

# RECENT DEVELOPMENT ON PALLADIUM CATALYZED SUZUKI TYPE CROSS COUPLING REACTIONS

# Shagun Goyat<sup>\*</sup>

#### Abstract

The Suzuki coupling reaction, which forms carbon-carbon bonds between aryl halides and compounds containing boron, is a well-known and often employed process. Palladium is the catalyst in this reaction. The goal of the current research is to broaden the understanding of the Suzuki coupling reaction, especially with regard to difficult substrates and asymmetric variations. The Suzuki coupling reaction can be included into multicomponent and cascade processes, which could facilitate the quick synthesis of a variety of chemical structures. We present a brief review of the main features of the Suzuki coupling reactions in this article.

Keywords: Palladium catalysis, Suzuki reaction, Cross-coupling, Aryl halides, Boron-compounds

\*AIJHM College Rothak

DOI: 10.53555/ecb/2022.11.8.113

# Introduction

One of the mainstays of contemporary organic synthesis, the palladium-catalyzed Suzuki coupling process transformed the synthesis of carbon-carbon bonds. This transformative reaction was created by Akira Suzuki<sup>1</sup> in the 1970s and is now a vital tool in the toolbox of synthetic chemists all over the world. The Suzuki reaction makes it easier to couple aryl halides with compounds that include boron. This allows for the quick and effective assembly of biaryl motifs, which are a common structural component in materials science. agrochemicals, and medicines.<sup>2</sup> Fundamentally, the Suzuki coupling reaction is a prime example of transition metal catalysis, utilizing palladium's special reactivity to facilitate the joining of several building components.<sup>3</sup> The ligand design of the mechanistic route fine-tunes the successive steps of oxidative addition, transmetalation, and reductive elimination to achieve optimal efficiency and selectivity. The creation of advanced catalytic systems that can precisely orchestrate intricate transformations has been made possible by this mechanistic insight.

The exceptional substrate universality of the Suzuki coupling reaction is one of its distinguishing characteristics. It may couple aryl halides with a variety of boron reagents, such as boronic acids, boronate esters, and heterocycles containing boron, and accepts a broad range of aryl halides, from electron-rich substrates basic to highly functionalized molecules.<sup>4</sup> The Suzuki reaction's adaptability has elevated it to the forefront of contemporary synthetic chemistry, enabling chemists to reach previously unreachable chemical spaces with never-before-seen ease. On the other hand, the palladium-catalyzed Suzuki coupling process is the result of decades of research and development.<sup>5</sup> By means of a thorough examination of its workings, reach, latest developments, and uses, we hope to demonstrate the long-lasting influence of this revolutionary synthesis approach. As we go deeper into the complexities of the Suzuki coupling reaction, which holds great significance for the growth of materials research, drug discovery, and other fields in addition to its scientific significance.

# Literature Review

Since its discovery by Akira Suzuki in the 1970s, the palladium-catalyzed Suzuki coupling reaction has attracted a great deal of interest and praise in the synthetic chemistry world. The scientific literature<sup>6</sup> has exhaustively documented its broad adoption and revolutionary impact on organic synthesis, highlighting its key role in the synthesis of biaryl compounds and the building of carboncarbon bonds. The broad substrate range of the Suzuki coupling reaction, which includes a variety of aryl halides and compounds containing boron, is indicative of its adaptability.7 Its broad compatibility has aided in its widespread use in a variety of synthetic chemistry fields, such as materials science, natural product synthesis, and medicinal chemistry. The fact that it can be used to build intricate molecular structures, including functional materials and pharmacological intermediates, emphasizes the importance of this synthetic tool, which is incredibly effective and versatile.

Suzuki coupling research has recently advanced by tackling long-standing issues and expanding the limits of reaction efficiency and selectivity.8 Techniques like flow chemistry, catalyst immobilization, and the creation of bioorthogonal variations have made the Suzuki reaction more widely applicable and have allowed it to reach previously unreachable areas. Moreover, attempts to include the Suzuki coupling reaction into reactions and multistep cascade synthesis methodologies have simplified synthetic pathways and made it possible to quickly and precisely assemble complex molecular scaffolds. The goals of ongoing research are to improve catalytic efficiency, broaden the scope of reactions, and create ecologically safe and long-lasting catalytic systems.9

Chemists can advance the boundaries of chemical innovation and discovery by fully utilizing this revolutionary synthetic process to tackle present and future difficulties in drug development, materials science, and other related fields.

# **Catalyst Design**

Synthesis of new palladium catalysts that are more selective, stable, and active. Investigation of ligand changes to improve catalytic performance and make difficult substrates easier to couple.<sup>10</sup>

# Sustainable Catalysis

Suzuki coupling techniques incorporate green chemistry ideas, emphasizing the use of environmentally friendly ligands, solvents, and reaction conditions. creation of reusable or recyclable catalytic systems to cut down on waste and lessen the impact on the environment.<sup>11</sup>

#### **Base-Free Conditions**

Investigation of base-free environments to expedite reaction methods and steer clear of the usage of pricy or potentially hazardous bases. Utilizing novel cocatalysts or reaction additives to enable effective catalysis in moderate circumstances.<sup>12</sup>

#### **Heterogeneous Catalysis**

Application of metal-organic frameworks (MOFs), silica, polymers, and other solid materials as supports for heterogeneous palladium catalysts. improved catalyst recovery and recyclability, making Suzuki coupling methods more economical and environmentally friendly.<sup>13</sup>

#### Asymmetric Suzuki Coupling

creation of enantioselective Suzuki coupling techniques using catalysts or chiral ligands. facilitating the production of highly stereocontrolled, optically pure biaryl compounds, which are useful in the synthesis of natural products and medicinal chemistry.<sup>14</sup>

#### **Dual Catalysis Approaches**

amalgamation of organocatalysis, photoredox, enzymatic, and palladium catalysis with other catalytic systems. synergistic effects in Suzuki coupling processes that improve selectivity, reactivity, and functional group tolerance.<sup>15</sup>

#### **C-H Activation**

Investigation of C-H activation techniques catalyzed by palladium to facilitate the direct coupling of arenes or heteroarenes with boron reagents. enhancement of Suzuki coupling techniques that are more atom-efficient and stepefficient while avoiding the requirement for prefunctionalized substrates.<sup>16</sup>

#### **Cascade Reactions**

Design of cascade reactions utilizing palladium as a catalyst for several steps in the formation of C-C bonds. enabling one procedure to access complex molecular scaffolds, thus increasing the synthetic value of Suzuki coupling.<sup>17</sup>

# **Bioorthogonal Suzuki Coupling**

Creation of bioorthogonal Suzuki coupling techniques to modify biomolecules site-selectively in a physiological environment. enabling selective labeling and functionalization for uses in chemical biology, imaging, and drug administration.<sup>18</sup>

# **Flow Chemistry**

The Suzuki coupling reaction's incorporation into continuous flow systems to improve reaction efficiency and allow for exact control over reaction parameters. assistance with process intensification and scale-up for industrial applications.<sup>19</sup> Palladium-catalyzed Suzuki coupling approach continues to be an important tool in modern synthetic chemistry, with many applications in both academia and industry. These latest developments in the field highlight this.

#### Conclusion

In conclusion, the palladium-catalyzed Suzuki coupling reaction represents a versatile and indispensable tool in modern organic synthesis, offering efficient access to biaryl compounds with widespread applications. Its continued development and application hold significant implications for the advancement of synthetic chemistry and the discovery of new bioactive compounds.

#### References

- 1. Miyaura, N. and Suzuki, A., 1995. Palladiumcatalyzed cross-coupling reactions of organoboron compounds. *Chemical reviews*, 95(7), pp.2457-2483.
- 2. Suzuki, A., 2004. Organoborane coupling reactions (Suzuki coupling). *Proceedings of the Japan Academy, Series B*, 80(8), pp.359-371.
- 3. Miyaura, N., Yamada, K. and Suzuki, A., 1979. A new stereospecific cross-coupling by the palladium-catalyzed reaction of 1alkenylboranes with 1-alkenyl or 1-alkynyl halides. *Tetrahedron Letters*, 20(36), pp.3437-3440.
- 4. Suzuki, A., 1999. Recent advances in the crosscoupling reactions of organoboron derivatives with organic electrophiles, 1995–1998. *Journal of Organometallic Chemistry*, 576
- Farhang, M., Akbarzadeh, A.R., Rabbani, M. and Ghadiri, A.M., 2022. A retrospectiveprospective review of Suzuki–Miyaura reaction: From cross-coupling reaction to pharmaceutical industry applications. *Polyhedron*, 227, p.116124.
- 6. Malav, R., Yadav, S. and Ray, S., 2024. Catalytic efficacy of Schiff base-palladium complexes in important C–C cross coupling reactions: a brief overview. *Proceedings of the Indian National Science Academy*, pp.1-13.
- Srivastava, A., Kaur, H., Pahuja, H., Rangarajan, T.M., Varma, R.S. and Pasricha, S., 2024. Optimal exploitation of supported heterogenized Pd nanoparticles for CC crosscoupling reactions. *Coordination Chemistry Reviews*, 507, p.215763.
- 8. Wei, D. and Lin, G.Q., 2024. Recent Advances in Pd-Catalyzed Reactions Involving the "On-Water" Mechanism. *Chinese Journal of Chemistry*, 42(5), pp.533-542.
- 9. Markham, S., Crans, D.C. and Atwater, B., 2024. Using speciation to gain insight into sustainable coupling reactions and their catalysts. *Catalysis for a Sustainable Environment: Reactions, Processes and Applied Technologies*, pp.559-576.

- 10.Kalikadien, A.V., Mirza, A., Hossaini, A.N., Sreenithya, A. and Pidko, E.A., 2024. Paving the road towards automated homogeneous catalyst design. *ChemPlusChem*, p.e202300702.
- 11. Markham, S., Crans, D.C. and Atwater, B., 2024. Using speciation to gain insight into sustainable coupling reactions and their catalysts. *Catalysis for a Sustainable Environment: Reactions, Processes and Applied Technologies*, pp.559-576.
- 12.Lennox, A.J. and Lloyd-Jones, G.C., 2013. Transmetalation in the Suzuki–Miyaura coupling: the fork in the trail. *Angewandte Chemie International Edition*, 52(29), pp.7362-7370.
- 13.Phan, N.T., Van Der Sluys, M. and Jones, C.W., 2006. On the nature of the active species in palladium catalyzed Mizoroki–Heck and Suzuki–Miyaura couplings–homogeneous or heterogeneous catalysis, a critical review. Advanced Synthesis & Catalysis, 348(6), pp.609-679.
- 14.Baudoin, O., 2005. The asymmetric Suzuki coupling route to axially chiral biaryls. *European journal of organic chemistry*, 2005(20), pp.4223-4229.
- 15.Martin, R. and Buchwald, S.L., 2008. Palladium-catalyzed Suzuki– Miyaura crosscoupling reactions employing dialkylbiaryl phosphine ligands. *Accounts of chemical research*, 41(11), pp.1461-1473.
- 16.Chen, X., Engle, K.M., Wang, D.H. and Yu, J.Q., 2009. Palladium (II)-catalyzed C-H activation/C-C cross-coupling reactions: versatility and practicality. *Angewandte Chemie International Edition*, 48(28), pp.5094-5115.
- 17. Kotha, S., Lahiri, K. and Kashinath, D., 2002. Recent applications of the Suzuki–Miyaura cross-coupling reaction in organic synthesis. *Tetrahedron*, *58*(48), pp.9633-9695.
- 18. Ramil, C.P. and Lin, Q., 2013. Bioorthogonal chemistry: strategies and recent developments. *Chemical communications*, 49(94), pp.11007-11022.
- 19.Len, C., Bruniaux, S., Delbecq, F. and Parmar, V.S., 2017. Palladium-catalyzed Suzuki– Miyaura cross-coupling in continuous flow. *Catalysts*, 7(5), p.146.