



Chiral Recyclable Dimeric and Polymeric Cr(III) Salen Complexes Catalyzed Aminolytic Kinetic Resolution of *trans*-Aromatic Epoxides Under Microwave Irradiation

RUKHSANA I. KURESHY,* K. JEYA PRATHAP, SURENDRA SINGH, SANTOSH AGRAWAL, NOOR-UL H. KHAN, SAYED H.R. ABDI, AND RAKSH V. JASRA

Discipline of Inorganic Materials and Catalysis, Central Salt and Marine Chemicals Research Institute (CSMCRI), Bhavnagar 364 002, Gujarat, India

ABSTRACT Aminolytic kinetic resolution (AKR) of *trans*-stilbene oxide and *trans*- β -methyl styrene oxide proceeded smoothly under microwave irradiation using chiral dimeric and polymeric Cr(III) salen complexes as efficient catalysts, giving regio-, diastereo-, and enantioselective *anti*- β -amino alcohols in high yields (49%) and chiral purity (ee up to 94%) in case of 4-methylaniline within 2 min. The kinetic resolution system is approximately five times faster than traditional oil bath heating at 70°C and 420 times faster than the reaction conducted at room temperature with concomitant recovery of respective chirally enriched epoxides (ee, 92%) in excellent yields (up to 48%). The catalyst **1** worked well in terms of enantioselectivity than the catalyst **2**, but both the catalysts were easily recovered and reused five times with the retention of its efficiency. *Chirality* 19:809–815, 2007. © 2007 Wiley-Liss, Inc.

KEY WORDS: aminolytic kinetic resolution; *trans* epoxides; chiral; Cr(III) dimeric and polymeric salen; recyclable

INTRODUCTION

Optically pure β -amino alcohols are important structural units for many biologically active, natural synthetic products and are valuable chiral auxiliaries/ligands in the area of asymmetric synthesis.^{1–4} Several strategically different approaches exist toward the direct synthesis of β -amino alcohols such as (a) via sharpless osmium catalyzed aminohydroxylation^{5,6} of alkenes (b) direct addition of α -hydroxy ketones^{7–9} to imines, and (c) ring opening of *meso*-epoxides with amines.^{10,11} Though these methods gave products with high enantioselectivity for *syn*- β -amino alcohols, very few methods favored the formation of optically pure *anti*- β -amino alcohols.^{12–19} In this direction, aminolytic kinetic resolution (AKR) of racemic 1,2-disubstituted epoxides with amines as nucleophiles is an attractive method for the synthesis of optically pure *anti*- β -amino alcohols with high diastereoselectivity.^{20–22} Bartoli et al. for the first time has reported the ring opening of *trans*- and *meso*-epoxides with anilines for the synthesis of *anti*/*syn*- β -amino alcohols using chiral monomeric Cr(III) salen as a catalyst, but separation and recycling of the catalyst is not feasible under homogeneous catalytic system.²³ As chiral catalysts are expensive, their separation and recycling is an important aspect. In order to make the catalytic process recyclable, attempts were made in the past for the anchoring of chiral homogeneous catalyst,²⁴ either on solid supports²⁵ or by making use of ionic liquids.²⁶ Though these approaches are interesting but usually demand additional modifications in the structure of the catalyst for making the catalyst compatible with the

desired support, such approaches frequently lead to partial loss of activity and/or enantioselectivity.

The use of microwave (MW) irradiations to speed up the sluggish chemical reactions is of current interest.^{27–31} The use of MW irradiations has been reported for the ring opening of vinyl and terminal epoxides by NH_4OH ,^{32,33} thiols³⁴ pyrazole, imidazole,³⁵ and amines.^{36–38} Yet, its use in the area of transition metal catalyzed asymmetric reactions³⁹ has not been successful, probably because of the small difference in the activation energy of the two enantiomers involved in the asymmetric reaction that is insignificant in comparison to the energy supplied by the MWs.²⁹ Consequently, the enantioselectivity of the reactions could be adversely affected by the use of MWs as a source of energy. Nevertheless, Dicos and Jacobs have recently reported the asymmetric ring opening of epoxides using TMSN_3 as nucleophile under MW irradiation.⁴⁰ In continuation of our ongoing research toward developing the recyclable chiral dimeric and polymeric catalysts with different metal ions for various organic transformations, namely asymmetric epoxidation,^{41–43} cyanation,^{44,45} hydrolytic kinetic resolution,⁴⁶ oxidative kinetic resolution,^{47,48}

Contract grant sponsor: DST and CSIR Network Project on Catalysis

*Correspondence to: Rukhsana I. Kureshy, Discipline of Inorganic Materials and Catalysis, Central Salt and Marine Chemicals Research Institute (CSMCRI), Bhavnagar 364 002, Gujarat, India.

E-mail: rukhsana93@yahoo.co.in

Received for publication 30 May 2007; Accepted 20 July 2007

DOI: 10.1002/chir.20472

Published online 4 September 2007 in Wiley InterScience (www.interscience.wiley.com).

and asymmetric ring opening of *meso* and *trans* epoxides with anilines,^{49,50} in the present study we are reporting the use of recyclable dimeric and polymeric chiral Cr(III)X salen complexes **1** and **2** as catalysts for the production of enantioselective *trans* 1,2 aminoalcohols through AKR of racemic *trans*-stilbene oxide and *trans*- β -methyl styrene oxide, with anilines under MW irradiation as an alternate source of energy to step up the reaction rate. Significantly, the rate of reactions was greatly enhanced and there was no loss in enantioselectivity under MW irradiation in comparison to the same reaction when conducted at room temperature. On the contrary, in some cases there was an improvement in the ee of the products. Chiral dimeric Cr(III) salen complex **1** gave better results in term of enantioselectivity than Jacobsen monomeric Cr(III) salen complex and polymeric Cr(III) salen complex **2**. The catalysts used in the present study were conveniently recovered and reused for five successive catalytic runs.

EXPERIMENTAL

Materials and Methods

Reagents and solvents were of laboratory grade and were used without further purification. *trans*-Stilbene oxide, *trans*- β -methyl styrene, aniline, 2-methoxyaniline, 4-methoxyaniline, 4-methylaniline, 4-chloroaniline, and 4-nitroaniline were purchased from Aldrich and were used as received. Racemic *trans*- β -methyl styrene oxide was prepared by the oxidation of respective alkene with *m*-CPBA (Spectrochem).⁵¹ (*R,R*) Polymeric [Cr(Salen)Cl] **2** was prepared and characterized by the reported procedure.⁵⁰ ¹H and ¹³C NMR spectra were recorded at 200 MHz and 50 MHz, respectively on a Bruker F113V. The chemical shifts (δ) for ¹H and ¹³C are given in ppm relative to the signal of TMS. FTIR spectra were recorded on a Perkin Elmer Spectrum GX spectrophotometer in KBr/nujol mull. Microanalysis of the complex was done on CHNS analyzer, Perkin Elmer model 2400. MW reactor with temperature controller used in the present study was from ETHOS 1600 Advanced Microwave Lab station. Purification of reaction products was carried out by flash column chromatography on silica gel (60–200 mesh). Melting points reported in the present study are uncorrected. Diastereomeric purity and enantiomeric excesses (ee) were determined by NMR analysis of the crude mixture and by HPLC (Shimadzu SCL-10AVP) analysis using Daicel Chiralpak OD and OJ chiral columns at wavelength 243 nm, with 2-propanol/hexane as mobile phase. Optical rotations were measured on a Digipol 781 automatic Polarimeter, Rudolph Instruments and are reported as: α_D^{27} (c = in g per 100 ml, solvent). HPLC traces of aminoalcohols were compared with authentic racemic samples prepared by the ring opening of respective epoxides with required amine using racemic Jacobsen Cr(III) salen as a catalyst. High-resolution mass spectra were obtained with a LC-MS (Q-TOFF) LC (Waters), MS (Micromass) instruments.

All *anti*- β -amino alcohol (**6a–f** and **7a–c**) and *trans*-epoxides (**3** and **4**) were characterized by ¹H and ¹³C NMR, LCMS, and IR. These spectroscopic data are comparable

with literature values.²³ The enantiomeric excess (ee) of *anti*- β -amino alcohol (**6a–f** and **7a–c**) and *trans*-stilbene oxide (**3**) were determined by HPLC using Chiralpak OD column except for **6b**, **7b**, where Chiralpak OJ column was used. The enantiomeric excess of *trans*- β -methyl styrene oxide was determined on GC using GTA-type column.

Synthesis of 5,5-Methylene Di-[(*R,R*)-1-*N*-(3-*tert*-butyl salicylidene)-*N'*-(3',5'-di-*tert*-butyl salicylidene))-cyclohexane-1,2-diaminato (2-)] Cr(III) Chloride **1**

Chiral dimeric ligand namely 5,5-methylene di-[(*R,R*)-{*N*-(3-*tert*-butyl salicylidene)-*N'*-(3',5'-di-*tert*-butyl salicylidene))-1,2-cyclohexanediamine}]^{41,42} (0.4 mmol) was dissolved in dry degassed THF (20 ml). The resulting yellow solution was interacted with anhydrous Cr(II) chloride (0.8 mmol) in a glove box under nitrogen atmosphere to give a brown solution. The brown solution turned dark green during 4 h of stirring under a blanket of nitrogen after which it was exposed to air with stirring for further 3 h. The resulting dark green solution was treated with *tert*-butyl methyl ether to precipitate out the dimeric Cr(III) complex **1**. The precipitate was filtered and washed with water to remove the unreacted chromium chloride and dried overnight under vacuum (yield 65%). The filtrate was sequentially washed with saturated solution of NH₄Cl and brine. The resulting solution was dried over anhydrous Na₂SO₄, filtered, and the solvent was removed to give another crop (~22%) of complex **1** as brown solid. The overall yield was found to be 87%. mp >250°C; Anal. Calcd for C₆₅H₈₈Cl₂Cr₂N₄O₄ (Calcd) C, 67.05; H, 7.62; N, 4.81 Found: C, 67.01; H, 7.60; N, 4.78%; IR in KBr (cm⁻¹): 2946, 2363, 2342, 1617 (C=N), 1536, 1435, 1350, 1316, 1161, 831, 738, 658, 563; [α_D^{25}] = -324 (c = 0.024 g in 100 ml of CH₂Cl₂); Λ_M (MeOH) 116 mho cm⁻¹ mol⁻¹; UV-vis: (CH₂Cl₂) λ_{max} (ϵ) 232 (52030), 263 (32125), 296 (19513), 345 (7689), 435 (3354) nm.

General Procedure for AKR of *trans*-aromatic epoxides

The AKR reactions were carried out in a corning glass test tube placed in a sealed reactor-ETHOS 1600 (Advanced Microwave Lab station) having temperature and MW power output control. In a typical AKR reaction catalysts **1/2** (0.01 mmol, based on monomeric salen unit), appropriate epoxide (0.20 mmol) and aniline (0.10 mmol) were taken in a mixed solvent CH₂Cl₂:MeOH (9:1, 200 μ l), and the mixture was irradiated with MW (900 W) at 70°C for 2 min. The conversion of the product was determined on a HPLC equipped with Chiralpak OD column, using calibration of peak area % of the amino alcohol and *trans*-stilbene oxide at 243. The catalyst was precipitated out from the reaction mixture by the addition of 5 ml of hexane:diethyl ether (1:1), filtered, vacuum-dried, and kept in a desiccator for further use. The filtrate was concentrated and the desired product was purified by column chromatography using silica gel 60–200 mesh as a stationary phase and hexane:ethyl acetate (8:2) as a mobile phase. All the products were characterized by ¹H and ¹³C NMR spectroscopy.

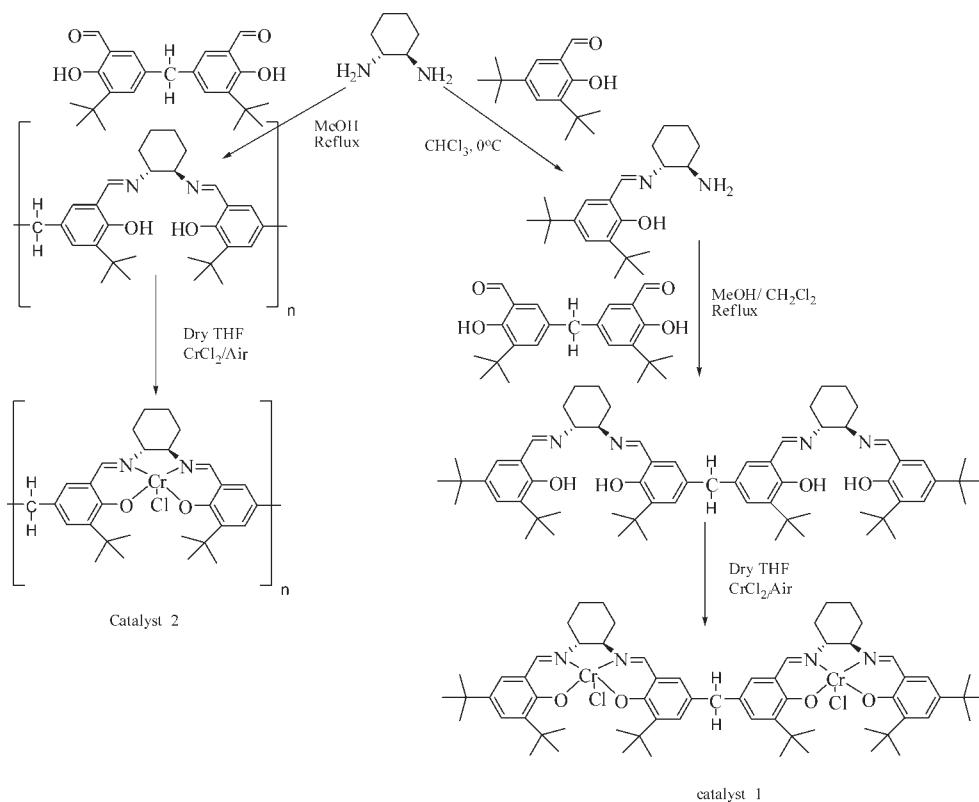
Recycling Experiment

After one catalytic cycle (checked on HPLC), the catalysts were precipitated out with hexane:diethyl ether (1:1) followed by thorough washing with hexane (10 ml). They were dried under reduced pressure and used as recovered catalyst for successive catalytic runs, using *trans*-stilbene oxide as a representative substrate for AKR reaction.

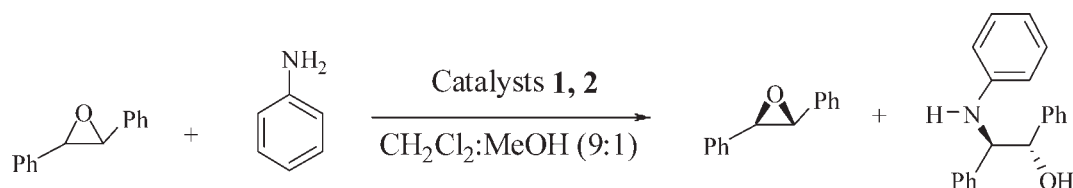
RESULTS AND DISCUSSION

Synthesis of chiral dimeric Cr(III) salen complex, namely, 5,5-methylene di-[(*R,R*)-{*N*-(3-*tert*-butyl salicylidene)-*N'*-(3',5'-di-*tert*-butyl salicylidene)}-1,2-cyclohexanediaminato (2-)] chromium (III) chloride] **1** (Scheme 1), which carries two active catalytic metal centers was obtained by the interaction of 5,5-methylene di-[(*R,R*)-{*N*-(3-*tert*-butyl salicylidene)-*N'*-(3',5'-di-*tert*-butyl salicylidene)}-1,2-cyclohexanediamine with anhydrous Cr(II) chloride in dry THF under nitrogen atmosphere followed by its auto oxidation in air. The characterization data for complex **1** is given in Experimental section. Whereas polymeric Cr(III) salen complex **2** namely poly [(*R,R*)-*N,N'*-bis{3-(1,1-dimethylethyl)-5-methylene salicylidene} cyclohexene-1,2-diaminato-chromium(III) chloride with 12 repetitive units was synthesized by the reported method^{50,52,53} (Scheme 1). Chiral dimeric and polymeric Cr(III) salen complexes **1** and **2** were used as catalysts for the AKR of *trans*-stilbene oxide, with aniline as nucleophile

in a model reaction for optimizing reaction conditions such as temperature and MW output. To begin with we first carried out AKR of *trans*-stilbene oxide with aniline, using complexes **1** and **2** as catalysts over a temperature range of (40–80°C) exposed to 900 W MW irradiation (Table 1, entries 1–10) in CH₂Cl₂:MeOH (9:1) as solvent. From the data in Table 1 it is evident that 70°C is the optimum temperature to achieve the highest yield (49%) with high enantioselection (92%) in the product (entry 7). Next, we studied the effect of MW output (800–1000 W) on the efficacy of AKR keeping the reaction temperature 70°C as constant. However, a MW output of 800 W caused reduction in both yield (32–35%) and enantioselectivity (37–42%) (Table 1, entries 11 and 12). On the other hand, 1000 W MW output resulted in an increase in the yield (72–74) of the product amino alcohol but with lower enantioselectivity (Table 1, entries 13 and 14). Consequently, 900 W MW output and 70°C (Table 1, entries 7 and 8) reaction temperature were found to be ideal for AKR of *trans*-stilbene oxide with aniline. Further, in an ideal situation, 0.5 equiv of amine with respect to 1 equiv of racemic epoxide should react with only one enantiomer of epoxide to give the product *anti*-β-aminoalcohol in 100% ee with a maximum theoretical yield of 50%. However, practically this situation is seldom achievable. It is known in the literature that the equivalence of the nucleophile with respect to racemic epoxide greatly influences the product distribution in a kinetic resolution reaction.⁵⁰ Therefore, we varied the equivalence of nucleophile (aniline, 0.4–0.75 equiv) with respect



Scheme 1. Synthesis of the catalysts **1** and **2**.

TABLE 1. Aminolytic kinetic resolution (AKR) of *trans*-stilbene epoxides using recyclable catalysts, with aniline under different reaction conditions^a

Entry	Temp. (°C)	Time (min)	MW (W)	<i>trans</i> -Epoxide		<i>anti</i> -β-Amino alcohol		<i>s</i> ^b
				ee ^c (%)	Yield ^d (%)	ee ^c (%)	Yield ^d (%)	
1 (2)	40	2	900	78 (73)	78 (83)	94 (81)	22 (16)	36 (10)
3 (4)	50	2	900	81 (74)	62 (65)	94 (80)	35 (30)	39 (10)
5 (6)	60	2	900	84 (72)	63 (62)	92 (78)	42 (37)	30 (9)
7 (8)	70	2	900	90 (72)	48 (48)	92 (78)	49 (49)	31 (10)
9 (10)	80	2	900	81 (69)	35 (37)	92 (67)	65 (62)	32 (7)
11 (12)	70	2	800	35 (25)	65 (67)	42 (37)	35 (32)	3 (2)
13 (14)	70	2	1000	40 (31)	37 (26)	55 (55)	72 (74)	10 (5)
15 (16)	70	2	900	75 (68)	65 (67)	96 (80) ^e	36 (32)	59 (10)
17 (18)	70	2	900	92 (79)	45 (46)	68 (70) ^f	60 (57)	7 (7)
19 (20)	70	2	900	96 (83)	39 (41)	61 (62) ^g	69 (68)	5 (6)
21	70	2	900	30	72	75 ^h	27	8
22 (23)	70	5	–	42 (40)	65 (68)	75 (73) ⁱ	35 (32)	8 (7)
24 (25)	70	10	–	79 (73)	51 (50)	61 (67) ⁱ	48 (49)	5 (6)
26 (27)	27	840	–	85 (80)	47 (48)	93 (87) ^j	49 (49)	36 (18)

^aThe 0.01 mmol catalyst was taken in 200 μl CH₂Cl₂:MeOH (9:1), 0.2 mmol epoxides and 0.1 mmol aniline were added and the reaction mixture was irradiated for 2 min in a microwave oven at 70°C. Data given in parentheses are for catalyst **2**.

^b*s* = selectivity factor = $\ln[1 - c(1 + ee)] / \ln[1 - c(1 - ee)]$, where *ee* is enantiomeric excess of the aminoalcohol product and *c* is the conversion (set to the isolated yield).²³

^cThe diastereoselectivity of product was found to be >99% determined by ¹H NMR and HPLC. The *ee* of amino alcohols and epoxides are determined on Chiralpak OD column and the absolute configuration was assigned by comparing the optical rotation with literature.²³

^dThe conversion of the product **7a** was determined on Chiralpak OD column, using a calibration curve of epoxide and aminoalcohols and rest given as isolated yield.

^eReaction was performed with 0.4 equiv aniline.

^fReaction was performed with 0.6 equiv aniline.

^gReaction was performed with 0.75 equiv aniline.

^hReaction with Jacobsen catalyst under microwave irradiation at 70°C

ⁱReaction carried out at 70°C on oil-bath heating in absence of MW and aliquot was taken at 5 and 10 min, respectively.

^jReaction carried out at 27°C in absence of MW.

to the substrate racemic *trans*-stilbene oxide (1.0 equiv), keeping the other reaction parameters same as mentioned in entry 7 of the Table 1. As the amount of nucleophile was reduced to 0.4 equiv, there was a slight improvement in the *ee* of the product but with significantly lower yield (Table 1, entries 15 and 16). On increasing the amount of nucleophile (0.6–0.75 equiv) there was a steady decrease in the *ee* of product *anti*-β-aminoalcohol with concomitant increase in the *ee* of *trans*-stilbene oxide (83–96%; Table 1, entries 17–20). Hence, the kinetic resolution reaction can be optimized depending upon which product (epoxide or aminoalcohol) is required in higher chiral purity by simply adjusting the amount of nucleophile (aniline) used. For the sake of comparison, we conducted the AKR of racemic *trans*-stilbene oxide with aniline, using monomeric Jacobsen Cr(III)Cl as catalyst (Table 1, entry 21) that gave far less yield (27%) of the product *anti*-β-aminoalcohol (**6a**); however, the *ee* was nearly the same as it was obtained

with polymeric complex **2** but inferior than dimeric Cr(III) salen complex **1** under identical conditions (Fig. 1). Further, when the aforementioned reaction was conducted in the absence of MW irradiation at 70°C on an oil-bath, there was a drop in yield (32–35%) for similar enantioselection in 5 min (Table 1, entries 22 and 23) as compared to the similar reaction under MW irradiation (Table 1, entries 7 and 8). In an attempt to increase the yield by way of increasing the time (10 min) of the reaction, there was a drop in *ee* (61–67%) of the product (Table 1, entries 24 and 25). On the other hand when the same reaction was conducted at room temperature, it took 840 min to give 49% *anti*-β-aminoalcohol with 87–93% *ee* (Table 1, entries 26 and 27), which is 420 times slower as compared to the reaction conducted under MW irradiation with comparable enantioselectivity. The rapid homogeneous energy transfer offered by MW irradiation as compared to traditional heating could be responsible for this remarkable reduction

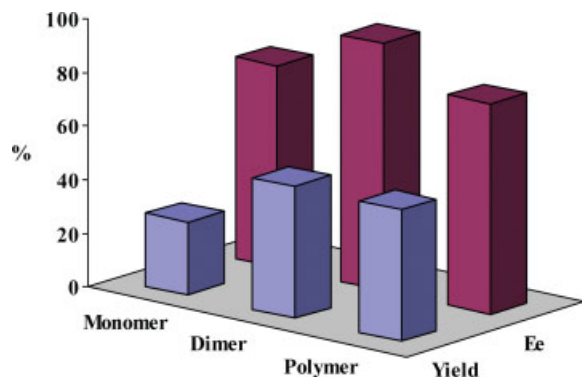
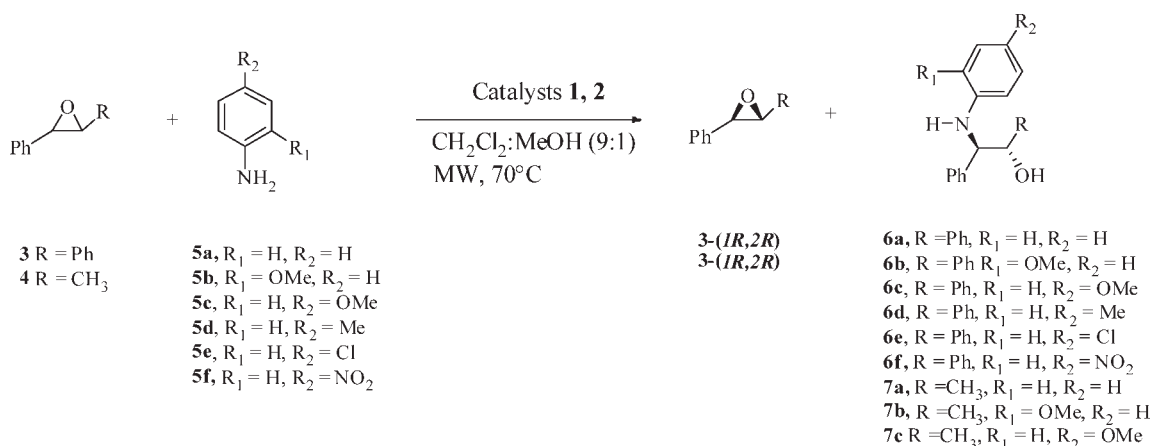


Fig. 1. 3D view representation of % yield and % ee of anti-β-amino alcohol of *trans*-stilbene oxide using monomer, dimer, and polymer Cr(III) salen complexes. [Color figure can be viewed in the online issue, which is available at www.interscience.wiley.com.]

of reaction time without significant reduction in enantioselectivity.

In a metal complex catalyzed ring opening of epoxides, electronic and steric features of nucleophiles play a very important role. In view of this, we conducted the AKR of *trans*-stilbene oxide with various substituted anilines under our optimized reaction conditions. Out of the *ortho*- and *para*-methoxy anilines used as nucleophile, the latter gave aminoalcohol with somewhat higher ee (Table 2, entry 6) with catalyst **2**, while for catalyst **1** the results are at par for both *ortho*- and *para*-methoxy anilines. On the other hand, *para*-methyl aniline produced aminoalcohol with excellent ee (94%) with catalyst **1** (Table 2, entry 7). *Para*-chloroaniline induced poor ee (Table 2, entries 9 and 10) possibly because of its poor nucleophilicity. In the case of anilines having strongly electron withdrawing group such as in the case of *para*-nitro aniline AKR, reaction with

TABLE 2. Aminolytic kinetic resolution (AKR) of *trans*-aromatic epoxides using recyclable catalysts with, anilines under microwave irradiations under optimized condition^a



Entry	Epoxide	Amine	<i>trans</i> -Epoxide		<i>anti</i> -β-Amino alcohols		<i>s</i> ^b
			ee ^c (%)	Yield ^d (%)	ee ^c (%)	Yield ^d (%)	
1 (2)	3	5a	90 (72)	48 (48)	92 (78)	49 (49)	31 (10)
3 (4)	3	5b	80 (65)	47 (46)	88 (67)	45 (39)	20 (5)
5 (6)	3	5c	82 (68)	48 (47)	89 (70)	47 (38)	22 (7)
7 (8)	3	5d	92 (69)	48 (48)	94 (78)	49 (47)	43 (10)
9 (10)	3	5e	65 (81)	86 (83)	19 (30)	25 (13)	2 (1)
11 (12)	3	5f	—	—	—	—	—
13 (14)	4	5a	86	40	59 (46)	46 (45)	5 (3)
15 (16)	4	5b	85	38	62 (47)	48 (46)	5 (2)
17 (18)	4	5c	88	36	69 (53)	49 (47)	7 (4)

^a0.01 mmol catalyst was taken in 200 μl CH₂Cl₂:MeOH (9:1), 0.2 mmol epoxides and 0.1 mmol aniline were added and the reaction mixture was irradiated for 2 min in a microwave oven at 70°C. Data presented in parentheses are for catalyst **2**.

^b*s* = selectivity factor = $\ln[1 - c(1 + ee)] / \ln[1 - c(1 - ee)]$, where ee is enantiomeric excess of the amino alcohol product and *c* is the conversion (set to the isolated yield).²³

^cThe diastereoselectivity of the product was found to be >99% determined by ¹H NMR and HPLC. The ee of aminoalcohols and epoxides are determined on Chiralpak OD column and the absolute configuration was assigned by comparing the optical rotation with literature.²³

^dThe conversion of the product **7a** was determined on ChiralPak OD column, using a calibration curve of epoxide and amino alcohols and rest given as isolated yield.

TABLE 3. Recycling data of catalyst 1/(2) for AKR of *trans*-stilbene oxide, using aniline as nucleophile^a

Run	Time (min)	Epoxide 3		<i>anti</i> -β-Amino alcohol 6a	
		ee (%)	Yield (%)	ee (%)	Yield (%)
1	2.0	90 (72)	48 (48)	92 (78)	49 (49)
2	2.0	89 (70)	47 (49)	91 (78)	48 (48)
3	2.0	89 (71)	46 (48)	92 (77)	48 (48)
4	2.0	90 (72)	46 (48)	91 (76)	47 (47)
5	2.0	90 (72)	48 (48)	92 (77)	46 (46)

^aThe 0.01 mmol catalyst was taken in 200 μl CH₂Cl₂:MeOH (9:1), 0.2 mmol epoxides and 0.1 mmol aniline were added and the reaction mixture was irradiated for 2 min in a microwave oven at 70°C. Results in parentheses is for catalyst 2.

trans-stilbene oxide does not commence (Table 2, entries 11 and 12). AKR of *trans*-β-methyl styrene oxide with aniline gave exclusively *anti*-β-aminoalcohol with moderate ee and yield (Table 2, entries 13–18). Based on the optical rotation studies for the products **7b–7c**, their absolute configuration was shown to be 1*S*,2*R*.²³

The catalyst recycle experiments were conducted for the AKR of *trans*-stilbene oxide, using aniline as a nucleophile with dimeric and polymeric Cr(III) salen complexes **1** and **2** as catalysts. In the postcatalysis work-up step, the catalysts were precipitated out by the addition of hexane: diethylether (1:1), which were filtered, washed with *n*-hexane, and dried in vacuum. The recovered catalysts were repeatedly used for four more catalysis runs which showed no apparent loss in the reactivity and enantioselectivity of the catalysts. However, there was some physical loss in the quantity of the catalysts during the catalyst recovery process (Table 3).

CONCLUSION

In summary, we have developed an efficient AKR protocol for the ring opening of *trans*-stilbene oxide, *trans*-β-methyl styrene oxide with various aromatic amines, to achieve respective *trans* β-amino alcohols in high enantio and diastereoselectivity and yields in a very short time under microwave irradiation. Unlike the monomeric Cr(III) salen complex, catalyst **1** gave higher enantioselectivity up to 94% with 4-methylaniline. Both dimeric and polymeric Cr(III) salen complexes were easily recyclable and were reused five times with retention of enantioselectivity.

ACKNOWLEDGMENT

RIK is thankful to Dr. P. K. Ghosh, the Director, of the Institute for providing instrumentation facilities.

LITERATURE CITED

- Jacobsen EN, Wu MH. Comprehensive asymmetric catalysis. In: Jacobsen EN, Pfaltz A, Yamamoto H, editors. Ring opening of epoxides and related reactions in comprehensive asymmetric catalysis, Vol. 3. New York: Springer; 1999. p 1309.
- Hodgson DM, Gibbs AR, Lee GP. Enantioselective desymmetrisation of achiral epoxides. *Tetrahedron* 1996;52:14361–14384.
- Bergmeier SC. The synthesis of vicinal amino alcohols. *Tetrahedron* 2000;56:2561–2576.
- Ager DJ, Prakash I, Schaad DR. 1,2-Amino alcohols and their heterocyclic derivatives as chiral auxiliaries in asymmetric synthesis. *Chem Rev* 1996;96:835–875.
- Li G, Chang HT, Sharpless KB. Catalytic asymmetric aminohydroxylation (AA) of olefins. *Angew Chem Int Ed Engl* 1996;35:451–454.
- Brien PO. Sharpless asymmetric aminohydroxylation: Scope, limitations, and use in synthesis. *Angew Chem Int Ed* 1999;38:326–329.
- List B. The direct catalytic asymmetric three-component Mannich reaction. *J Am Chem Soc* 2000;122:9336–9337.
- Corrêdo A, Notz W, Zhong G, Betancort JM, Barbas CF. A highly enantioselective amino acid-catalyzed route to functionalized α-amino acids. *J Am Chem Soc* 2002;124:1842–1843.
- Trost BM, Terrell LR. A direct catalytic asymmetric Mannich-type reaction to *syn*-amino alcohols. *J Am Chem Soc* 2003;125:338–339.
- Kobayashi S, Hishitani H, Ueno M. Catalytic asymmetric synthesis of both *syn*- and *anti*-β-amino alcohols. *J Am Chem Soc* 1998;120:431–432.
- Matsunaga S, Kumagai N, Harada S, Shibasaki M. *anti*-Selective direct catalytic asymmetric Mannich-type reaction of hydroxyketone providing β-amino alcohols. *J Am Chem Soc* 2003;125:4712–4713.
- Fu XL, Wu SH. A regio- and stereoselective synthesis of β-amino alcohols. *Synth Commun* 1997;27:1677–1683.
- Hou XL, Wu J, Dai LX, Xia LJ, Tang MH. Desymmetric ring-opening of *meso*-epoxides with anilines: a simple way to chiral β-amino alcohols. *Tetrahedron: Asymmetry* 1998;9:1747–1752.
- Sagawa S, Abe H, Hase Y, Inaba T. Catalytic asymmetric aminolysis of 3,5,8-trioxabicyclo[5.1.0]octane providing an optically pure 2-amino-1,3,4-butanetriol equivalent. *J Org Chem* 1999;64:4962–4965.
- Sekaine A, Ohshima T, Shibasaki M. An enantioselective formal synthesis of 4-demethoxydaunomycin using the catalytic asymmetric ring opening reaction of *meso*-epoxide with *p*-anisidine. *Tetrahedron* 2002;58:75–82.
- Carrée F, Gil R, Collin J. Samarium iodides catalyzed *meso*-epoxides ring opening by aromatic amines. *Tetrahedron Lett* 2004;45:7749–7751.
- Carrée F, Gil R, Collin J. Enantioselective ring opening of *meso*-epoxides by aromatic amines catalyzed by lanthanide iodo binaphtholates. *Org Lett* 2005;7:1023–1026.
- Schneider C, Sreekanth AR, Mai E. Scandium-bipyridine-catalyzed enantioselective addition of alcohols and amines to *meso*-epoxides. *Angew Chem Int Ed Engl* 2004;43:5691–5694.
- Kureshy RI, Singh S, Khan NH, Abdi SHR, Suresh E, Jasra RV. Facile enantioselective ring-opening reaction of *meso*-epoxides with anilines using (S)-(-)-BINOL-Ti complex as a catalyst. *Eur J Org Chem* 2006;1303–1309.
- Keith JM, Larrow JF, Jacobsen EN. Practical considerations in kinetic resolution reactions. *Adv Synth Catal* 2002;343:5–26.
- Label H, Jacobsen EN. Chromium catalyzed kinetic resolution of 2,2-disubstituted epoxides. *Tetrahedron Lett* 1999;40:7303–7306.
- Bandani M, Cozzi PG, Melchiorre P, Umani-Ronchi A. Kinetic resolution of epoxides by a C=C bond-forming reaction: Highly enantioselective addition of indoles to *cis*, *trans*, and *meso* aromatic epoxides catalyzed by [Cr(salen)] complexes. *Angew Chem Int Ed* 2004;43:84–87.
- Bartoli G, Bosco M, Carlone A, Locatelli M, Massaccesi M, Melchiorre P, Sambri L. Asymmetric aminolysis of aromatic epoxides: A facile catalytic enantioselective synthesis of *anti*-β-amino alcohols. *Org Lett* 2004;6:2173–2176.
- Song CE, Lee S. Supported chiral catalysts on inorganic materials. *Chem Rev* 2002;102:3495–3524.
- Xia QH, Ge HQ, Ye CP, Liu ZM, Su KX. Advances in homogeneous and heterogeneous catalytic asymmetric epoxidation. *Chem Rev* 2005;105:1603–1662.
- Song CE, Oh CR, Roh EJ, Choo DJ. Cr(salen) catalysed asymmetric ring opening reactions of epoxides in room temperature ionic liquids. *Chem Commun* 2000;1743–1744.
- Loupy A. Microwaves in organic synthesis. Weinheim: Wiley-VCH; 2002.

28. Kappe CO, Dallinger D. The impact of microwave synthesis on drug discovery. *Nat Rev Drug Discov* 2006;5:51–63.
29. Kappe CO. Controlled microwave heating in modern organic synthesis. *Angew Chem Int Ed* 2004;43:6250–6284.
30. Perreux L, Loupy A. A tentative rationalization of microwave effects in organic synthesis according to the reaction medium and mechanistic considerations. *Tetrahedron* 2001;57:9199–9223.
31. Adam D. Microwave chemistry: Out of the kitchen. *Nature* 2003;421:571–572.
32. Lindström UM, Olofsson B, Somfai P. Microwave-assisted aminolysis of vinyl epoxides. *Tetrahedron Lett* 1999;40:9273–9276.
33. Favretto L, Nugent WA, Licini G. Highly regioselective microwave-assisted synthesis of enantiopure C_3 -symmetric trialkanolamines. *Tetrahedron Lett* 2002;43:2581–2584.
34. Mojtahedi MM, Ghasemi MH, Abaee MS, Bolourtchian M. Solvent-free solid-supported recovery of carbonyl compounds from *N,N*-dimethylhydrazones under microwave irradiation. *ARKIVOC* 2005;15:68–73.
35. Glas H, Thiel WR. Microwave-assisted synthesis of chiral imidazolyl and pyrazolyl alcohols. *Tetrahedron Lett* 1998;39:5509–5510.
36. Gupta R, Paul S, Gupta AK, Kachroo PL. Opening of oxirane ring with *N*-nucleophiles under microwave irradiation. *Indian J Chem* 1997;36B:281–283.
37. Khosropous AR, Khodaei MM, Ghazati K. $Bi(OTf)_3$ - and $Bi(TFA)_3$ -catalyzed ring opening of epoxides with anilines under microwave irradiation. *Chem Lett* 2004;33:304–305.
38. Mojtahedi MM, Saidi MR, Bolourtchian M. Microwave-assisted aminolysis of epoxides under solvent-free conditions catalyzed by montmorillonite clay. *J Chem Res (S)* 1999;128–129.
39. Lutsenko S, Moberg C. Microwave-mediated ruthenium-catalyzed asymmetric hydrogen transfer. *Tetrahedron: Asymmetry* 2001;12:2529–2532.
40. Diao BML, Jacobs PA. Microwave-assisted Cr(salen)-catalysed asymmetric ring opening of epoxides. *J Catal* 2005;235:428–430.
41. Kureshy RI, Khan NH, Abdi SHR, Patel ST, Jasra RV. Dimeric chiral Mn(III) Schiff base complex-catalysed enantioselective epoxidation of non-functionalised alkenes. *Tetrahedron Lett* 2001;42:2915–2918.
42. Kureshy RI, Khan NH, Abdi SHR, Patel ST, Jasra RV. Enantioselective epoxidation of non-functionalised alkenes using a urea–hydrogen peroxide oxidant and a dimeric homochiral Mn(III)-Schiff base complex catalyst. *Tetrahedron: Asymmetry*, 2001;12:433–438.
43. Kureshy RI, Khan NH, Abdi SHR, Singh S, Ahmad I, Jasra RV. Catalytic asymmetric epoxidation of non-functionalised alkenes using polymeric Mn(III) salen as catalysts and NaOCl as oxidant. *J Mol Catal A: Chem* 2004;218:141–146.
44. Khan NH, Agrawal S, Kureshy RI, Abdi SHR, Mayani VJ, Jasra RV. Asymmetric synthesis of *O*-acetylcyanohydrins by reaction of aldehydes with NaCN/KCN catalyzed by recyclable chiral dimeric titanium(IV)/vanadium(V) salen complexes. *Eur J Org Chem* 2006;14:3175–3180.
45. Khan NH, Agrawal S, Kureshy RI, Abdi SHR, Mayani VJ, Jasra RV. Asymmetric addition of trimethylsilyl cyanide to aldehydes promoted by chiral polymeric vanadium(V) salen complex as an efficient and recyclable catalyst. *Tetrahedron: Asymmetry* 2006;17:2659–2666.
46. Kureshy RI, Singh S, Khan NH, Abdi SHR, Ahmad I, Bhatt A, Jasra RV. Improved catalytic activity of homochiral dimeric cobalt–salen complex in hydrolytic kinetic resolution of terminal racemic epoxides. *Chirality* 2005;17:1–5.
47. Kureshy RI, Ahmad I, Pathak K, Khan NH, Abdi SHR, Prathap KJ, Jasra RV. Easily recyclable chiral polymeric Mn(III) salen complexes for oxidative kinetic resolution of racemic secondary alcohols. *Chirality* 2007;19:352–357.
48. Pathak K, Ahmad I, Abdi SHR, Kureshy RI, Khan NH, Jasra RV. Oxidative kinetic resolution of racemic secondary alcohols catalyzed by recyclable chiral dimeric Mn(III) salen catalysts. *J Mol Catal A: Chem* 2007;274:120–126.
49. Kureshy RI, Singh S, Khan NH, Abdi SHR, Agrawal S, Mayani V, Jasra RV. Microwave-assisted asymmetric ring opening of *meso*-epoxides with aromatic amines catalyzed by a Ti-S(–)-BINOL complex. *Tetrahedron Lett* 2006;47:5277–5279.
50. Kureshy RI, Singh S, Khan NH, Abdi SHR, Agrawal S, Jasra RV. Enantioselective aminolytic kinetic resolution (AKR) of epoxides catalyzed by recyclable polymeric Cr(III) salen complexes. *Tetrahedron: Asymmetry* 2006;17:1638–1643.
51. G. Moyna, H. J. Williams, A. L. Scott. An improved procedure for the epoxidation of methyl cinnamate derivatives and production of acid-sensitive epoxides. *Synth Commun* 1996;26:2235–2240.
52. Daly AM, Renahan MF, Gilheny DG. High enantioselectivities in an (*E*)-alkene epoxidation by catalytically active chromium salen complexes. Insight into the catalytic cycle. *Org Lett* 2001;3:663–666.
53. Martinez LE, Leighton JL, Carsen DH, Jacobsen EN. Highly enantioselective ring opening of epoxides catalyzed by (salen)Cr(III) complexes *J Am Chem Soc* 1995;117:5897–5898.