Finding sustainable and environmentally friendly approaches to products is one of the major challenges facing chemists. Some important developing methodologies for producing target compounds in fewer steps with less waste are catalyzed multicomponent coupling reactions, which allow access to structurally complex compounds in a single step.

In one project, our group is developing titanium-catalyzed multicomponent coupling procedures to make nitrogen-based heterocycles either in a single pot or in a single step. Titanium catalysis is advantageous in that the metal is both abundant and nontoxic.

In the exploding diagram are some of the procedures developed for heterocyclic synthesis. These new protocols are applied to natural product synthesis and investigated for their biological activity. For example, with the Tepe group, we have discovered a new class proteasome inhibitor based on the quinoline core structure with potential applications in inflammatory disease and cancers like multiple myeloma.

To evaluate ligands for early transition metal catalysis, like in the project above, we have developed a chromium(VI), \( d^2 \)-system that is very synthetically versatile, \( \text{NCr}(\text{NPr}^i_2)_2X \), where \( X \) is the ligand under scrutiny. Using this system, we parameterize ligands based on their stericus and electronics. In one application, we were able to model the reactivity of a series of titanium hydroamination catalysts and determine quantitatively how the stericus and electronics of the ancillary ligand contributes to reaction rate. Once the model was established, we could anticipate what the reaction rate would be in many cases. This methodology is not being applied to a variety of different catalytic reactions.

In these projects, and others, we are attempting to widen and optimize the applications of transition metals, and we are investigating new possibilities for applications in human health and other areas.

Selected Publications


