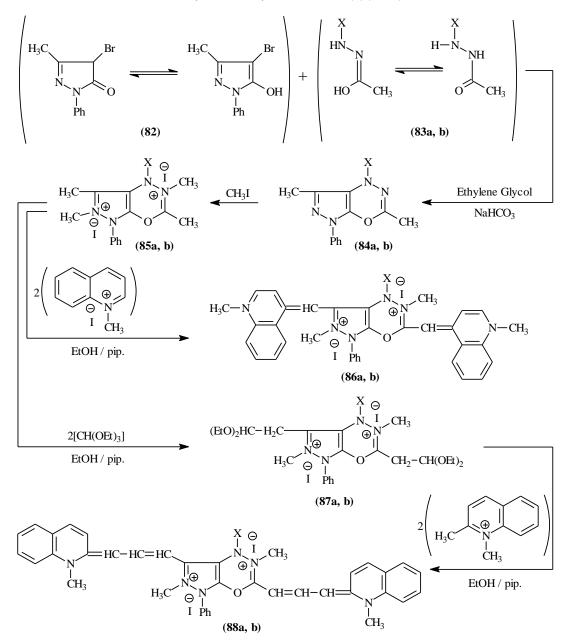
c- N-iodomethane and/or N-iodoethane quaternary salts of pyridine, quinoline and/or isoquinoline,

d- N-iodomethane and/or N-iodoethane quaternary salts of thiazole, bcnzothiazole, oxazole, benzoxazole, selenazole, benzoselenazole, imidazole and/or benzoimidazole.

4-The quaternary salts in a and b characterized by the presence of active (acidic) methyl group at 2 and/or 4 positions, while the quaternary salts in c and d characterized by the presence of active (acidic) hydrogen atom in 4, 1 and/or 2 positions, Scheme (16).

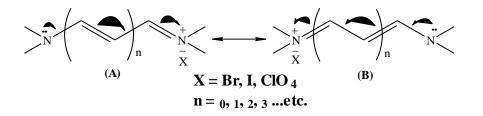
5-The activity (acidity) of the 2 (4)-methyl group in the quaternary salts of 2-picoline, quinaldine, 4-picoline and/or lepidine is related to the presence of the highly electron attracting positive charge on the quaternary nitrogen atom at position 1 of these compounds, Scheme (16).

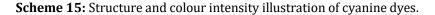
iscientic.org.



**Scheme 14:** Synthesis of monomethine and trimethine cyanine dyes.

Substituents in Scheme 14: (83a, b); (84a, b); (85a, b); (86a, b); (87a, b); (88a, b): X = H (a); Ph (b).



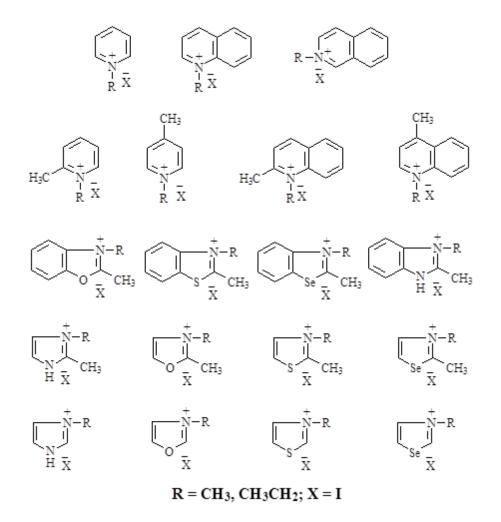


6-The activity (acidity) of the 2-methyl group in the quaternary salts of 2-methyl thiazole, 2-methyl oxazole, 2-methyl selenazole and/or 2-methyl imidazole is related to the presence of the highly electron attracting positive charge on the quaternary nitrogen atom at position 3 of these compounds in addition to the electron pulling characters of the hetero atom present, Scheme (16)..

7-The activity (acidity) of the 2-methyl group in the quaternary salts of 2-methyl bcnzothiazole, 2-methyl benzoxazole, 2-methyl benzoselenazole and/or 2-methyl imidazole is related to the presence of the highly electron attracting positive charge on the quaternary nitrogen atom at position 3 of these compounds in addition to the electron pulling characters of the hetero atom present and the electron attracting character of the condensed benzene ring system, Scheme (16).

8-The activity (acidity) of the 4 (1)-hydrogen atom in the quaternary salts of pyridine, quinoline and/or isoquinoline is related to the presence of positive charge on the quaternary nitrogen atom at positions 1 and/or 2 of these compounds, Scheme (16). 9-The activity (acidity) of the 2-hydrogen atom in the quaternary salts of thiazole, oxazole, selenazole and/or imidazole is related to the presence of the highly electron attracting positive charge on the quaternary nitrogen atom at position 3 of these compounds in addition to the electron pulling characters of the hetero atom present, Scheme (16).

10- The activity (acidity) of the 2-hydrogen atom in the quaternary salts of benzothiazole, benzoxazole, benzoselenazole and/or benzoimidazole is related to the presence of the highly electron attracting positive charge on the quaternary nitrogen atom at position 3 of these compounds in addition to the electron pulling characters of the hetero atom present and the electron attracting character of the condensed benzene ring system (Scheme 16). Cyanine dyes also might have enhanced biological properties, which need to be tested evaluated (Abate et al., 2019; Ayare et al., 2019; Karcı et al., 2013; Manjunatha et al., 2019; Mishra et al., 2019; Negm et al., 2016; Rizk et al., 2015; Rizk et al., 2017; Sadeghi-Kiakhani et al., 2019; Şener et al., 2018; Silva et al., 2018; Sun et al., 2019; Zhang et al., 2019).



Scheme 16: Most common active heterocyclic quaternary salts residue used in the preparation of cyanine dyes.

#### **FUTURE PROSPECTS**

The current and the future research developments aim to provide novel synthetic methods for the preparation of different classes of highly antimicrobial (antibacterial and anti-fungi) active, anti-tumor, anti-cancer, pH sensitive, highly photographic sensitizers, non-toxic, high stability, light fastness, near IR (Infrared), fluorescent, anticorrosion, strong labeled DNA and extra conjugated cyanine dyes. Such as oxadiazine cyanine dyes, thiazole cyanine dyes, metal stabilized cyanine dyes, pentamethine cyanine dyes, undecamethine cyanine dyes and tridecamethine cyanine dyes.

Also, the current and/or the future research developments aimed to provide new, novel and/or patent review papers in the field of color, dyes and pigments chemistry. The aimed review papers will covers and/or includes topics like the origin of color, the relation between color and constitutions, synthesis of dyes, properties of dyes, classification of dyes, uses and/or applications of dyes. Also, additional important topics for the current and/or the future research developments for the aimed review papers will includes methine cyanine dyes, hemicyanine dyes (styryl cyanine dyes), merocyanine dyes, apocyanine dves, monoheterocyclic cyanine dves, biheterocyclic cyanine dyes, polyheterocyclic cyanine dyes, six membered heterocyclic cyanine dyes, five/six membered heterocyclic cyanine dyes, five membered heterocyclic cyanine dyes and benz(naphth)/five membered heterocyclic cyanine dyes.

A very bright future for cyanine dyes chemistry can be expected through joint efforts (collaboration) of a large heterogenous community groups composed of synthetic dyes chemists, biologists, physicists, biotechnologists, pharmacologists, technological engineerists and medical scientists.

#### ACKNOWLEDGEMENT

I am thankful to the Chemistry department, Faculty of Science, Aswan University, Aswan, Egypt for supporting this work.

#### REFERENCES

- Abate, M.T., Ferri, A., Guan, J., Chen, G., Ferreira, J.A., Nierstrasz, V., 2019. Single-step disperse dyeing and antimicrobial functionalization of polyester fabric with chitosan and derivative in supercritical carbon dioxide. The Journal of Supercritical Fluids 147, 231-240.
- Abd E1-Aa1, R.M., 1999. Synthesis and characterization of new photosensitizer bridgehead cyanine dyes. Proc. Journal of Chemical Sciences (Indian Academy of Sciences) 111(2), 343-352.

- Abd E1Aal, R.M., Abd El-Latif, F.M., Koraiem, A.I.M., 1998. Proc. Synthesis and solvatochromic behaviour of new asymmetrical mono-(tri)-methine cyanine dyes. Journal of Chemical Sciences (Indian Academy of Sciences) 110 (4), 433-444.
- Abd El-Aal, R.M., Koraiem, A.I.M., Khalil, Z.H., El-Kody, A. M. M., 2005. The use of oxonium salts in the synthesis of mono-,  $\beta$ -substituted dimethine and styryl cyanine dyes. Dyes and Pigments 66, 201-209.
- Abu El-Hamd, R.M., Koraiem, A.I.M., 1990. Studies on The Synthesis of Conjucted Five-Six Biheterocyclic Cyanine Dye Series. Journal of Islamic Academy of Sciences 3(4), 262-268.
- Arjona, A., Stolte, M., F. Wilerthner, F., 2016. Conformational switching of  $\pi$  -conjugated junctions from merocyanine to cyanine statesby solvent polarity. Angewandte Chemie, 55(7), 2470-2473.
- Ayare, N.N., Ramugade, S.H., Sekar, N., 2019. Photostable coumarin containing azo dyes with multifunctional property. Dyes and pigments 163, 692-699.
- Boto, R.E.F., Santos, P.F., Reis, L.V., Almeida, P., 2008. Synthesis and characterization of new mono- and dicarboxyalkylselenacarbocyanines. Dyes and Pigments 76, 165-172.
- Chen, H., Sun, H., Zhang, X., Sun, X., Shi, Y., Xu, S., Tang, Y., 2016. A colorimetric and fluorometric dual-modal DNA logic gate based on the response of a cyanine dye supramolecule to G-quadruplexes. New Journal of Chemistry 40, 1940-1943.
- Deligeorgiev, T.G., Zaneva, D.A., Kim, S.H., Sabnis, R.W., 1998. Preparation of monomethine cyanine dyes for nucleic acid detection. Dyes and Pigments 37(3), 205-211.
- Eissa, F.M., 2009. Preparation, antibacterial activity and absorption spectra of pyrazolo-oxadiazine cyanine dyes. Journal of the Chinese Chemical Society 56, 843-849.
- El-Hamd, R.A., Koraiem, A., Abd El-Aal, R., El-Bahnasawy, A.A., 1994. Synthesis of asymmetrical-2(3)oxazolino(pyrazolino)-2(4)-methine cyanine dyes. Zagazig Journal of Pharmaceutical Science 3(2), 13-22.
- Fasiulla, M.H., Khan, M., Harish, M.N.K., Keshavayya, J., Reddy, K.R.V., 2008. Synthesis, spectral, magnetic and antifungal studies on symmetrically substituted metal(II)octaiminophthalocyanine pigments. Dyes and Pigments 76, 557-563.
- Gadjev, N.I., Deligeorgiev, T.G., Kim, S.H., 1999. Preparation of monomethine cyanine dyes as noncovalent labels for nucleic acids. Dyes and Pigments 40, 181-186.
- Hamer, F.M., 1964 The cyanine dyes and related compounds, New York, Interscience publishers Inc., p. 198.

- Hassan, M.E., El-Maghraby, M.A., Khalafalla, A.K., Khalil, M.A.,1991. Asymmetrical dimethine and styryl cyanines from pyrazolo-pyran moiety. Journal für praktische Chemie 333 (2), 361-368.
- Kabatc, J., Paczkowski, J., 2010. Monomeric asymmetric two and three- cationic monomethine cyanine dyes as novel photoinitiations for free radical polymerization. Dyes and Pigments 86, 133-142.
- Karcı, F., Karcı, F., Demirçalı, A., Yamaç, M., 2013. Synthesis, solvatochromic properties and antimicrobial activities of some novel pyridone-based disperse disazo dyes. Journal of Molecular Liquids 187, 302-308.
- Klochko, O.P., Fedyunyayeva, I.A., Khabuseva, S.U., Semenova,
  O.M., Terpetschnig, E.A., Patsenker, L.D., 2010.
  Benzodipyrrolenine-based biscyanine dyes: Synthesis,
  molecular structure and spectroscopic characterization.
  Dyes and Pigments 85, 7-15.
- Komljenovic, D., Wiessler, M., Waldeck, W., Ehemann, V., Pipkorn, R., Shrenk, H., Debus, J., Braun, K., 2016. NIRcyanine dye linker: a promising candidate for isochronic fluorescence imaging in molecular cancer diagnostics and therapy monitoring. Theranostics 6(1), 131-141.
- Koraiem, A., Girgis, M., Khalil, Z., Abu El-Hamd, R., 1990. Electronic absorption spectral studies on new synthesised monomethine cyanine dyes. Indian Journal of Technology 28, 580-586.
- Koraiem, A.I.M., 1984. Apocyanine dyes from 4,5-dioxo-3methyl-1-phenyl-pyrazoline. Journal für praktische Chemie 326(5), 811-816.
- Koraiem, A.I.M., Abu El-Hamd, R.M., Shindy, H.A., 1995. Studies on some new heterocyclic quinone monomethine cyanine dyes. Chemical paper 49(4), 192-197.
- Koraiem, A.I.M., Abu El-Hamd, R.M., Khalafalah, A.K., Hammam, A.S., El-Maghraby, M.A., Gomaa, M.M., 1996. Synthesis and properties of some naphtho (quinolino)quinone heterocyclic dimethine cyanine dyes. Dyes and Pigments 30(2), 89-98.
- Koraiem, A.I.M., El-Maghraby, M.A., Khalafallah, A.K., Shindy, H.A., 1998. Studies on The synthesis and solvatochromic behaviour of mono- and tri methine cyanines: Methine cyanine dyes, synthesis and solvatochromism. Dyes and Pigments 11, 47-63.
- Koraiem, A.I.M., Girgis, M.M., Khalil, Z.H., Abu El-Hamd, R. M.,1991. Electronic absorption spectral studies on new dimethine cyanine dyes. Dyes and Pigments 15(2), 89-105.
- Kovalska, V. B., Volkova, K. D., Manaev, A. V., Losytskky, M. Y., Okhrimenko, I. N., Traven, V. F., Yarmoluk, S. M., 2010. 2-Quinolone and Coumarin polymethines for the detection

of proteins using fluorescence. Dyes and Pigments 34, 159-164.

- loan, P., Ioan, C., Valeria, S., Viorica, A., Rodica, C., 1998. Rom.
  Ro 106, 266 (Cl.C09B23/02), 31 Mar 1993, Appl. 147,736, 10 Jun 1991; 8 pp. (Rom). Chemical Abstract, 128, 295825h.
- Manjunatha, C.R., Nagabhushana, B.M., Raghu, M.S., Pratibha, S., Dhananjaya, N., Narayana, A., 2019. Perovskite lanthanum aluminate nanoparticles applications in antimicrobial activity, adsorptive removal of Direct Blue 53 dye and fluoride. Materials Science and Engineering: C 101, 674-685.
- Matsuoka, M., 1990. (Ed.), Infrared Absorbing dyes, Plenum Press, New York, p. 220.
- Mishra, V.R., Ghanavatkar, C.W., Sekar, N., 2019. UV protective heterocyclic disperse azo dyes: Spectral properties, dyeing, potent antibacterial activity on dyed fabric and comparative computational study. Spectrochimica Acta Part A: Molecular and Biomolecular Spectroscopy 223, 117353.
- Negm, N.A., Abou Kana, M.T.H., Abd-Elaal, A.A., Elwahy, A.H.M., 2016. Fluorescein dye derivatives and their nanohybrids: Synthesis, characterization and antimicrobial activity. Journal of Photochemistry and Photobiology B: Biology 162, 421-433.
- Rizk, H.F., Ibrahim, S.A., El-Borai, M.A., 2015. Synthesis, fastness properties, color assessment and antimicrobial activity of some azo reactive dyes having pyrazole moiety. Dyes and Pigments 112, 86-92.
- Rizk, H.F., Ibrahim, S.A., El-Borai, M.A., 2017. Synthesis, dyeing performance on polyester fiber and antimicrobial studies of some novel pyrazolotriazine and pyrazolyl pyrazolone azo dyes. Arabian Journal of Chemistry 10, S3303-S3309.
- Sadeghi-Kiakhani, M., Tehrani-Bagha, A.R., Gharanjig, K., Hashemi, E., 2019. Use of pomegranate peels and walnut green husks as the green antimicrobial agents to reduce the consumption of inorganic nanoparticles on wool yarns. Journal of Cleaner Production 231, 1463-1473.
- Şener, N., Mohammed, H.J.A., Yerlikaya, S., Celik Altunoglu, Y., Gür, M., Baloglu, M.C., Şener, İ., 2018. Anticancer, antimicrobial, and DNA protection analysis of novel 2,4dihydroxyquinoline dyes. Dyes and Pigments 157, 11-19.
- Shindy, H. A., 2015. Different methods in the synthesis of mono(bi)-heterocyclic six membered cyanine dyes: A Review. Organic Chemistry: An Indian Journal 11(1), 26-36.
- Shindy, H. A., El-Maghraby, M. A., Goma, M. M. and Harb, N. A.,
  2018a. Synthesis, electronic transitions and antimicrobial activity evaluation of novel monomethine

ISSN: 2410-9649

and trimethine cyanine dyes. European Journal of Molecular Biotechnology 6(2), 83-95.

- Shindy, H.A., 2012. Synthesis of different classes of five/five membered heterocyclic cyanine dyes: A Review paper. Mini-Reviews in Organic Chemistry 9(2), 209-222.
- Shindy, H.A., 2014. Synthesis of different classes of six membered heterocyclic cyanine dyes: A Review. Revue Roumaine de Chimie 59(2), 117-12.
- Shindy, H.A., 2015a. Synthesis of different classes of benz (naphth) / five membered heterocyclic cyanine dyes: A review. Revue Roumaine de Chimie 60(1), 5-13.
- Shindy, H.A., 2016. Basics in colors, dyes and pigments chemistry: A review. Chemistry International 2(1), 29-36.
- Shindy, H.A., 2016a. Characterization, mechanisms and applications in the chemistry of cyanine dyes: A review. European Journal of Molecular Biotechnology 14(4), 158-170.
- Shindy, H.A., 2017. Problems and solutions in colors, dyes and pigments chemistry: A review. Chemistry International 3(2), 97-105.
- Shindy, H.A., 2017a. Fundamentals in the chemistry of cyanine dyes: A review. Dyes and Pigments 145, 505-513.
- Shindy, H.A., 2018. Novel polyheterocyclic cyanine dyes: synthesis, photosensitiztion and solvent/electronic transitions correlation. European Reviews of Chemical Research 5 (1), 30-39.
- Shindy, H.A., 2018a. Structure and solvent effects on the electronic transitions of some novel furo/pyrazole cyanine dyes. Dyes and Pigments 149, 783-788.
- Shindy, H.A., El-Maghraby, M. A., Goma, M. M. and Harb, N. A., 2018. Synthesis and visible spectra studies of novel pyrazolo/oxazole merocyanine dyes. European Reviews of Chemical Research 5 (2), 75-85.
- Shindy, H.A., El-Maghraby, M.A., Goma, M.M., Harb, N.A., 2019. Novel styryl and aza-styryl cyanine dyes: Synthesis and spectral sensitization evaluation. Chemistry International 5, 117-125.
- Shindy, H.A., El-Maghraby, M.A., Goma, M.M., Harb, N.A., 2020. Dicarbocyanine and tricarbocyanine dyes: Novel synthetic approaches, photosensitization evaluation and antimicrobial screening. Chemistry International 6, 30-41.
- Shindy, H.A., Goma, M.M., Harb, N.A., 2015. Synthesis, Spectral Behavior and Biological Activity of some Novel 1,3,4-Oxadiazine Cyanine Dyes. European Journal of Chemistry 6(2), 151-156.
- Shindy, H.A., Goma, M.M., Harb, N.A., 2016a. Novel carbocyanine and bis carbocyanine dyes: synthesis,

visible spectra studies, solvatochromism and halochromism, ,Chemistry International 2(4), 222-231.

- Shindy, H.A., Khalafalla, A.K., Goma, M.M., Eed, A.H., 2016. Synthesis, photosensitization and antimicrobial activity evaluation of some novel Merocyanine dyes. Chemistry International 2 (3), 114-120.
- Shindy, H.A., Khalafalla, A.K., Goma, M.M., Eed, A.H., 2016b. Polyheterocyclic compound in the synthesis and spectral studies of some novel methine cyanine dyes., Revue Roumaine de Chimie 61(3), 139-145.
- Shindy, H.A., Khalafalla, A.K., Goma, M.M., Eed, A.H., 2016c. Novel hemicyanine and aza-hemicyanine dyes: synthesis, spectral investigation and antimicrobial evaluation. European Journal of Molecular Biotechnology 13(3), 94-103.
- Shindy, H.A., Khalafalla, A.K., Goma, M.M., Eed, A.H., 2017. Synthesis and studies on some new dimethine and tetramethine cyanine dyes. European Reviews of Chemical Research 4(2), 52-65.
- Shindy, H.A., Khalafalla, A.K., Goma, M.M., Eed, A.H., 2017a. Synthesis, spectral sensitization, solvatochromic and halochromic evaluation of new monomethine and trimethine cyanine dyes. European Journal of Molecular Biotechnology 5(1), 30-42.
- Silva, M.G.d., Barros, M.A.S.D.d., Almeida, R.T.R.d., Pilau, E.J., Pinto, E., Soares, G., Santos, J.G., 2018. Cleaner production of antimicrobial and anti-UV cotton materials through dyeing with eucalyptus leaves extract. Journal of Cleaner Production 199, 807-816.
- Soriano, E., Holder, C., Levitz, A., Henary, M., 2016. Benz [c,d] indolium-containing monomethine cyanine dyes: synthesis and photophysical properties. Molecules 21(1), 23.
- Sun, J., Wang, H., Zheng, C., Wang, G., 2019. Synthesis of some surfactant-type acid dyes and their low-temperature dyeing properties on wool fiber. Journal of Cleaner Production 218, 284-293.
- Volkova, K.D., Kovalska, V.B., Balanda, A.O., Losytskyy, M.Y., Golub, A.G., Subramaniam, R.J., Tolmachev, O.I., Yarmoluk, S.M., 2008. Specific fluorescent detection of fibrillar α-synuclein using mono- and trimethine cyanine dyes. Bioorganic and Medicinal Chemistry 16(3), 1452-1459.
- Volkova, K.D., Kovalska, V.B., Balanda, A.O., Vermeij, R.J., Subramaniam, V., Slominski, Y., Yarmoluk, S.M., 2007.
  Cyanine dye-protein interactions: Looking for fluorescent probes for amyloid structures. Journal of Biochemical and Biophysical Methods 70(5), 727-733.
- Xu, C., Yu, M., Losytskyy, V.B., Kovalska, D.V., Kryvorotenko, S. M., Yarmoluk, S., Bianco, P.R., 2007. Novel monomeric

cyanine dyes as reporters for DNA helicase activity. Journal of Fluorescence 17(6), 671- 685.

- Yagupolski, L.M., Kondratenko, N.V., Chernega, O.N., Chernega, A.N., Buth, S.A., Yagupolskil, Y.L., 2008. A novel synthetic route to thiacyanine dyes containing a perfluorinated polymethine chain. Dyes and Pigments 79 (3), 242-246.
- Yakubovskyi, V.P., Shandura, M.P., Kovtum, Y.P., 2010. Boradipyrromethenecyanines derived from conformationally restricted nuclei. Dyes and Pigments 87, 17-21.
- Yashchuk, V.M., Gusak, V.V., Dmytruk, I.M., Prokopets, V.M., Kudrya, V.Y., Losytskyy, M.Y., Tokar, V.P., Gumenyuk, Y.O., Yarmoluk, S.M., Kovalska, V.B., Kryvorotenko, D.V., 2007. Two-photon excited luminescent styryl dyes as probes for the DNA detection and imaging. photostability and phototoxic influence on DNA. Molecular Crystals and Liquid Crystals 467 (1), 325-338.
- Zhang, G., Li, X., Xu, X., Tang, K., Vu, V.H., Gao, P., Chen, H., Xiong, Y.L., Sun, Q., 2019. Antimicrobial activities of irradiation-degraded chitosan fragments. Food Bioscience 29, 94-101.
- Zhang, H., Niesen, B., Hack, E., Jenatasch, S., Wang, L., Veron, A.
  C., Makha, M., Schneider, R., Arroyo, Y., Hany, R., Nuesch,
  F., 2016. Cyanine tandem and triple-junction solar cells.
  Organic Electronics 30, 191-199.

Visit us at: http://bosaljournals.com/chemint/ Submissions are accepted at: editorci@bosaljournals.com