Differentiation of Epoxide Enantiomers in the Confined Spaces of an Homochiral Cu(II) Metal-Organic Framework by Kinetic Resolution

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Abstract: TAMOF-1, a homochiral metal-organic framework (MOF) constructed from an amino acid derivative and Cu(II), was investigated as a heterogeneous catalyst in kinetic resolutions involving the ring opening of styrene oxide with a set of anilines. The branched products generated from the ring opening of styrene oxide with anilines and the unreacted epoxide were obtained with moderately high enantiomeric excesses. The linear product arising from the attack on the non-benzylic position of styrene oxide underwent a second kinetic resolution by reacting with the epoxide, resulting in an amplification of its final enantiomeric excess and a concomitant formation of an array of isomeric aminodiols. Computational studies confirmed the experimental results, providing a deep understanding of the whole process involving the two successive kinetic resolutions. Furthermore, TAMOF-1 activity was conserved after several catalytic cycles. The ring opening of a meso-epoxide with aniline catalyzed by TAMOF-1 was also studied and moderate enantioselectivities were obtained.

Introduction

Metal-organic frameworks (MOFs) are porous coordination polymers composed of organic linkers and metal centers or clusters. With high porosities and thermal stabilities, these materials have captured the attention of many researchers due to their remarkable[1] chemical and physical versatility. They have been used as gas storage[2] and (chiral) separation materials,[3,4] sensors,[5] LEDs,[6] drug delivery materials[7] and catalysts,[8] among other applications.

Homochiral MOFs may be obtained from enantiopure precursors. Well-known enantiopure linkers have been used in the construction of homochiral MOFs, such as BINOL- or salen-based porous materials (BINOL = 1,1’-binaphthalene)-2,2’-diol; salen = 2,2’-[(1,1’E,1’E)-ethane-1,2-diylbis(azaneylylidene)]bis-(methaneylylidene)ditophenol).[9] These MOFs, possessing rigid structures and tunable pores, have been exploited in asymmetric catalysis and chiral separations, two areas of high interest in the pharmaceutical industry. Homochiral MOFs have already been used as enantioselective catalysts for directing the preferential formation of a stereoisomer from a pro-chiral substrate (for instance, in the cyanosilylation of aldehydes,[10] epoxidation of alkenes,[11] additions to carbonyl groups,[12] cyclopropanations,[13] and Diels-Alder reactions,[14] aldol reactions,[15] and the ring opening of meso-epoxides with nucleophiles[16]). Kinetic resolution is also an interesting tool in enantioselective catalysis for the generation of enantiopure (or highly enantioenriched) products.[17] The success of a catalytic kinetic resolution relies on distinct reaction rates occurring in the presence of a catalyst for the reaction between the two enantiomers of the starting material.
and an additional reactant/reagent. There are very few examples in the literature of kinetic resolutions of epoxides with homochiral MOFs as catalysts. Moreover, these examples mainly comprise the reaction of oxygen-containing nucleophiles and epoxides outside the confined spaces of the MOFs.\textsuperscript{[19,20]}

Our research group has recently reported a new homochiral MOF, \textit{TAMOF-1} (Figure 1), constructed from copper(II) and a linker prepared from natural L-histidine. The strong metal-nitrogen coordination bonds of \textit{TAMOF-1} produce a structurally robust porous material, with a very high permanent porosity (ca. 1200 m\textsuperscript{2}g\textsuperscript{-1}) and excellent thermal and chemical stabilities. \textit{TAMOF-1} has been used as a chiral stationary phase in chromatographic separations for the quantitative separation of racemic mixtures of drugs (for instance, \textit{rac}-ibuprofen and \textit{rac}-thalidomide), as well as in HPLC columns for highly efficient analytical separations of racemic mixtures. The 3D intersected channels with a diameter $\leq$ 10 Å and the internal binding sites for chiral molecular recognition led to the fabrication of in-house HPLC columns that outperformed several commercial HPLC columns. The interesting properties of \textit{TAMOF-1}\textsuperscript{[21]} (i.e., capability of chiral recognition inside wide channels combined with broad tolerance to solvents) prompted us to study its catalytic activity in kinetic resolutions within confined spaces. Herein, we describe our interesting findings for the kinetic resolution of styrene oxide with a set of anilines in the presence of \textit{TAMOF-1}. The unreacted epoxide and products generated from the ring opening at the benzylic position were kinetically resolved in an efficient way. Computational studies provided a deep understanding of the whole process, based on the different reactivity of the substrates when confined in the helical channels of \textit{TAMOF-1}.

### Results and Discussion

#### Reaction conditions for \textit{TAMOF-1}-catalyzed ring openings of epoxides with anilines

We first explored the resolution of racemic styrene oxide (\textit{rac}-1) with aniline (2) to assess the potential of \textit{TAMOF-1} as a catalyst in the corresponding ring-opening reaction (Scheme 1). The ring opening of \textit{rac}-1 with 2 can generate four different products arising from the nucleophilic attack on the benzylic methine carbon of each enantiomer of the epoxide (branched products, \textit{B1}) or on the non-benzylic methylene carbon (linear products, \textit{L1}). We used standard screening conditions (10 mol\% of catalyst and equimolar amounts of the reagents) in different solvents at different temperatures (Table 1, Supporting Information). Control experiments in the absence of the catalyst revealed that ring opening did not take place to a measurable extent. In the presence of \textit{TAMOF-1}, the ring-opening reaction proceeded smoothly, yielding variable amounts of the four possible products depending on the solvent used and the reaction temperature. In all cases, ring opening took place mostly by the nucleophilic attack on the benzylic carbon and involved the formation of \textit{B1}. Conversions of the ring opening with solvents such as toluene, tetrahydrofuran and ethyl acetate were poor, both at low and high temperatures. Consequently, these solvents were not used any further in our studies. Acetonitrile (ACN) was an adequate solvent, with conversions ranging from 26 \% to 84 \% in the temperature range of 25 °C to 100 °C, respectively. As expected, the enantiomeric excesses (ee's) of the branched and linear products increased by lowering the temperature. The best ee’s with ACN were found at rt (36.6 \% for (\textit{R})-\textit{B1} and 15.2 for (\textit{S})-\textit{L1}), although the ee's at...
40°C were quite similar (33.4% for (R)-B1 and 7.1% for (S)-L1). As 
the conversion was significantly higher at 40°C than at rt (53% 
vs. 26%), further studies were performed in ACN at 40°C, as 
these reaction conditions provided a good balance between 
conversion and stereoselectivity.

At this point of the study, the effects of the epoxide/amine 
ratio were also assessed (Table 1, Supporting Information, 
entries 17–18). There were no effects on the regioselectivity or 
enantioselectivity of the final products. Thus, the concentrations 
indicated above were used in subsequent catalytic studies. 
Other control experiments were performed to determine 
whether racemization of the starting materials or products was 
taking place. These processes were studied by reacting 
enantiopure (R)-1 or (S)-1 separately with aniline (2) in the 
presence of TAMOF-1 under the same reaction conditions as 
those in the catalytic assays. The products obtained were 
diastereomerically and enantiomerically pure, as expected for 
an epoxide ring opening process through a stereospecific S

In order to expand the scope of this reaction, 10 mol% of 
TAMOF-1 was used as the catalyst in the reaction of rac-1 and 
ø-anisidine 3, with similar results to those reported for aniline 
being observed (51% conv.; 60% ee for (R)-4b and 35% ee for 
(S)-4l; see Scheme 2 and Table 2, Supporting Information, for 
details). In the reaction of the dialkyl-substituted meso-epoxide 
5 and aniline with 10 mol% of TAMOF-1 as catalyst, lower 
reactivity and stereoselectivity were observed than those for 
the phenyl-substituted analogue 1 (15% conversion and 35% 
ee in favor of product (15,25)-6; see Scheme 2 and Table 3 in 
the Supporting Information for details).

Monitoring of the kinetic resolution

Using the optimized conditions, we studied the kinetic 
resolution of rac-1 by measuring conversions and enantiomeric 
excesses at different reaction times (4, 12, 17 and 48 h). The 
results are summarized in Figure 2. The concentrations of 1 
and 2 decreased rapidly, yielding B1 as the major product and small 
amounts of L1. Regarding the stereoselectivity of the kinetic 
resolution, the ee of the unreacted starting material increased 
with the conversion, whilst the ee of the products dropped as 
the conversion increased. These trends are in agreement with 
what is expected in a kinetic resolution.19 Ring opening of the 
(R)-enantiomer of 1 with aniline proceeded at a lower rate 
when using TAMOF-1 as the catalyst, with the concentration of
(R)-1 increasing with time. As shown in Figure 2, an ee of 25.3 % in favor of (R)-1 was recorded after 48 h. As (S)-1 reacts faster with aniline, the major product formed was the branched product derived from (S)-1 by the attack of aniline on its benzylic carbon with an inversion of the configuration at the attacked carbon and retention at the other.

Consequently, the branched product (R)-B1 was mostly formed and its absolute configuration was confirmed by comparing with previous data. The ee for this compound after four hours was 34.3 %. The product arising from the attack of aniline on the non-benzylic carbon of the two enantiomers of the starting epoxide (L1) was also observed in the reaction mixture. However, L1 was formed in lower amounts compared to B1. A comparison of the yields of B1 and L1 as a function of time (Figure 2a) shows that the yield of L1 was lower than that for B1 throughout the whole reaction coordinate. The stereoselectivity of L1 after four hours was slightly lower than that for B1 (ee of 20.0 % for (S)-L1 and 34.3 % for (R)-B1). This indicates a slightly lower degree of stereodifferentiation of the two enantiomers of the starting material by TAMOF-1 for the non-benzylic position. Surprisingly, an inversion of the configuration of the product L1 was observed at long reaction times (formation of (R)-L1 with an extended ee of 9.1 %). This finding was not consistent with a simple kinetic resolution. To further understand the stereochemical outcome of this reaction, we performed analogous experiments with increased TAMOF-1 amounts (from 10 mol% to 50 mol%) and extended reaction times. The results of this study are summarized in Figure 3. With the new reaction conditions, the conversion of the ring opening increased up to 97 % after 96 h.

Despite both the ee of the unreacted starting material and the stereoselectivity pattern of B1 as a function of time being in agreement with a standard kinetic resolution, the final configuration of L1 using 50 mol% of TAMOF-1 was again opposite to the one expected for ring opening at the non-benzylic position of the fastest reacting enantiomer of 1 with 2, as observed with 10 mol% of TAMOF-1. Analyses in an HPLC system equipped with an atmospheric pressure chemical ionization detector in the positive mode (APCI+) of crude reaction mixtures identified unexpected reaction products, which were consistent with adducts formed from one molecule of aniline and two molecules of styrene oxide. These findings suggested that a second ring-opening reaction was taking place, since the products arising from the first ring-opening (i.e., B1 and L1) still had the ability to attack a second molecule of styrene oxide to form dialkyl aryl tertiary aminodiols. It should be noted that the ring opening of a second unit of a racemic epoxide, at the benzylic and non-benzylic positions, by amino alcohols (R)-B1, (S)-B1, (R)-L1 or (S)-L1 may generate up to 10 diastereoisomeric aminodiols, whose structures are shown in Figure 4. Interestingly, five peaks with [M + H]+ ions that were consistent with the suggested aminodiol structures were observed. In particular, LB1, LL1, LB2 and LL2 were identified in the reaction mixtures by comparative HPLC analyses of the crude reaction mixtures and pure samples of these compounds.

In addition, the aminodiol BB1 arising from the ring opening of the fastest reacting enantiomer (S)-1 by the most
abundant amino alcohol (R)-B1 was unequivocally identified in the reaction mixture by comparative HPLC analysis.[24]

Matched double catalytic kinetic resolutions via subtractive Horeau amplification

Intrigued by the origin of the inversion of the configuration of L1 at high conversions, we looked closely at the concentration profile of both enantiomers of L1 vs. time (see Figure 3b). At short reaction times, there was an excess of (S)-L1 with respect to (R)-L1. As previously indicated, formation of (S)-L1 in the presence of TAMOF-1 was slightly preferred in the reaction between rac-1 and 2. However, (S)-L1 reacted faster than (R)-L1 with styrene oxide to form the corresponding aminodiol. This can be seen in Figure 3b, where the concentration of (S)-L1 decreased with time (dotted curve in Figure 3b), whilst that of (R)-L1 reached a plateau and remained quite constant (dashed line in Figure 3b). As (S)-L1 was abstracted from the reaction media forming LL1, the whole process led to the formation of L1 with an increased ee of 57.9% in favor of the (R)-enantiomer (orange curve in Figure 3c). These observations were confirmed by performing the kinetic resolution of rac-L1 with rac-1 in the presence of TAMOF-1 (Figure 5). As shown in this figure, the kinetic resolution of rac-L1 led to (R)-L1, leaving (S)-1 as the unreacted enantiomer of the starting epoxide. It should be noted that (R)-1 is the fastest reacting enantiomer of the epoxide with the amino alcohol (S)-L1, whilst the other enantiomer of styrene oxide is the fastest reacting enantiomer with aniline. It is also interesting to note the chiral amplification effect observed in the formation of (R)-L1 (from an ee of 20.0% with one kinetic resolution to 57.9% after two kinetic resolutions). The most abundant enantiomer of the linear product formed after one kinetic resolution (i.e., (S)-L1) reacted faster through a second kinetic resolution process than (R)-L1, leading to an accumulation of the latter (by abstraction of the former). Overall, through two matched kinetic resolutions, the stereoselectivity of the formation of (R)-L1 is enhanced. These results constitute an interesting example of subtractive Horeau amplification within the pores of TAMOF-1 (see Scheme 3 for a summary of the processes involved).[25]

To demonstrate that kinetic resolutions were occurring inside the pores of TAMOF-1, reactants with more demanding steric profiles were used (i.e., racemic trans-2,3-diphenyloxirane instead of its mono-substituted analogue rac-1 and tert-butylamine instead of aniline). Interestingly, kinetic resolution of trans-2,3-diphenyloxirane and aniline under identical reaction conditions to those used for its monophenyl-substituted analogue rac-1 did not take place at any measurable extent. In an analogous manner, the kinetic resolution of styrene oxide with tert-butylamine did not proceed either. These results confirmed that kinetic resolution processes take place inside the pores of TAMOF-1.

Reaction mechanism

DFT calculations have been employed to investigate the reaction pathways leading to the experimentally observed amino alcohol and aminodiol products in both kinetic reso-
The reaction mechanism for the TAMOF-1-catalyzed ring opening of both (S)-1 and (R)-1 styrene oxides by aniline (2) (Figure 6) to produce (R)/(S)-L1 or (R)/(S)-B1 has been explored. The reaction starts by the coordination of the epoxide (Figure 6) to produce (S)-1, the distance between the oxygen and the copper is around 2.4 Å. In intermediates I R and I S, the distance between the oxygen and the copper is around 2.4 Å. In addition, other weak contacts contribute to the stabilization of these structures (e.g., the hydrogen atoms of the epoxide coordination to the copper, as their C–H–C distances remain practically the same in the free and copper-bound styrene oxides). Subsequent aniline attachment to one of the remaining two copper atoms leads to the formation of intermediates II R/II S. The C–N distance in these complexes is around 2.35 Å and, although one of the Cu–H epoxide contacts is lost, the other one is maintained, helping to stabilize these intermediates. This coordination process is also exothermic, and the associated energy gain varies depending on which enantioomer of styrene oxide is already present within TAMOF-1. Our results indicate that the binding of aniline is ca. 12 kJ mol⁻¹ stronger in the presence of (R)-styrene oxide, indicating that the MOF differentiation for both styrene oxide enantiomers takes place in the presence of aniline. After the addition of aniline, the whole cavity is practically full and both substrates lie quite close to each other; additional contacts, shorter than 2.6 Å, are established between the styrene oxide and aniline in both II R and II S.

Nucleophilic attack of 2 onto the epoxide takes place producing either the linear or the branched product. The energy barriers leading to the latter are lower for both styrene oxides by 2.2 and 2.8 kJ mol⁻¹ for the (R)-B1/(S)-L1 and (S)-B1/(R)-L1 pairs, respectively. Although these energy differences are not very large, they are enough to favor the formation of one regioisomer over the other in agreement with the experimental observations, which state that the branched products (B1) are favored. The energy barrier heights range between 98.2 and 102.7 kJ mol⁻¹, indicating that the reaction should not be very fast, as observed experimentally. It should be noted that the S2 barriers for the reaction of (S)-1 with aniline are lower than for its (R)-1 counterpart: 98.2 and 100.5 kJ mol⁻¹ for the branched products, and 101.1 and 102.7 kJ mol⁻¹ for the linear products. These differences in barrier heights are responsible for the observed enantioselectivity, with products (R)-B1 and (S)-L1 being preferably formed. The intramolecular proton transfer steps to transform the zwitterionic amino alcohols into B1 and L1 have not been studied, since they are expected to be fast processes not determining the reaction outcome. The release of the final product is exothermic in all cases, except for (S)-B1. In this case, this product should be easily displaced by an incoming styrene oxide or aniline molecules, which would enable the catalytic process to proceed smoothly. As observed, TAMOF-1 favors the formation of the branched products B1 over their –more stable– linear counterparts L1, as expected for a transformation that takes place under kinetic control.
The transition states for the nucleophilic attack of aniline onto R-1 and S-1 have also been computed without TAMOF-1. In these cases, the energy barriers leading to the branched and linear products are as high as 155.3 and 148.9 kJ mol\(^{-1}\), respectively, around 50 kJ mol\(^{-1}\) higher than those computed within TAMOF-1. Interestingly, the uncatalyzed reaction favors the formation of the linear products, which are in all cases the preferred thermodynamic products (see Figure 7 and Figure 8).

These results indicate that the copper atoms in TAMOF-1 cooperatively mediate the formation of the products in the first kinetic resolution, probably by bringing styrene oxide and aniline close together inside the TAMOF-1 cavities. This effect has been already observed in reactions with supramolecular catalysts such as cucurbiturils, where the encapsulation of the reactants largely reduces the entropic cost of bringing reagents together and enhances reaction rates.\(^{23}\) In addition, the coordination of one aniline and one styrene oxide molecules within TAMOF-1 has a positive accumulation effect on the reaction rate; the formation of species II-S and II-R is energetically favored over other possible non-productive bimolecular arrangements containing either one aniline and one acetonitrile, one styrene oxide and one acetonitrile, and two aniline molecules.\(^{26}\) Therefore, TAMOF-1 has two different functions in this process: first, it lowers the reaction barrier leading to the formation of all products, and second, allows for the preferential formation of the kinetic products over their thermodynamic counterparts, in contrast to the uncatalyzed process. The observed final configuration of L1 enantiomers at long reaction times, producing a second kinetic resolution of aminodiol, has also been computed. In particular, we have studied the TAMOF-1-catalyzed reaction between (R)/S-L1 intermediates and (R)/S-1 to deliver the linear products LL1, LL2, and their enantiomers, which constitute the most abundant products at this stage (Figure 7). The second kinetic resolution reaction starts by coordination of styrene oxide to one of the copper atoms in TAMOF-1 with this process being exactly the same than that found in the first kinetic resolution.

The reaction should subsequently proceed by coordination of L1 products to TAMOF-1. The computed transition states indicate that the fastest reaction is that between (S)-L1 and (R)-1 to produce LL1, as experimentally observed (see Figure 7 and Figure 8). This reaction has a barrier of 100.5 kJ mol\(^{-1}\); in contrast, the reaction between (S)-L1 and (S)-1 to produce LL2...
has a larger barrier of 112.9 kJ mol\(^{-1}\), confirming the preference of (S)-L1 to engage with (R)-1.

These results indicate that, under experimental conditions, (S)-L1 would engage faster in a second C–N bond formation process, hence producing the accumulation of (R)-1 and the reversal of the expected enantioselectivity for these amino alcohol intermediates at long reaction times.

The energy barriers for the formation of LL1 and LL2 from (S)-L1 and (R)/(S)-1 have also been computed in the absence of TAMOF-1. The reaction of (S)-L1 with (R)-1 and (S)-1 produces high barriers of 150.4 and 161.4 kJ mol\(^{-1}\), respectively. In this case, the reactivity order with and without TAMOF-1 is not altered but it can be observed that the presence of the MOF catalyst drastically reduces the energy barriers for most of the reaction pathways.

Recyclability of TAMOF-1

Recyclability is another important feature for heterogeneous catalysts. To assess the performance of TAMOF-1 as a recoverable and reusable catalyst, a series of reaction cycles were performed. After each run (ACN, 40°C, 16 h), the catalyst was collected through filtration, dried in the oven at 40°C and preferential consumption of (R)-1 was recorded in CDCl\(_3\) using a Solvent Purification System (SPS). Silica gel 60 (230–400 mesh) was used for column chromatography. NMR spectra are quoted in ppm relative to residual solvent peaks. Data for \(^{1}\)H NMR are given as follows: chemical shift, multiplicity (s = singlet, d = doublet, t = triplet, m = multiplet, dd = doublet of doublets, tt = triplet of triplets), coupling constants and intensities. High-resolution mass spectra (HRMS) were recorded by using ESI as

**Conclusion**

In conclusion, TAMOF-1 is a recyclable catalyst for the efficient kinetic resolution of rac-styrene oxide (1) with a set of anilines, yielding secondary amino alcohols. The linear products arising from the attack on the non-benzylic position underwent a second kinetic resolution, resulting in an amplification of its final enantiomeric excess and a concomitant formation of an array of isomeric aminodiols (subtractive Horeau amplification effect). The ring opening of a meso-epoxide with aniline catalyzed by TAMOF-1 was also studied and moderate enantioselectivities were obtained. Computational studies on the whole process have provided deep insight into the ring opening steps and stereochemical outcome of the sequential kinetic resolution processes. These studies have revealed that TAMOF-1 lowered the reaction barriers of all ring opening processes taking place inside channels. The calculated energy barriers of the reactions leading to the ring opening products were in complete agreement with the experimental observations: preferential formation of (R)-B1 in the first kinetic resolution and preferential consumption of (S)-L1 in the second kinetic resolution (with concomitant amplification of the chirality for the unreacted (R)-L1 amino alcohol). The regioselectivity and enantioselectivity of TAMOF-1 remained constant throughout five consecutive reaction cycles.

**Experimental Section**

**Materials and general**

All syntheses were carried out on chemicals as purchased from commercial sources, unless otherwise indicated. Air and moisture sensitive manipulations or reactions were run under inert atmosphere using anhydrous solvents and Schlenk techniques. All solvents were either purchased commercially anhydrous or dried by using a Solvent Purification System (SPS). Silica gel 60 (230–400 mesh) was used for column chromatography. NMR spectra were recorded in CDCl\(_3\), as solvent on a Bruker Advance 300, 400 or 500 MHz Ultrashield spectrometers. \(^{1}\)H and \(^{13}\)C NMR chemical shifts are quoted in ppm relative to residual solvent peaks. Data for \(^{1}\)H NMR are given as follows: chemical shift, multiplicity (s = singlet, d = doublet, t = triplet, m = multiplet, dd = doublet of doublets, tt = triplet of triplets), coupling constants and intensities. High-resolution mass spectra (HRMS) were recorded by using ESI as
ionization method in positive mode. IR spectra were recorded using Attenuated Total Reflection (ATR) spectroscopy. Specific optical ionization method in positive mode. IR spectra were recorded using a Jasco P1030 Polarimeter with a wavelength of 589 nm (sodium, D line) at 25 °C in the corresponding solvent, in a 10 cm cell, unless otherwise indicated. Crude mixtures of the kinetic resolution or rac-1 were analyzed by HPLC chromatography with an Agilent 1200 Series system equipped with a Chiralpak IC column (4.6 x 250 mm) and a diode array detector, at a flow rate of 1 mL/min and a gradient solvent system of Hex/IPA 99:1 to 95:5 (min, %IPA): (0, 1); (7, 1); (14, 5); (40, 5), unless otherwise indicated.

General procedure for the kinetic resolution of rac-1
Prior to start the reaction, TAMOF-1 (10 or 50 mol%) was activated at 130 °C in vacuo for at least 3 h in a Schlenk flask. The Schlenk flask was filled with nitrogen and left to cool at rt, then, a solution of rac-1 (1 equiv.) and the corresponding amine (1 equiv.) in ACN (0.25 M) was added to the Schlenk flask and the reaction mixture was stirred under nitrogen atmosphere at the desired temperature for the required time. The reaction mixture was left to cool at rt, then the suspension was filtered using a syringe filter with PTFE membrane (0.2 µm) and the filter was washed three times with ACN. Crude products of catalytic reactions were analyzed in an Agilent 1200 series HPLC system equipped with diode array detector and quantified using external standards.

**Computational details**

**General procedure for the kinetic resolution of rac-1**
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**(R)-B1: ^1^H NMR (400 MHz, CDCl$_3$) δ 7.41–7.31 (m, 4H), 7.31–7.24 (m, 2H), 6.63–6.55 (m, 2H), 4.51 (dd, J = 7.0, 4.2 Hz, 1H), 3.95 (dd, J = 11.1, 4.2 Hz, 1H), 3.76 (dd, J = 11.1, 7.0 Hz, 1H). ^13^C{^1^H} NMR (101 MHz, CDCl$_3$) δ 147.37 (C), 140.25 (C), 129.30 (CH), 128.99 (CH), 127.77 (CH), 126.87 (CH), 118.05 (CH), 114.00 (CH), 67.53 (CH), 60.02 (CH).

**Keywords**

Keywords: amino acids · chirality · epoxides · kinetic resolution · metal-organic framework

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