

Review in Effect of Catalysis in Any Organic Reaction

*Dr. Nagham Mahmood Aljamali, **Hussein Ali Ahmed

* Professor, Ph.D., Organic Chemistry, Synthetic Chemistry Field, Iraq

** Assist. Lecturer, Dept of Chemistry, College of Education for Pure Science, University of Kerbala, Iraq.

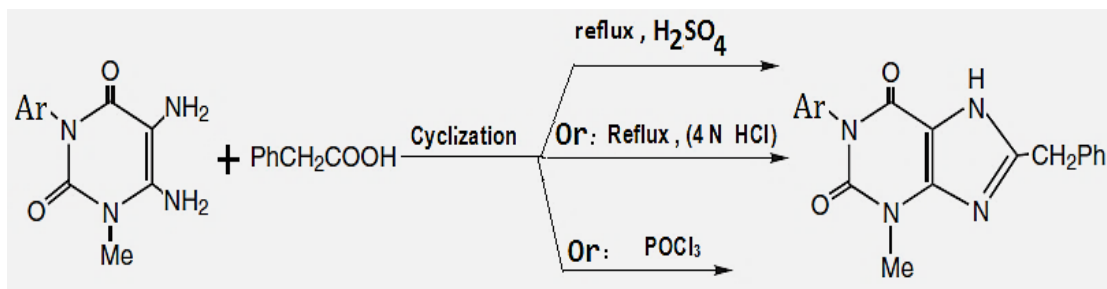
ABSTRACT

This review involved effect of catalysts on any chemical reaction, any reaction proceeds because the reaction products are more stable than the reactants and starting materials. The un catalyzed reaction is slow. In fact, simplest example: the decomposition of hydrogen peroxide is so slow that hydrogen peroxide solutions are commercially available. This reaction is strongly affected via catalysts like [manganese dioxide](#), or the enzyme [peroxidase](#) in organisms, reduction and oxidation reactions.

Keywords: catalysts, catalysis, organic reaction, reduction

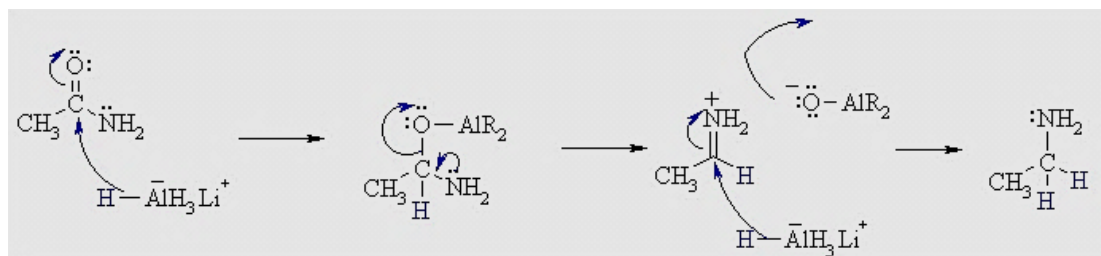
I. INTRODUCTION

Catalysis is the process of increasing the [rate](#) of a [chemical reaction](#) via adding a substance known as a catalyst. Catalysts are not consumed in the catalyzed reaction while can act repeatedly. Often only very small amounts of catalyst are required. In general, chemical reactions occur faster in the presence of a catalyst because the catalyst provides an alternative reaction pathway- or mechanism- with a lower activation energy than the non-catalyzed mechanism. In catalyzed mechanisms, the catalyst usually reacts to form intermediate, which then regenerates the original catalyst in a process



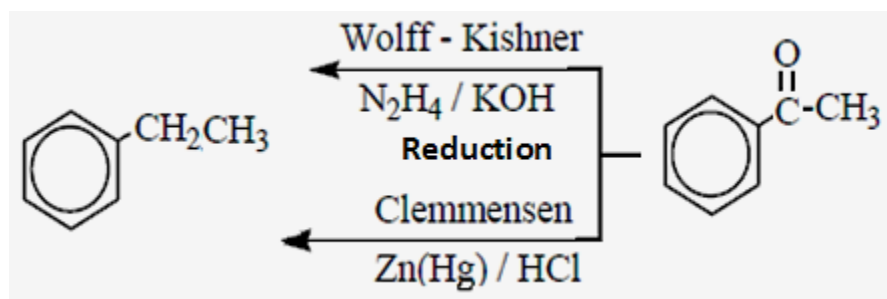
Reduction and Oxidation Catalysts :

- ❖ Amides (RCONHR) can be reduced to the amine (RCH₂NHR) by conversion of the carbonyl (C=O) to methyl group (-CH₂-).
- ❖ Amides can be reduced with LiAlH₄ to amine :



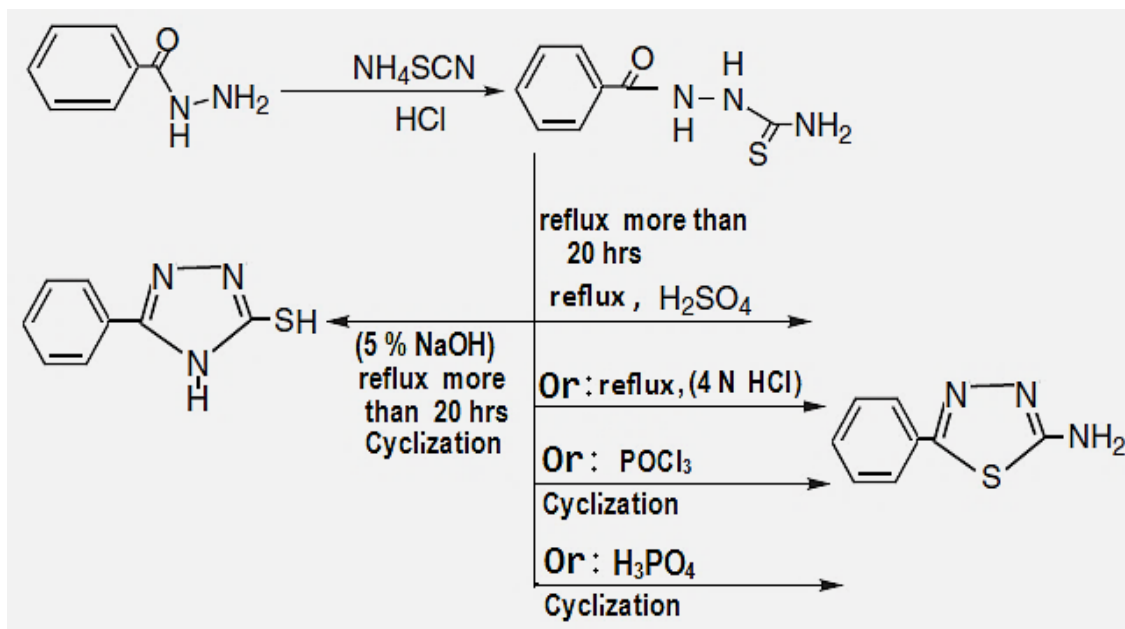
- ❖ Reduce carbonyl group to Alkanes (**Zn /Hg, HCl**).
- ❖ Reduce carbonyl group to Alkanes (**Na₂Cr₂O₇, HCl, Δ**).
- ❖ Substitute a carboxylic group on a aromatic compound to for a methyl group by (**NaNO₂, HCl**).
- ❖ **Fe** (Iron metal) to reduce nitro group to amines in the presence of an acid like (HCl).
- ❖ Hydrogen gas (**H₂**) is used to the reduction of alkenes, alkynes with catalysts like (**Pd and Pt**).
- ❖ Chromic acid (**H₂CrO₄**) is a strong acid and an oxidant (oxidize secondary alcohols to ketones and primary alcohols to carboxylic acids)
- ❖ Hydrogen peroxide (**H₂O₂**) to (oxidize aldehydes to carboxylic acids).
- ❖ (**KMnO₄**) Potassium permanganate, which used to oxidize primary alcohol and aldehyde to carboxylic acid while secondary alcohol to ketone.
- ❖ **LiAlH₄** (Lithium aluminum hydride) is a very strong reducing reagent. It used to reduce ((aldehydes, ketones, esters, and carboxylic acids to alcohols)) while ((amides and nitriles to amines)).
- ❖ Sodium borohydride (**NaBH₄**) is a reagent for the reduction of ketones and aldehydes.
- ❖ (**Sn**) or (**Zn**) with acid, used to reduce nitro groups to amines.
- ❖ (**Zn - Hg**) with acid, zinc amalgam used to reduce ketones alkanes (Clemmensen reaction).

In general, chemical reactions occur faster in the presence of a catalyst because the catalyst provides an alternative reaction pathway - or mechanism - with a lower [activation energy](#) than the non-catalyzed mechanism. In catalyzed mechanisms, the catalyst usually reacts to form a [intermediate](#), which then regenerates the original catalyst in a process., There are more than reduction methods like :

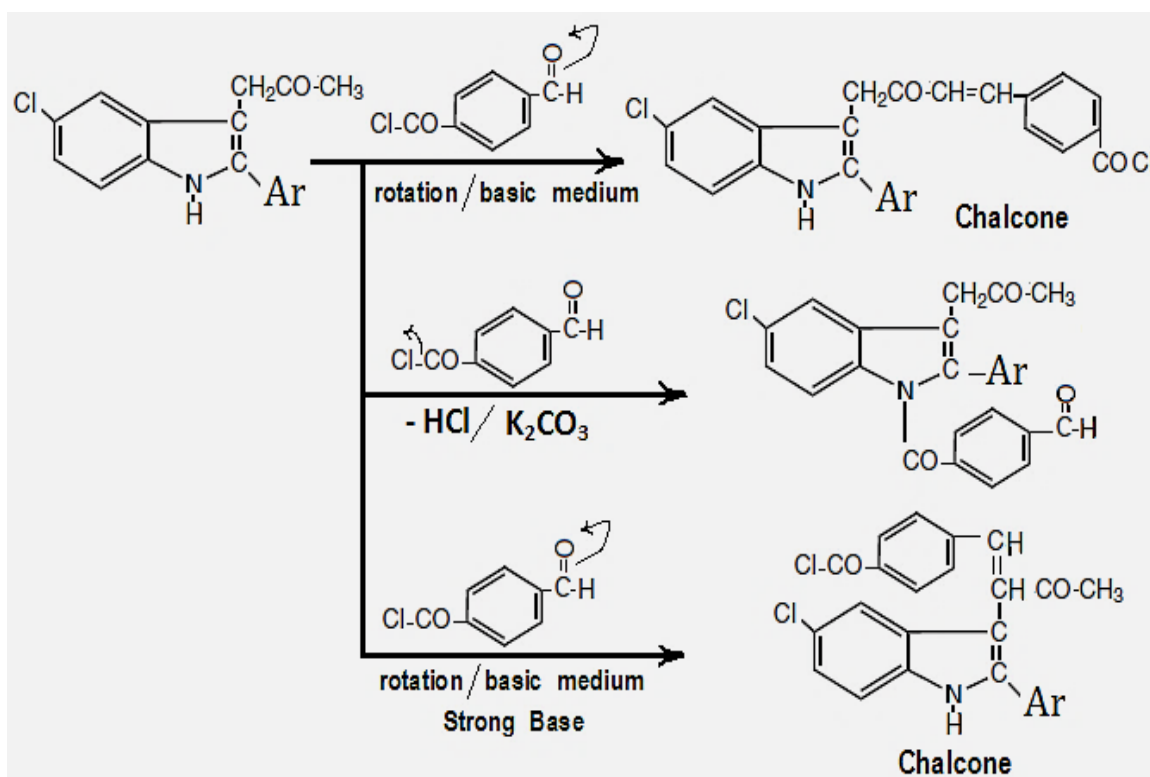


II. TYPES OF CATALYSTS:

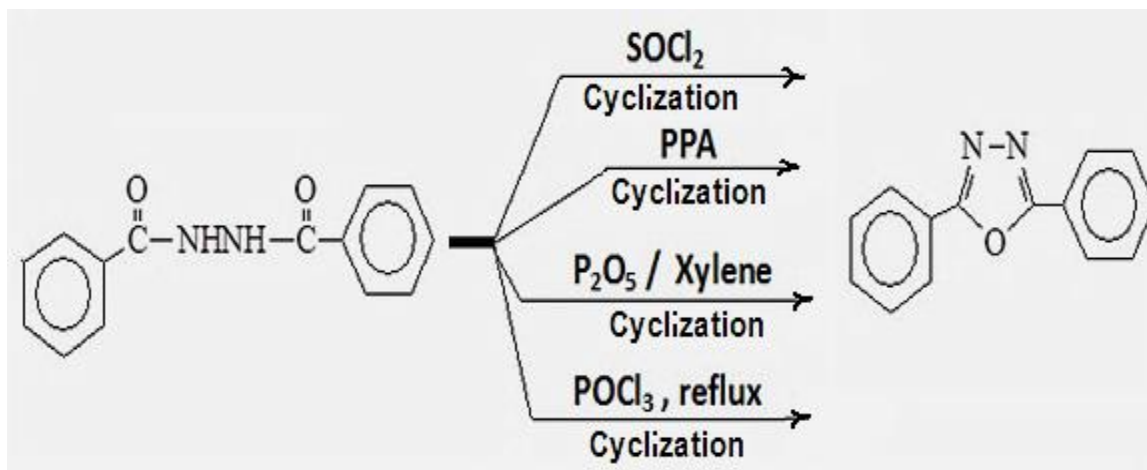
There are various catalysis give same product but in different conditions like the reaction of carbonylhydrazine with ammonium thiocyanate followed by cyclization reaction via many methods like (dehydration with sulfuric acid ., 5N of HCl ., acidic medium with H₃PO₄ ., or in basic medium in 5 % of NaOH solution) .



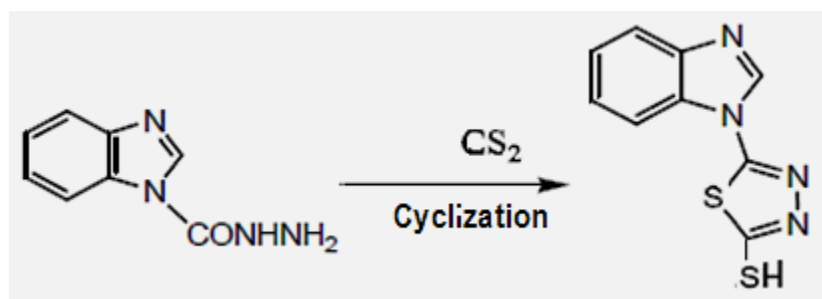
Other reaction involves two active group in same compound ,carbonyl of aldehyde and carbonyl of Aroyl halide, the product depends on catalysis:



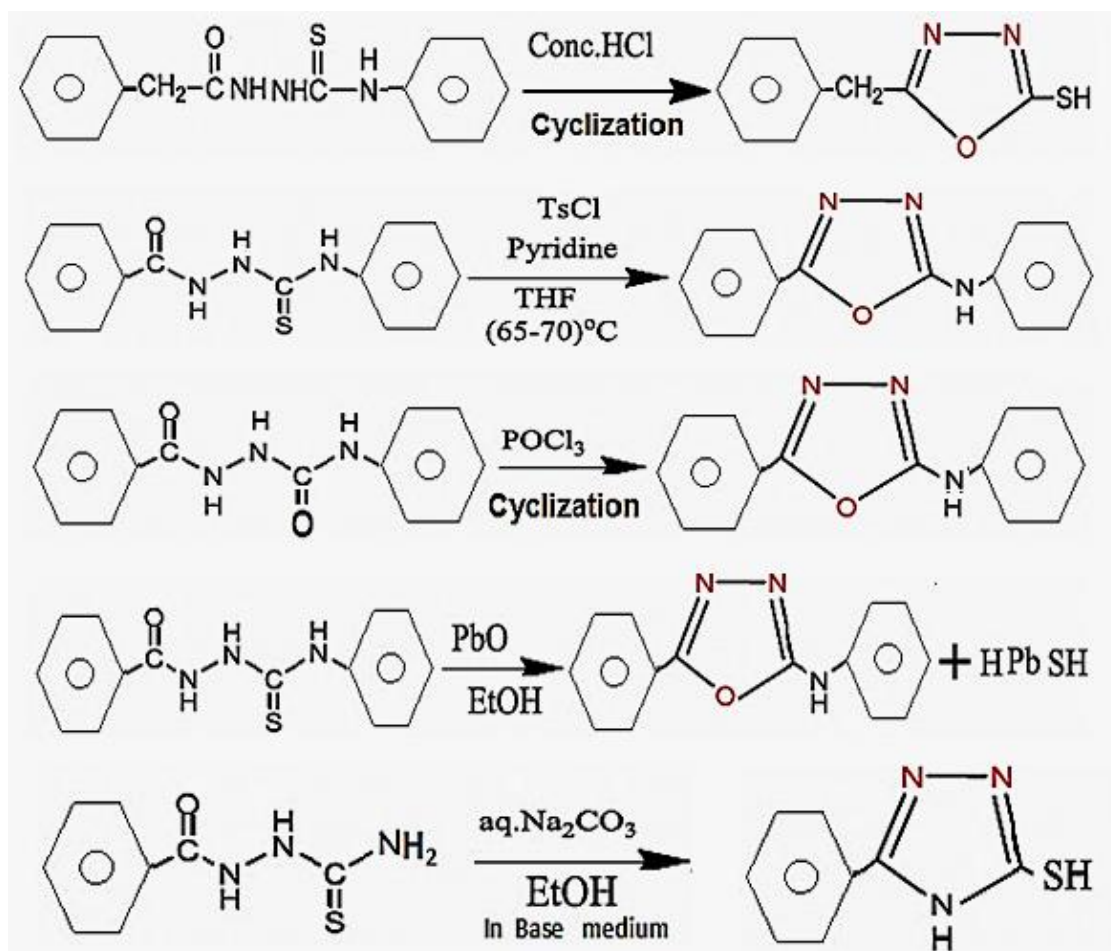
Other methods for cyclization process of hydrazine derivatives in presence of new catalysts:



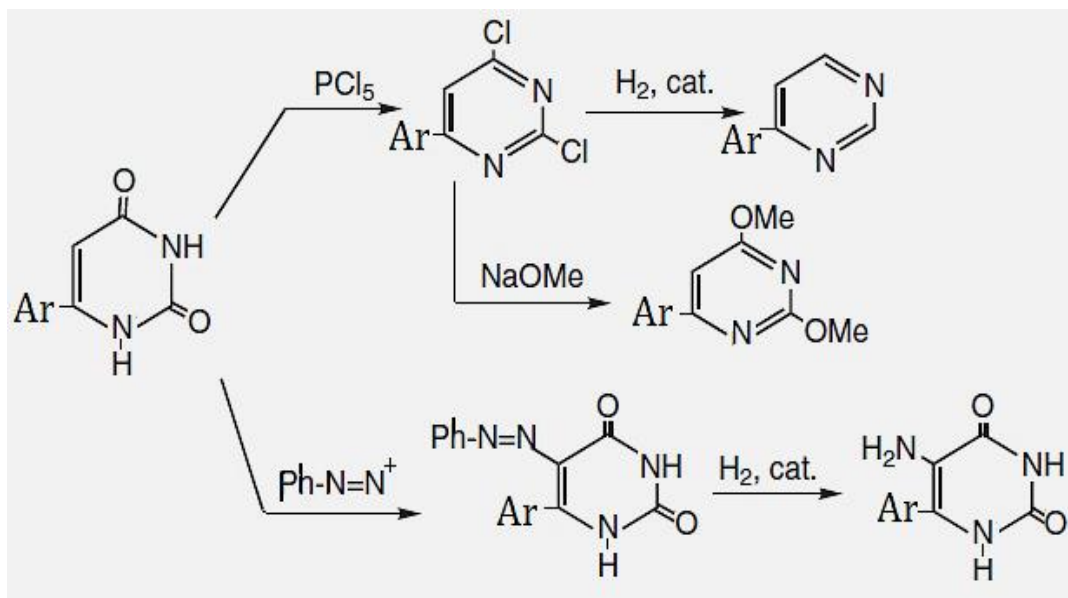
There is other catalyst represented by (CS_2) for ring closure of Hydrazone derivatives:



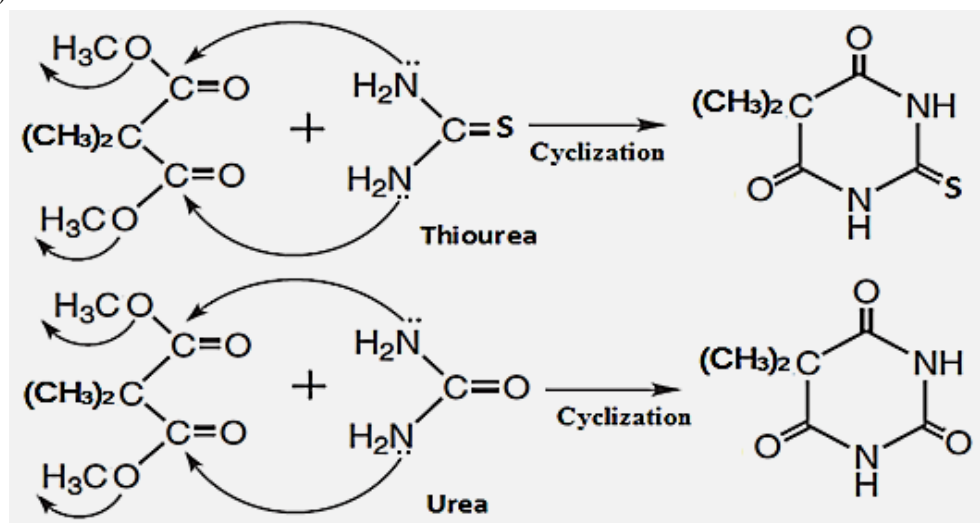
Also thiosemicarbazide and semicarbazide proceed to cyclization reaction by formation oxadiazole or thiadiazole or triazole ring via various catalyst :



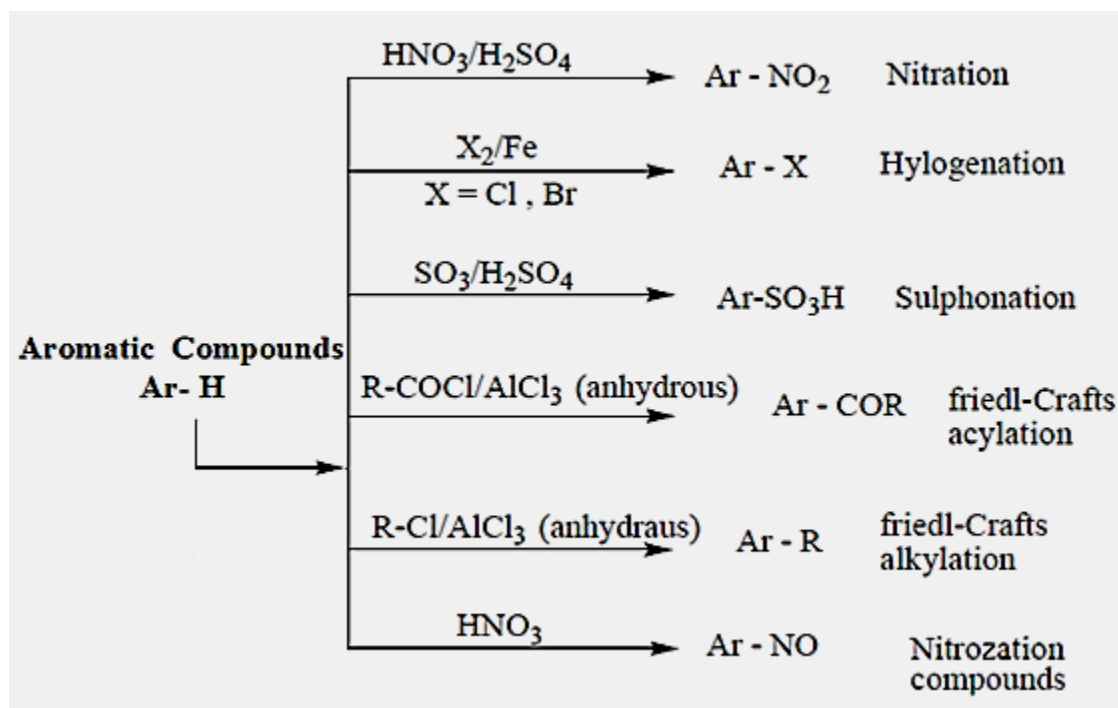
Also any product depends on type of functional group in any reactant not only catalysis or condition of reaction which affect directly on any products:



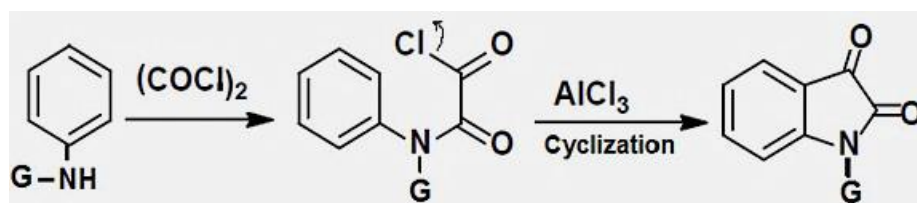
Reaction of any di-carbonyl compounds in cyclization reaction occurs in presence of acidic medium as a catalyst like (H_2SO_4 , HCl , ...)



In the other hand, there are several catalysts for substitution reaction in aromatic compounds represented by (HNO_3 , H_2SO_4 , SO_3H , AlCl_3 ,



The reaction with any alkyl or acyl halide proceed toward alkylation process by Aluminum chloride :



III. CONCLUSIONS

The chemical reactions occur faster in the presence of a catalyst because the catalyst provides an alternative reaction pathway- or mechanism- with a lower activation energy than the non-catalyzed mechanism.

REFERENCES

1. P. Degennes . **Soft Matter** ., Rev. Mod. Phys., 64 (3) (1992), pp. 645-648.
2. M. Lattuada, T.A. Hatton. **Synthesis, properties and applications of Janus nanoparticles.**, Nano Today, 6 (3) (2011), pp. 286-308.
3. Y. Song, S. Chen . **Janus nanoparticles: preparation, characterization, and applications** ., Chem. Asian J., 9 (2) (2014), pp. 418-430.
4. F. Liang, C. Zhang, Z. Yang. **Rational design and synthesis of Janus composites** Adv. Mater., 26 (40) (2014), pp. 6944-6949.
5. A. Perro, F. Meunier, V. Schmitt, S. Ravaine. **Production of large quantities of “Janus” nanoparticles using wax-in-water emulsions.**, Colloids Surf. Physicochem. Eng. Asp ., 332 (1) (2009), pp. 57-62.

6. Miad Mohamed ,Nagham Mahmood Aljamali ,Sabreen Ali Abdalrahman., Wassan Ala Shubber ., "Formation of Oxadiazole Derivatives Ligands from Condensation and Imination Reaction with References To Spectral Investigation, Thermal and Microbial Assay"., Biochem. Cell. Arch., 2018 ,18, 1, pp. 847-853.
7. S. Pradhan, L.-P. Xu, S. Chen. **Janus nanoparticles by interfacial engineering.** Adv. Funct. Mater., 17 (14) (2007), pp. 2385-2392.
8. Wang, Cuiling; Yan, Jiayu; Du, Mo; Burlison, Joseph A.; Li, Chi; Sun, Yanni; Zhao, Danqing; Liu, Jianli (2017). "One step synthesis of indirubins by reductive coupling of isatins with KBH 4" . *Tetrahedron* . 73 (19): 2780–2785. doi:10.1016/j.tet.2017.03.077.
9. Nagham Mahmood Aljamali., "The Various Preparation Methods in Synthetic Chemistry".,1 Edt. ,Evincepub Publishing house, 2019., ISBN :978-93-88277-82-2.
10. Nagham Mahmood Aljamali. "Reactions and Mechanisms".,1 Edt., IJMRA Publication ,2018 .,ISBN : 978- 93-87176-25-6 .
11. Matheus ME, de Almeida Violante F, Garden SJ. Isatins inhibit cyclooxygenase-2 and inducible nitric oxide synthase in a mouse macrophage cell line. *Eur J Pharmacol.* 2007; 556 :200–6.
12. Nagham Mahmood Aljamali . "*Experimental Methods for Preparation of Mannich_Bases, Formazan, Normal and Cyclic Sulfur Compounds*", 1st edition Evince pub Publishing House;2018, ISBN: 978-93-87905-19-1.
13. Nagham Mahmood Aljamali., "Alternative Methods in Organic Synthesis" .,1th-Edition, Eliva Press SRL, 2020 ., ISBN: 9798680201176.
14. Nagham Mahmood Aljamali. 2016. " Synthesis and Biological Study of Hetero (Atoms and Cycles) Compounds", *Der PharmaChemica*, 8,6, 40-48.
15. A. Perro, S. Reculusa, E. Bourgeat-Lami, E. Duguet, S. Ravaine. **Synthesis of hybrid colloidal particles: from snowman-like to raspberry-like morphologies** *Colloids Surf, Physicochem. Eng. Asp.*, 284 (2006), pp. 78-83.
16. M. Pera-Titus, L. Leclercq, J.-M. Clacens, F. De Campo, V. Nardello-Rataj**Pickering interfacial catalysis for biphasic systems: from emulsion design to green reactions.**, *Angew Chem. Int. Ed.*, 54 (7) (2015), pp. 2006-2021
17. . S.U. Pickering. **CXCVI.—emulsions.** *J. Chem. Soc. Trans.*, 91 (0) (1907), pp. 2001-2021.
18. M. Destribats, S. Gineste, E. Laurichesse, H. Tanner, F. Leal-Calderon, V. Heroguez. **Pickering emulsions: what are the main parameters determining the emulsion type and interfacial properties?** *Langmuir*, 30 (31) (2014), pp. 9313-9326.
19. Z. Fan, A. Tay, M. Pera-Titus, W.-J. Zhou, S. Benhabbari, X. Feng, *et al.***Pickering Interfacial Catalysts for solvent-free biomass transformation: physicochemical behavior of non-aqueous emulsions.** *J. Colloid Interface Sci.*, 427 (2014), pp. 80-90.
20. Imad Kareem Alwan Alsabri, Hasaneen Kudhair Abdullabass ,Nagham Mahmood Aljamali ., Invention of (Gluta.Sulfazane-Cefixime) Compounds as Inhibitors of Cancerous Tumors., *Journal of Cardiovascular Disease Research*, 2020,11, 2., 44-55 ., DOI: 10.31838/jcdr.2020.11.02.09 .
21. B.P. Binks. **Particles as surfactants: similarities differences.**, *Curr. Opin. Colloid Interface Sci.*, 7 (1-2) (2002), pp. 21-41
22. B.P. Binks, P.D.I. Fletcher**Particles adsorbed at the oil-water interface: a theoretical comparison between spheres of uniform wettability and “Janus” particles** *Langmuir* , 17 (16) (2001), pp. 4708-4710.
23. T. Ondarucu, P. Fabre, E. Raphael, M. Veyssie. **Specific properties of amphiphilic particles at fluid interfaces.**, *J. Phys.*, 51 (14) (1990), pp. 1527-1536.
24. C. Casagrande, P. Fabre, E. Raphael, M. Veyssie**Janus beads: realization and behavior at water oil interfaces.**, *Europhys. Lett.*, 9 (3) (1989), pp. 251-255.
25. Nagham Mahmood Aljamali., 2015. Synthesis and Chemical Identification of Macro Compounds of (Thiazol and Imidazol)" ., *Research J. Pharm. and Tech*, 8,1, 78-84., DOI: 10.5958/0974-360X.2015.00016.5.

26. W.-J. Zhou, L. Fang, Z. Fan, B. Albela, L. Bonneviot, F. De Campo. **Tunable catalysts for solvent-free biphasic systems: pickering interfacial catalysts over amphiphilic silica nanoparticles.**, J. Am. Chem. Soc., 136 (13) (2014), pp. 4869-4872.
27. Mestaf M, Nawfel Muhammed Baqer Muhsin., NeuroQuantology, 2019.,17,11, 11-16 .,10.14704/nq.2019.17.11.NQ19108.
28. Miad Mohmed ,Naghah Mahmood Aljamali ,Sabreen Ali Abdalrahman., Wassan Ala Shubber ., "Formation of Oxadiazole Derivatives Ligands from Condensation and Imination Reaction with References To Spectral Investigation, Thermal and Microbial Assay"., Biochem. Cell. Arch., 2018 ,18, 1, pp. 847-853.
29. Nagham Mahmood Aljamali.,Synthesis of Antifungal Chemical Compounds from Fluconazole with (Pharmaceutical) Studying, Research journal of Pharmaceutical, biological and chemical sciences, 2017, 8 (3), 564 -573.
30. Mehta SL, Manhas N, Raghuz R. Molecular targets in cerebral ischemia for developing novel therapeutics . Brain Res Rev. 2007;54:34–66.
31. Nawfel Muhammed Baqer Muhsin, Hayder H K, Noor H D, Nagham Mahmood Aljamali., "Preparation of Chemical Inhibitors to Treat the Corrosion and Erosion of Machines", International Journal of Engineering, Applied and Management Sciences Paradigms.,2019, 54,3,89-93.
32. Meaad M ,Naghah Mahmood Aljamali ,Nadheema A A .,"Preparation,Spectral Investigation, Thermal Analysis, Biochemical Studying of New (Oxadiazole -Five Membered Ring)-Ligands"., Journal of Global Pharmacy Technology,2018;10,1,20-29.
33. J. Faria, M.P. Ruiz, D.E. Resasco**Phase-selective catalysis in emulsions stabilized by Janus silica-nanoparticles.**, Adv. Synth. Catal., 352 (14-15) (2010), pp. 2359-2364.
34. Y. Liu, J. Hu, X. Yu, X. Xu, Y. Gao, H. Li. **Preparation of Janus-type catalysts and their catalytic performance at emulsion interface.**, J. Colloid Interface Sci., 490 (2017), pp. 357-3564.
35. Z.W. Seh, S. Liu, S.-Y. Zhang, M.S. Bharathi, H. Ramanarayan, M. Low. **Anisotropic growth of titania onto various gold nanostructures: synthesis, theoretical understanding, and optimization for catalysis.**, Angew Chem. Int. Ed., 50 (43) (2011), pp. 10140-10143.
36. S. Pradhan, D. Ghosh, S. Chen. **Janus nanostructures based on Au-TiO₂ heterodimers and their photocatalytic activity in the oxidation of methanol.**, Acs Appl. Mater. Interfaces , 1 (9) (2009), pp. 2060-2065.
37. A.A. Ismail, D.W. Bahnemann, I. Bannat, M. Wark **Gold nanoparticles on mesoporous interparticle networks of titanium dioxide nanocrystals for enhanced photonic efficiencies**J. Phys. Chem. C., 113 (17) (2009), pp. 7429-7435.
38. Masel, Richard I. (2001) Chemical Kinetics and Catalysis. Wiley-Interscience, New York. [ISBN 0-471-24197-0](#).
39. Laidler, K.J. and Meiser, J.H. (1982) Physical Chemistry, Benjamin/Cummings, p. 425. [ISBN 0-618-12341-5](#).
40. Laidler, Keith J.; Meiser, John H. (1982). *Physical Chemistry. Benjamin/Cummings.* pp. 424–425. [ISBN 0-8053-5682-7](#).
41. Atkins, Peter; de Paula, Julio (2006). *Atkins' Physical Chemistry (8th ed.)*. W.H.Freeman. p. 839. [ISBN 0-7167-8759-8](#).
42. Steinfeld, Jeffrey I.; Francisco, Joseph S.; Hase, William L. (1999). *Chemical Kinetics and Dynamics (2nd ed.)*. Prentice Hall. pp. 147–150. [ISBN 0-13-737123-3](#).
43. Jacoby, Mitch (16 February 2009). ["Making Water Step by Step"](#). *Chemical & Engineering News*. p. 10.
44. Matthiesen J, Wendt S, Hansen JØ, Madsen GK, Lira E, Galliker P, Vestergaard EK, Schaub R, Laegsgaard E, Hammer B, Besenbacher F (2009). "Observation of All the Intermediate Steps of a Chemical Reaction on an Oxide Surface by Scanning Tunneling Microscopy". *ACS Nano*. 3 (3): 517–26.
45. Robertson, A.J.B. (1970) Catalysis of Gas Reactions by Metals. Logos Press, London.

46. Shafiq, Iqrash; Shafique, Sumeer; Akhter, Parveen; Yang, Wenshu; Hussain, Murid (2020-06-23). "[Recent developments in alumina supported hydrodesulfurization catalysts for the production of sulfur-free refinery products: A technical review](#)". *Catalysis Reviews*. **0**: 1–86. doi:10.1080/01614940.2020.1780824. ISSN 0161-4940.
47. Housecroft, Catherine E.; Sharpe, Alan G. (2005). *Inorganic Chemistry (2nd ed.)*. Pearson Prentice-Hall. p. 805. ISBN 0130-39913-2.
48. Knözinger, Helmut and Kochloefl, Karl (2002) "Heterogeneous Catalysis and Solid Catalysts" in Ullmann's Encyclopedia of Industrial Chemistry, Wiley-VCH, Weinheim . doi:10.1002/14356007.a05_313
49. Wei, Hui; Wang, Erkang (2013-06-21). "Nanomaterials with enzyme-like characteristics (nanozymes): next-generation artificial enzymes". *Chemical Society Reviews*. **42** (14): 6060–93. doi:10.1039/C3CS35486E. ISSN 1460-4744. PMID 23740388.
50. Behr, Arno (2002) "Organometallic Compounds and Homogeneous Catalysis" in Ullmann's Encyclopedia of Industrial Chemistry, Wiley-VCH, Weinheim . doi:10.1002/14356007.a18_215
51. Elschenbroich, C. (2006) *Organometallics*. Wiley-VCH: Weinheim. ISBN 978-3-527-29390-2
52. Nelson, D.L. and Cox, M.M. (2000) *Lehninger, Principles of Biochemistry 3rd Ed.* Worth Publishing: New York. ISBN 1-57259-153-6.
53. [Catalytic Antibodies Simply Explained](#). Documentroot.com (2010-03-06). Retrieved on 2015-11-11.
54. Solovev, Alexander A.; Sanchez, Samuel; Mei, Yongfeng; Schmidt, Oliver G. (2011). "[Tunable catalytic tubular micro-pumps operating at low concentrations of hydrogen peroxide](#)". *Physical Chemistry Chemical Physics*. **13** (21): 10131–35.
55. Hävecker, Michael; Wrabetz, Sabine; Kröhnert, Jutta; Csepei, Lenard-Istvan; Naumann d'Alnoncourt, Raoul; Kolen'Ko, Yury V.; Girgsdies, Frank; Schlögl, Robert; Trunschke, Annette (2012). "[Surface chemistry of phase-pure M1 MoVTeNb oxide during operation in selective oxidation of propane to acrylic acid](#)". *Journal of Catalysis*. **285**: 48–60.
56. Naumann d'Alnoncourt, Raoul; Csepei, Lénárd-István; Hävecker, Michael; Girgsdies, Frank; Schuster, Manfred E.; Schlögl, Robert; Trunschke, Annette (2014). "[The reaction network in propane oxidation over phase-pure MoVTeNb M1 oxide catalysts](#)". *Journal of Catalysis*. **311**: 369–385. doi:10.1016/j.jcat.2013.12.008. hdl:11858/00-001M-0000-0014-F434-5.
57. Mokrani, Touhami; van Reenen, Albert; Amer, Ismael (2015). "[Molecular weight and tacticity effect on morphological and mechanical properties of Ziegler–Natta catalyzed isotactic polypropylenes](#)". *Polímeros*. **25** (6): 556–563. doi:10.1590/0104-1428.2158 . ISSN 0104-1428.
58. [Dub, Pavel A.; Gordon, John C. \(2018\). "The role of the metal-bound N–H functionality in Noyori-type molecular catalysts". *Nature Reviews Chemistry*. **2** \(12\): 396–408. doi:10.1038/s41570-018-0049-z. S2CID 106394152.](#)
59. Clark, Jim (October 2013). "[Types of catalysis](#)". *Chemguide*. Bård Lindström and Lars J. Petterson (2003) "[A brief history of catalysis](#)" *Cattech*, **7** (4) : 130–38.
60. Rayner-Canham, Marelene; Rayner-Canham, Geoffrey William (2001). [Women in Chemistry: Their Changing Roles from Alchemical Times to the Mid-Twentieth Century](#). American Chemical Society. ISBN 978-0-8412-3522-9.
61. Berzelius, J.J. (1835) Årsberättelsen om framsteg i fysik och kemi [Annual report on progress in physics and chemistry]. Stockholm, Sweden: Royal Swedish Academy of Sciences. After reviewing Eilhard Mitscherlich's research on the formation of ether, Berzelius coins the word katalys (catalysis) on p. 245:
62. Mitscherlich, E. (1834). "[Ueber die Aetherbildung](#)" [On the formation of ether]. *Annalen der Physik und Chemie*. **31** (18): 273–82. Bibcode:1834 AnP...107..273M

63. *Döbereiner (1822). "Glühendes Verbrennen des Alkohols durch verschiedene erhitzte Metalle und Metalloxyde" [Incandescent burning of alcohol by various heated metals and metal oxides]. Journal für Chemie und Physik. 34: 91–92.*
64. *Döbereiner (1823). "Neu entdeckte merkwürdige Eigenschaften des Platinsuboxyds, des oxydirten Schwefel-Platins und des metallischen Platinstaubes" [Newly discovered remarkable properties of platinum suboxide, oxidized platinum sulfide and metallic platinum dust]. Journal für Chemie und Physik. 38: 321–26.*
65. *Davy, Humphry (1817). "Some new experiments and observations on the combustion of gaseous mixtures, with an account of a method of preserving a continued light in mixtures of inflammable gases and air without flame". Philosophical Transactions of the Royal Society of London. 107: 77–85. doi:10.1098/rstl.1817.0009.*
66. *Roberts, M.W. (2000). "Birth of the catalytic concept (1800–1900)". Catalysis Letters. 67 (1): 1–4. doi:10.1023/A:1016622806065. S2CID 91507819.*
67. *Nicholas, Christopher P. (21 August 2018). "Dehydration, Dienes, High Octane, and High Pressures: Contributions from Vladimir Nikolaevich Ipatieff, a Father of Catalysis". ACS Catalysis. 8 (9): 8531–39. doi:10.1021/acscatal.8b02310.*
68. *Dhara SS; Umare SS (2018). A Textbook of Engineering Chemistry. India: S. Chand Publishing. p. 66. ISBN 9789352830688.*
69. *Laidler, K.J. (1978) Physical Chemistry with Biological Applications, Benjamin/ Cummings . pp. 415–17. ISBN 0-8053-5680-0.*
70. *Lindlar, H. and Dubuis, R. (2016). "Palladium Catalyst for Partial Reduction of Acetylenes". Organic Syntheses. doi:10.15227/orgsyn.046.0089; Collective Volume, 5, p. 880*
71. *Jencks, W.P. (1969) Catalysis in Chemistry and Enzymology McGraw-Hill, New York. ISBN 0-07-032305-4*
72. *Bender, Myron L; Komiyama, Makoto and Bergeron, Raymond J (1984) The Bioorganic Chemistry of Enzymatic Catalysis Wiley-Inter. science, Hoboken, U.S. ISBN 0-471-05991-9.*
73. *Duran A, Dogan HN, Rollas S. Synthesis and preliminary anticancer activity of new 1,4-dihydro-3-(3-hydroxy-2-naphthyl)-4-substituted-5H-1,2,4-triazoline-5-thiones. Farmaco. 2002;57:559–564.*
74. *El Shehry MF, Abu-Hashem AA, El-Telbani EM. Synthesis of 3-((2,4-dichlorophenoxy)methyl)-1,2,4-triazolo (thiadiazoles and thiadiazines) as anti-inflammatory and molluscicidal agents. Eur J Med Chem. 2010;45:1906–1911.*
75. *Gadad AK, Noolvi MN, Karpoornath RV. Synthesis and anti-tubercular activity of a series of 2-sulfonamido /trifluoromethyl-6-substituted imidazo-[2,1-b]-1,3,4-thiadiazole derivatives. Bioorg Med Chem. 2004;12:5651–5659.*
76. *Gülerman NN, Doğan HN, Rollas S, Johansson C, Çelik C. Synthesis and structure elucidation of some new thioether derivatives of 1,2,4-triazoline-3-thiones and their antimicrobial activities. Farmaco. 2001;56:953–958.*
77. *Holmwood G, Buechel KH, Plempel M, Haller J (1982) Antimicrobial azoles. Chem Abstr 96:62979s. Patent RFN DE 3018865, 1981*
78. *Kaplaushenko AH, Panasenko OI, Knish EH, Scherbina RO. Synthesis, physicochemical and biological properties of 2-(5-R1-4-R2-1,2,4-triazol-3-ylthio)acetic acids. Farm Zh. 2008;2:67–72.*
79. *Klimešová V, Zahajská L, Waisser K, Kaustová J, Möllmann U. Synthesis and antimycobacterial activity of 1,2,4-triazole 3-benzylsulfanyl derivatives. Farmaco . 2004;59:279–288. doi: 10.1016/j.farmac.2004 .01.006 .*
80. *Kumar D, Kumar NM, Chang K-H, Shah K. Synthesis and anticancer activity of 5-(3-indolyl)-1,3,4-thiadiazoles . Eur J Med Chem. 2010;45:4664–4668. doi: 10.1016/j .ejmech.2010.07.023.*