 REVIEW ARTICLE

Synthesis and biological importance of amide analogues

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The Amide bond formation reactions are among the most important transformations in organic chemistry and biochemistry because of the widespread occurrence of amides in pharmaceuticals, natural products and biologically active compounds. The amide group is widely present in the drugs, intermediates, pharmaceuticals, and natural products. It is also available in a large number of industrial materials including polymers, detergents and lubricants (1). The common method for the preparation of amides involves the reaction of activated carboxylic acid derivatives, such as chlorides, anhydrides or esters, with amines or, alternatively, the direct union of the carboxylic acids with amines assisted by stoichiometric amounts of coupling reagents, such as carbodiimides or 1H-benzotriazole derivatives (1,2). However, these classical approaches are low in atom efficiency and generate large amounts of waste products, making their environmental profile unfavorable. Therefore, ACS Green Chemistry Institute and members of leading pharmaceutical corporations worldwide, identified “amide formation avoiding poor atom economy reagents” as one of the top challenges in organic chemistry (3). New efficient and sustainable synthetic routes to access this important class of compounds are therefore needed (4).

In the search of more atom-economical and cost-effective protocols, metal-catalyzed transformations have emerged in the last few years as attractive alternatives, offering the possibility to develop previously unavailable routes starting from substrates other than carboxylic acids and their derivatives (5). Thus, with the help of transition metals, a plethora of functional groups, such as nitriles, aldehydes, ketones, oximes, primary alcohols or amines, can be now conveniently employed as starting materials for the construction of the amide bond. There are three types of amides available in Chemistry: (i) an organic amide which is also referred as carboxamide, (ii) a sulfonamide, and (iii) a phosphoramid. Amides are usually regarded as derivatives of carboxylic acids in which the hydroxyl group has been replaced by an amine or ammonia.

There are many drugs available in the market which contains amide linkage in the nucleus and possesses various therapeutic activities. Few drugs with their structures (Figure 1) are mentioned below:

Figure 1) Structures of drugs with amide linkage.

ABSTRACT

The present research article deals with the amide analogues prepared by available very well-known name reactions. The author have studied the name reactions like Beckmann rearrangement, Schmidt reaction, Passerine reaction, Willgerodt-Kindler reaction and UGI reaction, which involves preparation of amide linkage containing compounds. The main purpose of article is to provide information on the development of novel amide derivatives to the scientific community. Doing so, it focuses on mechanisms of action and adverse events, and suggests measures to be implemented in the clinical practice according to bioethical principles.

Key Words: Novel amide derivatives; Amide analogues; Amide formation

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LITERATURE REVIEW

Beckmann rearrangement

The Beckmann rearrangement developed by German chemist Ernst Otto Beckmann since 1853-1923. It is an acid-catalyzed rearrangement of an oxime to an amide (6-8). Cyclic oximes yield lactams. Since the discovery of this Beckmann rearrangement (BR) in 1896, successive investigations have been largely carried out and applied in many ways. It is a skeletal rearrangement, which accomplishes both, the cleavage of C=C bond and formation of C-N bond. It has become a very useful and efficient method for incorporation of nitrogen atom in both cyclic and acyclic systems and also for the synthesis of various alkaloids. Several catalysts have been used for this rearrangement like zeolites, metal catalysts, FeCl₃/AgSbF₆ etc (9-12). Though BR is a very well-known and old reaction, rearrange cyclic oximes fused with heterocyclic ring systems. The reaction scheme of Beckmann rearrangement is mentioned below as Scheme-1:

Beckmann rearrangement reaction includes preparation of amide based compound from oxime derivatives using acid catalyst i.e., Con. H₂SO₄. The same reaction mechanism of Beckmann rearrangement is mentioned below as Scheme-2:

The reaction mechanism of the Beckmann rearrangement is in general believed to consist of an alkyl migration with expulsion of the hydroxyl group to form a nitrilium ion followed by hydrolysis.

Srivastava et al. (13) have synthesized amide compounds from aldoxide through performing an efficient Beckmann rearrangement at room temperature which involves an eosin Y catalyzed, visible-light-mediated in-situ formation of the Vilsmeier-Haack reagent from CBr₄ and a catalytic amount of DMF. This operationally simple method for the activation of ketoximes avoids the need for any corrosive, water-sensitive reagents and elevated temperatures. The reaction scheme of Beckmann Rearrangement performed by Srivastava et al. is mentioned below as Scheme-3:

Ganguly et al. (14) have synthesized amides from ketoximes using Mild Neutral Conditions through performing an efficient Iodine-Mediated Beckmann Rearrangement. The reaction scheme of Beckmann Rearrangement performed by Ganguli et al. is mentioned below as Scheme-4:

Sharghi et al. (15) have synthesized amides from Ketones and Aldehydes using Zinc Oxide via performing Solvent-Free and One-Step Beckmann rearrangement. The reaction scheme of Beckmann Rearrangement performed by Sharghi et al. is mentioned below as Scheme-4:

Ramon et al. (16) have synthesized Amides from Aldoximes using Au/Ag Cocatalyzed through performing Beckmann-Rearrangement under Solvent- and Acid-Free Conditions. The reaction scheme of Beckmann Rearrangement performed by Ramon et al. is mentioned below as Scheme-6:

Kusurkar et al. (17) have synthesized derivatives of biologically active scaffolds by performing Beckmann rearrangement successfully. The synthesis of two β-carbolinone, three new γ-carbolinone, new pyrolopyridinone and tetrahydroisquinolinone derivatives were achieved in good to moderate yields. Kusurkar et al. have synthesized following compounds i.e., Compound of formulae (Figure 2) I-IV by performing Beckmann Rearrangement:

Figure 2) Compound derivatives of Beckmann Rearrangement, Formulae 1-4

Shah et al. (18) have synthesized 97 various symmetrical, unsymmetrical, and N-substituted 1,4-dihydropyridines. The synthesized molecules were tested for their activity against M. tuberculosis H37Rv strain with rifampin as the standard drug. The percentage inhibition was found in the range 3-93%. In an effort to understand the relationship between structure and activity, 3D-QSAR studies were also carried out on a subset that is representative of the molecules synthesized. For the generation of the QSAR models, a training set of 35 diverse molecules representing the synthesized molecules was utilized. The molecules were aligned using the atom-fit technique like CoMFA and CoMSIA models. The QSAR models revealed the importance of the conformational flexibility of the substituents in antitubercular activity. Shah et al. have synthesized following compounds generically mentioned as compound of formula-V (Figure 3) by performing Beckmann Rearrangement:

Figure 3) Compound derivatives of Beckmann Rearrangement, Formula 5

Spring et al. (19) have developed a convenient, efficient, and general method for the synthesis of benzo-fused seven- and eight-membered ring lactams by the Beckmann rearrangement of cyclic oximes. The process is technically simple, proceeds under relatively mild conditions with complete regioselectivity, and forms biologically interesting products that are generally difficult to synthesize by other methods. The biological activities of the synthesized compounds were also evaluated. In addition, several structurally novel Narylated derivatives were prepared, demonstrating the capability to carry out structural elaboration around the lactam core. Overall, the work reported by Spring et al. represents a convenient approach to benzo-fused eight-membered ring lactams, a biologically interesting chemotype that is underrepresented in current small-molecule collections. The enrichment of screening libraries with compounds of this sort will allow the sampling of previously untapped regions of chemical space, which may thus facilitate the discovery of novel biologically active small-molecule agents. As such, author anticipated that this methodology will prove especially valuable in the diversity-oriented synthesis of structurally diverse small-molecule collections. Spring et al.
have synthesized following compounds as compound of formula-VI (Figure 4) by performing Beckmann Rearrangement:

![Formula VI](image)

**Figure 4** Compound derivatives of Beckmann Rearrangement, Formula 6

### Schmidt Reaction

The Schmidt reaction is an organic reaction in which an azide reacts with a carbonyl group to give an amine or amide, with expulsion of nitrogen. It is named after Karl Friedrich Schmidt (1887–1971), who first reported it in 1924 by successfully converting benzophenone and hydroxyacetic acid to benzaldehyde. Surprisingly, the intramolecular reaction was not reported until 1991 but has become important in the synthesis of natural products. The reaction scheme of Schmidt reaction is mentioned below as Scheme-7:

In the reaction mechanism for the ketone Schmidt reaction, the carbonyl group is activated by protonation for nucleophilic addition by the azide, forming intermediate compound, which loses water in an elimination reaction to temporary imine, over which one of the alkyl groups migrates from carbon to nitrogen with loss of nitrogen. Attack by water and proton loss converts intermediate compounds, which is a tautomer of the final amide (21). The said mechanism is mentioned below as Scheme-8:

Aube et al. (22) have performed Schmidt reaction via carrying out two variations in which ketone electrophiles in hexafluoroisopropanol (HFIP) solvent has been studied. When TMSN₃ is reacted in HFIP in the presence of triflic acid (TfOH) promoter, tetrozoles are obtained as the major products. This observation is in contrast to established methods, which usually lead to amides or lactams arising from formal NH insertion as the major products. The full product profiles of several examples of this reaction are also reported and found to include mechanistically interesting products (e.g., double ring expansion). Application of TfOH promoter in HFIP was also found to promote the reaction of a hydroxyalkyl azide with a ketone, which affords lactams following nucleophilic opening of initially formed imine ether more efficiently than previously reported methods. The reaction scheme of Schmidt reaction performed by Aube et al. is mentioned below as Scheme-9:

Iyengar et al. (23) have disclosed a total synthesis of (+)-aspidospermidine, which featuring an intramolecular Schmidt reaction as the key step. The effects of stereoselectivity and protecting group status on the regio- and chemoselectivity of this reaction were examined. Further motivation was provided by the opportunity to examine a challenging intramolecular Schmidt reaction, in which selectivity for insertion into only one of two regioisomeric ketones is required for the preparation of a key tricyclic lactam. The reaction scheme of Schmidt reaction performed by Iyengar et al. is mentioned below as Scheme-10:

Jia et al. (24) have disclosed the Schmidt reaction, which includes the treatment with hydroxyacetic acid in the presence of a strong acid, converts ketones to amides directly and has been broadly applied in organic synthesis. In this communication, the Schmidt reaction of ketones was carried out in DME solution in a continuous-flow micro reactor and gave the amide products. The enclosed small-volume feature of the micro reactor made this reaction safe, fast, and cost effective. The reaction scheme of Schmidt reaction performed by Jia et al. is mentioned below as Scheme-11:

Huntress et al. (25) have disclosed Schmidt reaction which includes preparation of amide from primary, secondary and tertiary acid. The organic acid is shown reacting with hydroxyacetic acid only in the form of a positive ion, since changes in the system which result in decreased ionization of the organic acid, also result in incomplete reaction. In reaction which yield both carbon monoxide and carbon dioxide, complex ionization of the organic acids is assumed, the dihydroxy-carbonium ions [29] reacting to form carbon dioxide and primary amines or cleavage products, the oxocarbonium ions [30] reacting to form carbon monoxide and cleavage products (but no primary amine). This interpretation is chosen to explain the following observations: as reaction conditions are changed to decrease the yield of carbon monoxide, the yield of primary amine is increased. The proportion of carbon monoxide formed increased by factors (dilution and accumulation of reaction products) which would reverse the mobile equilibrium shown. The proportion of primary amine to cleavage products was observed to vary on dilution of the reaction medium, if no carbon monoxide was evolved. The alternative assumption is that only one ion (29 or 30) can react in three different ways, only one of which is sensitive to dilution, appears a less satisfactory explanation.

Ion [30] is shown decomposing to form carbon monoxide and a carbonium ion only after reaction with hydroxyacetic acid and not directly as does triphenylacetic acid since no gas evolution occurred with these acids until sodium azide was added to the reaction mixture. The assumption that in complex [31] electron-withdrawal from the alkyl group and the dihydroxy-carbonium ions proceeds in preference to the cleavage of the nitrogen-nitrogen bond does not appear unlikely in view of the tendency for Carbonium ion formation when carbon dioxide is evolved. The reaction scheme of Schmidt reaction performed by Huntress et al. is mentioned below as Scheme-12:

### Willgerodt-Kindler reaction

The Willgerodt rearrangement or Willgerodt reaction is an organic reaction which converting an aryl alkyl ketone to the corresponding amide by reaction with ammonium polysulfide, named after Conrad Willgerodt. The related Willgerodt-Kindler reaction takes place with elemental sulfur and an amine like morpholine. The reaction is name after Karl Kindler. The Willgerodt Reaction allows the synthesis of amides from aryl ketones under the influence of a secondary amine and a thiating agent. The reaction mainly relates to the preparation of...
The reaction scheme of Willgerodt-Kindler reaction is mentioned below as Scheme-13a & 13b:

The reaction mechanism of Willgerodt-Kindler reaction involves the formation of an enamine which undergoes thiation, and the carbonyl group migrates to the end of the chain via a cascade of thio-substituted iminium-aziridinium rearrangements (27).

Bolm et al. (28) have disclosed that Willgerodt-Kindler reaction is an efficient method for the preparation of (thio) amidines from a range of substrates and additionally allows the terminal functionalisation of arylalkyl ketones. Probably due to concerns regarding both the yield and the complex reaction mixtures the Willgerodt-Kindler reaction has, to date, been relatively under utilised in organic synthesis. Bolm et al. have mainly disclosed the use of microwave techniques for the preparation of thio-amides using Willgerodt-Kindler reaction and proved that microwave techniques is application in organic synthesis and be no longer plagued by the poor reputation it has held in the past. Moreover, thioamides have been employed as valuable intermediates in the preparation of S- and N-heterocycles. To this end, the Willgerodt-Kindler reaction has also shown potential for the preparation of numerous sulfur-containing heterocycles, but this is yet to be fully exploited. Investigations into the preparation of many new sulfur- and/or nitrogen containing heterocyclic compounds under Willgerodt-Kindler reaction conditions are warranted. Although much progress has been made in the development of the Willgerodt-Kindler reaction there is still much work to be done. To allow this reaction process to find wider utility in organic synthesis, the challenge now stands for synthetic chemists to identify conditions that allow the selective functionalisation of certain moieties under Willgerodt-Kindler conditions. In addition, the further development of this reaction process to allow the efficient preparation of thioamides from arylalkyl ketones with much longer alkyl chains would be of significant synthetic utility and as such warrants investigation. The Bolm et al. have synthesized following compounds i.e., compound of formulae VII-XI (Figure 5) by performing Willgerodt-Kindler reaction:

![Figure 5](image)

**Figure 5** Compound derivatives of Willgerodt-Kindler reaction, Formulae 7-11

Feroci et al. (29) have disclosed the synthesis of thiobenzamides from benzaldehydes, elemental sulfur and cyclic secondary amines was carried out by Willgerodt-Kindler reaction following three different protocols, avoiding the harsh conditions usually reported in the literature. Protocol A was solventless, protocol B was in acetonitrile, while protocol C was electrochemical in acetonitrile (using electrogenerated acetonitrile anion). For all protocols the temperature was 6°C to 80°C. In all cases, a stoichiometric amount of reagents was used, with waste lessening with respect to literature syntheses. Good to high yields were obtained by chemical or electrochemical methods, near or in acetonitrile. Advantages and limits of these sustainable syntheses are highlighted. The reaction scheme of Willgerodt-Kindler reaction performed by Feroci et al. is mentioned below generically as Scheme-15a & specifically as Scheme-15b:

Saraei et al. (30) have disclosed a simple and efficient method for the synthesis of α-ketothioamides via the Willgerodt-Kindler reaction is developed. Reactions were carried out between arylglyoxal hydrates, amines and elemental sulfur in water at 80°C to afford corresponding ketothioamides in good to high yields in a short reaction time. The reaction scheme of Willgerodt-Kindler reaction performed by Saraei et al. is mentioned below as Scheme-16:

Moghaddam et al. (31) have disclosed that Aldehydes and aryl alkyl ketones were efficiently transformed to thioamides with the same number of carbon atoms via Willgerodt-Kindler reaction under microwave irradiation in solvent-free conditions and hydrolysis of same thiomides into carboxylic acids. Moghaddam et al. have synthesized following compounds i.e., compound of formulae XII-XVIII (Figure 6) via performing Willgerodt-Kindler reaction:

![Figure 6](image)

**Figure 6** Compound derivatives of Willgerodt-Kindler reaction, Formulae 12-18

Darabi et al. (32) have disclosed the use of Willgerodt-Kindler reaction for reaction of several aryl alkyl ketones with sulfur and morpholine under solvent-free conditions was performed in a domestic microwave oven. Good results including good yield and purity were obtained in a very short reaction time (between 3.5- minutes). Darabi et al. have synthesized following compounds i.e., compound of formula XIX (Figure 7) via performing Willgerodt-Kindler reaction:

![Figure 7](image)

**Figure 7** Compound derivatives of Willgerodt-Kindler reaction, Formula 19
Passerini Reaction

This three-component reaction between a carboxylic acid, a carbonyl compound such as a ketone or aldehyde, and an isocyanide, offers direct access to α-hydroxy carboxamides (33). The reaction scheme of Passerini Reaction is mentioned below as Scheme-17:

The Passerini reaction proceeds rapidly if the reaction is performed in aprotic solvents at room temperature. High yields are obtained with high concentrations of the starting materials in the reaction mixture. From these findings, it is assumed that the Passerini Reaction does not follow an ionic pathway. Hydrogen bonding is believed to play a crucial role in the formation of the presumed cyclic transition state for this reaction (34). The reaction mechanism of Passerini reaction is mentioned below as Scheme-18:

Schreiber et al. (35) have synthesized amide analogues by performing catalytic asymmetric Passerini reaction using trisubstrate indan (pybox) Cu(II) Lewis acid complex 4 with substrates capable of bidentate coordination. The reaction occurs via ligand-accelerated catalysis. Schreiber et al. have reacted different aliphatic and aromatic aldehydes, acids and isocyanides to form required amide analogues. In all cases, they were able to obtain the desired product in fair to excellent yields and enantioselectivities. The reaction scheme of Passerini reaction performed by Schreiber et al. is mentioned below as Scheme-19:

Andrade et al. (36) have α-Acyloxy carboxamides were easily assessed in one step by the multicomponent reaction between carboxylic acids, aldehydes and C-isocyanides (Passerini reaction) using ionic liquids or polyethylene glycol (PEG 400) as green reaction media. The reaction scheme of Passerini reaction performed by Andrade et al. is mentioned below as Scheme-20:

Zhu et al. (37) have discloses the alcohols instead of aldehydes were used in the Passerini three-component reaction under catalytic aerobic conditions. Mixing alcohols, isocyanides, and carboxylic acids in toluene in the presence of a catalytic amount of cupric chloride, NaNO₂, and TEMPO afforded, under an oxygen atmosphere, the P-3CR adducts in good yields. The reaction scheme of Passerini reaction performed by Zhu et al. is mentioned below as Scheme-21:

Soeta et al. (38) have synthesized a new method for a highly effective addition of isocyanides to aldehydes proceeded smoothly in the presence of a silanol to give the corresponding α-siloxyamides in high yields. A wide range of aldehydes and isocyanides are applicable in this reaction. The reaction scheme of Passerini reaction performed by Soeta et al. is mentioned below as Scheme-22:

It is to be noted that Soeta et al. have developed a direct O-silylative Passerini reaction consisting of aldehydes, isocyanides, and silanols. The said reaction is the first example of the isocyanide-based multicomponent reaction using a silanol instead of a carboxylic acid component, giving the corresponding R-siloxyamides in high yields. A wide range of aldehydes and isocyanides are applicable to this reaction.

Grimaud et al. have (39) discloses a three-component addition of isocyanides to phenol derivatives and aldehydes in methanol forms O-arylated compounds in a new Passerini-type reaction. The key step is an irreversible Smiles rearrangement of intermediate phenoxymidate adducts. The reaction scheme of Passerini reaction performed by Grimaud et al. is mentioned below as Scheme-23:

UGI reaction

The UGI reaction is a multi-component reaction in organic chemistry involving a ketone or aldehyde, an amine, an isocyanide and a carboxylic acid to form a bis-amide (40). The reaction is named after Ivar Ugi, who first reported this reaction in 1959. The reaction scheme of UGI reaction is mentioned below as Scheme-24:

The Ugi reaction is exothermic and usually complete within minutes of adding the isocyanide. High concentration (0.5M - 2.0M) of reactants gives the highest yields. Polar, aprotic solvents, like DMF, work well. However, methanol and ethanol have also been used successfully. This uncatalyzed reaction has an inherent high atom economy as only a molecule of water is lost and chemical yield in general are high. Recent research has shown that the Ugi reaction is accelerated in water.

In the reaction mechanism of Ugi reaction, the initial reaction is the formation of an imine [82] from the amine and the ketone. Subsequent reaction of the imine with the isocyanide and the carboxylic acid gives intermediate [84], which rearranges via an acyl transfer into the bis-amide [85]. The exact mechanism of the trimolementary reaction to form intermediate [84] is not known (41). The reaction mechanism of UGI reaction is mentioned below as Scheme-25:

Grimaud et al. (40) have disclosed a novel and very efficient four-component reaction from readily available substrates. The creation of four bonds that should be reasonably resistant to hydrolysis makes this process particularly attractive for the design of pharmaceutical and agrochemical libraries. Moreover, a straightforward elaboration of these adducts gives easy access to various heterocyclic structures. Further studies are under way to examine more thoroughly the scope and limitations of this MCR system. The reaction scheme of UGI reaction performed by Grimaud et al. is mentioned below as Scheme-26:

Tye et al. (42) have disclosed a utility of a Design of Experiments (DoE) approach for the rapid and efficient optimisation of a microwave assisted UGI reaction of levulinic acid is demonstrated. DoE methods have also been applied to the assessment of the reaction scope for a range of amine and isonitrile substrates. The optimal procedure developed using this approach has enabled the preparation of lactam derivatives in moderate to excellent yields (17-90%) in a reaction time of only 30 min compared to the conventional methodology which required up to 48 h. The reaction scheme of UGI reaction performed by Tye et al. is mentioned below:

Kazmaier et al. (43) have disclosed an Endothiopeptides can easily be obtained via Ugi reaction using thio acids as acid components. If isonitriles with an acetal group are applied, the endothiopeptides can directly be converted into thiazoles using TMSCl-Nal under microwave irradiation. In conclusion Kazmaier et al. have shown that the application of thio acids in Ugi reactions gives rise to endothiopeptides in one step with the option of combinatorial synthesis. If suitable isonitriles are used, these endothiopeptides can be converted into

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peptidic thiazoles also in one step under microwave irradiation. The reaction scheme of UGI reaction performed by Kazmaier et al. is mentioned below as Scheme-28a & 28b:

![Scheme 1]

![Scheme 2]

Suginome et al. (44) discloses a variety of secondary amines have become utilized in the Ugi reaction by using aminoborane as an iminium ion generator. Aldehydes, secondary amines, and isocyanides are coupled in the presence of aminoborane at room temperature, giving the corresponding amine amides in good yields. The nonacidic reaction conditions are beneficial for unique chemoselectivity, where the aldimine functionality is left intact in the present Ugi-type reaction. The reaction scheme of UGI reaction performed by Suginome et al. is mentioned below as Scheme-29:

![Scheme 29]

Orru et al. (45) have discloses a development of 2-isocyanopyridines as novel convertible isocyanides for multicomponent chemistry. Comparison of representatives of this class revealed 2-bromo-6-isocyanopyridine as the optimal reagent in terms of stability and synthetic efficiency. It combines sufficient nucleophilicity with good leaving group capacity of the resulting amide moiety under both basic and acidic conditions. To demonstrate the practical utility of this reagent, an efficient two-step synthesis of the potent opioid carfentanil is presented. The reaction scheme of UGI reaction performed by Orru et al. is mentioned below as Scheme-30:

![Scheme 30]

Sung et al. (46) have discloses a modified U-4CR reaction, which has been done by using commercially available 2-nitrobenzylamine as an ammonia equivalent and it involves a multi-component reaction, followed by photochemical cleavage of the 2-nitrobenzyl group, which can be done in one pot with good yields. The reaction scheme of UGI reaction performed by Sung et al. is mentioned below as Scheme-31:

**DISCUSSION AND CONCLUSION**

The present review article includes an overview of well-known name reactions like Beckmann rearrangement, Schmidt reaction, Passerine reaction, Willgerodt-Kindler reaction and UGI reaction, which contains amide group formation in nucleus of synthesized organic compounds.

Number of organic and a natural product which contains peptide linkage are possessing interesting biological activities. For this reason, the development of new peptide linkage containing compounds by using name reactions has become a most fascinating field of research for many organic chemists with various backgrounds. It is believed that this review presents a systematic overview of use of well-known name reactions, which contains peptide linkage serves as an excellent guideline for the organic synthesis of bioactive molecules bearing peptide linkages.

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Scheme 6) Preparation of amide derivatives from oxime derivatives by Ramon et al

Scheme 7) Reaction scheme of Schmidt reaction

Scheme 8) Reaction mechanism of Schmidt reaction

Scheme 9) Preparation of amide derivatives from ketone by Audhe et al

Scheme 10) Preparation of amide derivatives from ketone by Iyengar et al

Scheme 11) Preparation of amide derivatives from ketone by Jia et al

Scheme 12) Preparation of amide derivatives from acid derivatives by Huntress et al

Scheme 13) (A) Reaction scheme of Willgerodt-Kindler reaction

Scheme 13) (B) Reaction scheme of Willgerodt-Kindler reaction

Scheme 14) Reaction mechanism of Willgerodt-Kindler Reaction

Scheme 15) (A) Generic preparation of thiamide derivatives from aldehyde by Feroci et al

Scheme 15) (B) Specific preparation of thiamide derivatives from aldehyde by Feroci et al
Scheme 16) Preparation of Thiamide derivatives from Arylglucal hydrate by Saraei et al

Scheme 17) Reaction scheme of Passerini reaction

Scheme 18) Reaction mechanism of Passerini reaction

Scheme 19) Preparation of amide derivatives from reaction between acid, aldehyde and cyano derivatives by Schreiber et al

Scheme 20) Preparation of amide derivatives from reaction between acid, aldehyde and cyano derivatives by Andrade et al

Scheme 21) Preparation of amide derivatives from reaction between acid, aldehyde and cyano derivatives by Zhu et al

Scheme 22) Preparation of amide derivatives from reaction between acid, aldehyde and cyano derivatives by Soeta et al

Scheme 23) Preparation of amide derivatives from reaction between acid, aldehyde and cyano derivatives by Grimaud et al

Scheme 24) Reaction scheme of UGI reaction

Scheme 25) Reaction mechanism of UGI reaction

Scheme 26) Preparation of amide derivatives from reaction between aldehyde, amine, cyano and ester derivatives
Scheme 27) Preparation of amide derivatives from reaction of levulinic acid, amine and cyano derivatives by Tye et al

Scheme 28) (A) Preparation of amide derivatives from reaction between thioacid, amine, aldehyde and ester derivatives by Kazmaier et al

Scheme 28) (B) Preparation of amide endo-thiopeptides by Kazmaier et al

Scheme 29) Preparation of amide derivatives from reaction between aldehyde, Secondary amines and isocyanides in the presence of aminoborane by Suginome et al

Scheme 30) Preparation of amide derivatives from convertible Isocyanide by Orru et al

Scheme 31) Preparation of amide derivatives from reaction between acid, aldehyde, amide and cyano derivatives by Sung et al

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