Core-Shell Interface Oriented Synthesis of Bowl-Structured Hollow Silica Nanospheres Using Self-Assembled ABC Triblock Copolymeric Micelles

Md Nuruzzaman\textsuperscript{ab}, Yanju Liu\textsuperscript{ab}, Mohammad Mahmudur Rahman\textsuperscript{ab}, Ravi Naidu\textsuperscript{ab*}, Rajarathnam Dharmarajan\textsuperscript{ab}, Ho Kyong Shon\textsuperscript{bc}, and Yun Chul Woo\textsuperscript{d}

\textsuperscript{a}Global Centre for Environmental Remediation (GCER), Faculty of Science, The University of Newcastle, University Drive, Callaghan, NSW 2308, Australia

\textsuperscript{b}Cooperative Research Centre for Contamination Assessment and Remediation of the Environment (CRC CARE), ATC Building, The University of Newcastle, Callaghan, NSW 2308, Australia

\textsuperscript{c}Centre for Technology in Water and Wastewater, School of Civil and Environmental Engineering, University of Technology Sydney (UTS), P.O. Box 123, 15 Broadway, NSW 2007, Australia

\textsuperscript{d}Environment and Plant Research Institute, Korea Institute of Civil Engineering and Building Technology (KICT), 283, Goyangdae-Ro, Ilsanseo-Gu, Goyang-Si, Gyeonggi-Do 411-712, Republic of Korea
*Corresponding author: Ravi Naidu,

Email: Ravi.Naidu@newcastle.edu.au; Ravi.Naidu@crecare.com

Postal address: Room 147, ATC Building, The University of Newcastle, University Drive,
Callaghan, NSW 2308, Australia
Core-Shell Interface Oriented Synthesis of Bowl-Structured Hollow Silica Nanospheres Using Self-Assembled ABC Triblock Copolymeric Micelles

Md Nuruzzaman\textsuperscript{ab}, Yanju Liu\textsuperscript{ab}, Mohammad Mahmudur Rahman\textsuperscript{ab}, Ravi Naidu\textsuperscript{a,+}, Ho Kyong Shon\textsuperscript{bc}, Rajarathnam Dharmarajan\textsuperscript{ab}, and Yun Chul Woo\textsuperscript{c}

ABSTRACT

Hollow porous silica nanospheres (HSNs) are emerging classes of cutting-edge nanostructured materials. They have elicited much interest as carriers of active molecules delivery due to their amorphous chemical structure, non-toxic nature and biocompatibility. Structural development with hierarchical morphology is mostly required for obtaining the desired performance. In this context, large through-holes or pore openings on shells are desired so that the post-synthesis loading of active molecules HSNs via a simple immersion method can be facilitated. This study reports the synthesis of HSNs with large through-holes or pore openings on shells, which are subsequently termed bowl-structured hollow porous silica nanospheres (BHSNs). The synthesis of BHSNs was mediated by the core-shell interfaces of the core-shell-corona structured micelles obtained from a commercially available ABC triblock copolymer (polystyrene-b-poly(2-vinyl pyridine)-b-poly(ethylene oxide) (PS-P2VP- PEO)). In this synthesis process, polymer@SiO\textsubscript{2} composite structure was formed because of the deposition of silica (SiO\textsubscript{2}) on the micelles’ core. The P2VP block played the significant role in: firstly, hydrolysis and condensation of the silica precursor i.e. tetraethyorthosilicate (TEOS); and secondly, maintaining the shell’s growth. The
PS core of the micelles built the void spaces. Transmission electron microscopy (TEM) images revealed a spherical hollow structure with an average particle size of 41.87 ± 3.28 nm. The average diameter of void spaces was 21.71 ± 1.22 nm and shell thickness was 10.17 ± 1.68 nm. According to the TEM image analysis the average large pore was determined as 15.95 nm. Scanning electron microscopy (SEM) images further confirmed the presence of large single pores or openings in shells. These were formed due to the accumulated ethanol on PS core acting to prevent the growth of silica.

KEYWORDS. Silica nanospheres, Bowl-structure, Interfacial synthesis, ABC triblock copolymer, Hollow structure.

INTRODUCTION

Inorganic hollow-structured nanomaterials have emerged as potentially useful functional phenomena. Such nanomaterials have generated increasing research interest as carriers of active molecules, because they have potential in helping develop controlled delivery systems in various fields. This is due to their large specific surface area, low density, mechanical and thermal stabilities, surface permeability, high loading capacity, controllable structure, and a wide range of applications.1-3 Among the various hollow-structured inorganic nano-carriers, silica-based nanoparticles such as hollow porous silica nanospheres (HSNs) are considered most promising because of their amorphous chemical structure, non-toxic nature4 and biocompatibility5. As well as drug delivery their potential applications have been extended to agrochemical delivery, especially pesticides, particularly over the last decade.6-9 It is expected that such carrier materials
can be utilized for the preparation of controlled-release formulations (CRFs) due to the novel properties of nano-size particles or materials.\textsuperscript{10}

It is important to ensure the quality and uniform particle size of nanospheres during their synthesis so that the desired performance will be obtained.\textsuperscript{11} Recent studies suggest that the size of pores in the shells of hollow nanosphere material is important because they play a significant role in the encapsulation and delivery of functional substances.\textsuperscript{12} For example, HSNs’ performance declines during post-synthesis loading by simple immersion methods because of small mesopores (2-5 nm).\textsuperscript{8,9} Loading active molecules after synthesis mainly depends on diffusion processes and the efficiency of this loading varies according to the size of mesopores in the shell. Therefore, technology such as high-pressure supercritical fluid loading is required to facilitate post-synthesis loading of active molecules into the inner core or void space of HSNs.\textsuperscript{8,9} More importantly, to serve as an ideal carrier, the HSNs must be structured with an appropriate pore size that is larger than the kinetic diameter of the respective active molecules.\textsuperscript{13,14} It was reported that pore size less than 10 nm often obstructs the application of ordered mesoporous nanomaterials in many fields especially where it requires a fast transport and efficient loading of large-size active molecules such as genetic material, proteins, drugs, organic bioactive molecules etc.\textsuperscript{15}

In this aspect, hollow nanospheres or microspheres with large through-holes (HMLS), or pore openings wider than 10 nm, are highly desirable for facilitating the loading of active guest molecules.\textsuperscript{16} HMLS can even encapsulate large molecules, for example proteins or DNA\textsuperscript{17}, with diffusion this processes producing any limitations. The guest molecules could also be delivered in a controlled way from such nanocarriers where initially a fast release rate is observed which is desired in many applications such as drug delivery, pesticide delivery etc. to compensate rapidly against the target germs or pests. Later on, a sustained releasing profile is observed due to
electrostatic interaction and/or hydrogen bonding interaction between the guest molecules and Si-
OH groups of HMLs. However, to accomplish sustained release profile surface functionalization
seems to be essential where large through-holes facilitate the surface functionalization process to
modify the inner core surfaces on demand. In this case, with increasing amounts of surface
functionalization agents may reduce the loading capacity but slow releasing profile is observed. In
addition, the guest molecules also could be delivered to the target more efficiently preparing
environmentally responsive controlled release system through post loading surface
functionalization or coating with polymers.

To date, various synthetic methods for producing hollow microspheres with a large shell pore
size (>10 nm) have utilized approaches such as template-directed synthesis, shell-breaking,
Ostwald ripening and galvanic replacement reaction. These methods have produced various
types of HMLS with single, multiple and hierarchal shell holes, and depending on their
morphology, the spheres have been termed bowl-like, mouth-like, golf ball-like, or cage-
like. It is apparent that the physico-chemical properties of synthesized hollow-structured
nanomaterials vary depending on their synthesis technique. Several methods have been employed
for synthesizing HSNs, such as sacrificial hard templates, polymeric soft templates,
surfactant/vesicle templates and emulsion templating technique, producing various shell
thicknesses and porosity of structures. Among these methods, hard templating method suffers
from certain limitations. In the hard templating approach, removing the template is complicated,
time-consuming and unsuitable for mass production due to the utilization of costly hard
template. While the soft templating approach offers more opportunities for large scale
synthesis of mesoporous materials compared to hard templating approach. Of the various soft
templates based on cationic surfactants or Pluronic amphiphilic block copolymers were employed
initially for the synthesis of mesoporous nanomaterials.\textsuperscript{15} However, they can produce only a small pore size due to their short chain length.\textsuperscript{27} Thus, non-Pluronic based amphiphilic AB diblock or symmetric ABA triblock copolymers consisting of hydrophilic segments, such as PEO, poly(L-glutamic acid) (PLGA), poly(2-vinylpyridine) (P2VP) or poly(4-vinylpyridine) (P4VP), and hydrophobic segments such as polystyrene (PS), polyisoprene (PI), polybutadiene (PB) and polyacrylonitrile (PAN) created much attention for synthesizing various mesoporous materials.\textsuperscript{27,28} With such block copolymers the mesoporous structure, shape, pore size is also considered to be adjustable by controlling the synthesis conditions (such as temperature, pH, ionic strength etc.) and properties of the block copolymers (such as composition, block sequence, molecular weights, volume fractions etc.).\textsuperscript{14,29,30} Along with block copolymer, various amphiphilic polypeptides are also utilized for the synthesis of mesoporous materials through biomineralization process.\textsuperscript{31-34} Nonetheless, aggregation of nanoparticles is a common phenomenon observed in these processes.\textsuperscript{35} Generally, non-Pluronic and polypeptide based amphiphilic AB diblock or symmetric ABA triblock copolymers were found mostly suited for synthesizing 2-D or 3-D structured mesoporous materials. Synthesis of zero dimensional nanomaterial is still challenging utilizing such block copolymers. Therefore, HSNs with large through-holes with several morphologies that have been synthesized using these templates were not uniform, even when the particle size was observed larger than 100 nm.

Advances in interfacial chemistry have enabled HSNs synthesis by more effectively controlling the nucleation and growth of silica on templates. Recent studies have shown that ABC triblock copolymers are able to form micelles with a core-shell-corona structure due to their self-assembling properties. This capacity makes them excellent templates for synthesizing several uniform and monodispersed hollow porous nanomaterials including HSNs.\textsuperscript{11,35-38} For example,
micelles of poly(styrene-\textit{b}-2-vinyl pyridine-\textit{b}-ethylene oxide) (PS-P2VP-PEO) were successfully utilized for the synthesis of HSNs where tetramethyl orthosilicate was used as a silica precursor.\textsuperscript{11,35,36} Nonetheless, no large pore was observed in the shell of HSNs during this synthesis process.

It is well investigated that each chemical involved in the synthesis of HSNs can play a significant role in their morphology and behavior. It is expected that changing the precursor to tetraethylorthosilicate (TEOS) instead of tetramethylorthosilicate (TMOS) for the synthesis of HSNs using core-shell-corona structured micelles of PS-P2VP-PEO triblock copolymer utilized as soft template will facilitate the production of large through-holes or pore openings in the shell of HSNs. Generally, TMOS and TEOS are the most widely used precursors for the synthesis of silica using the sol-gel process.\textsuperscript{39} During the hydrolysis of these precursors, TEOS generates more hydrophobic by-products such as ethanol (log\textit{P} = -0.18) compared to methanol (log\textit{P} = -0.69), which is the by-product of hydrolyzed TMOS. In contrast the hydrolysis and condensation rate of TMOS is faster than that of TEOS.\textsuperscript{40} Thus, it is expected that during the hydrolysis process the ethanol will be able to interact with the hydrophobic PS block more efficiently. Accumulation of ethanol as droplets will be further promoted by hydrogen bonding interaction with the P2VP block. Consequently, deposition of silica will be prevented and as a result, large through-holes or pores in the shell of HSNs will be observed after calcination.

Considering the hypothetical mechanisms, this study was conducted to synthesize HSNs with large through holes or pore openings in shells termed as bowl-structured hollow porous silica nanospheres (BHSNs) using the core-shell interfacial chemistries of the micelles. This study presents a detailed synthesis process concerning BHSNs. Specifically, we determined the effects of changing the precursor to TEOS instead of TMOS on particle morphology while the triblock
copolymers, PS-P2VP-PEO with a core-shell-corona structure serve as a soft template.

This study also presents plausible mechanisms for the synthesis of BHSNs and investigates the characteristics of synthesized BHSNs to establish their potential for future applications.

**EXPERIMENTAL SECTION**

**Materials and reagents**

PS-P2VP-PEO triblock copolymer with block sizes PS(20k)–P2VP(15k)–PEO(27k) was purchased from Polymer Source Inc., Canada. The number in parentheses represents the average molar mass (Mn) of the block chains, e.g., 20k denotes 20,000 g mol⁻¹. The poly-dispersity index (PDI), or ratio of weight average molecular weight (Mw) and number average molecular weight (Mn) of the tri-block copolymer, was 1.11. The chemical structure of the block copolymer is presented in Supporting Information (Scheme S1). Tetraethylorthosilicate (TEOS, >98%), dimethylformamide (DMF, 99.8%) and hydrochloric acid (37%, analytical grade) were purchased from Sigma-Aldrich, Australia. All chemicals were used without further purification.

**Preparation of polymeric soft template**

Micelles of PS-P2VP-PEO triblock copolymer were prepared according to the method described earlier. For the typical preparation of micelles, 2 wt.% of the copolymer was dissolved in 5 ml DMF containing 10 wt.% water. The mixture was stirred with a magnetic stirrer until the copolymer was completely dissolved. Subsequent formation of micelles was achieved by eliminating DMF from the solution by dialysis against regularly replaced Milli-Q (MQ) water. Completion of dialysis was confirmed by measuring the surface tension of the MQ water. Finally, the micelles were transferred to a 250 ml standard flask and diluted with MQ water to make the
concentration to 1 g L\(^{-1}\). The micelle structure’s formation was confirmed by images obtained using a FEI Quanta 450 FEG field emission scanning electron microscope (FESEM) with an accelerating voltage of 10 kV.

**Synthesis of polymer@silica nanocomposite**

After preparing the micelles the functional groups were activated by adjusting the pH value of the micelle suspension to 4 with 0.1 M HCl. After adding the desired amount of silica precursor (TEOS) to maintain a molar ratio of TEOS to P2VP around 30:1, the micelle suspension was slowly stirred for 2 days at room temperature. To ensure slow hydrolysis, TEOS was added drop-wise at 0.1 ml min\(^{-1}\). The suspension of the template micelle and the precursor was stored for 4 more days at room temperature without stirring to allow the silica network to be formed by the sol-gel reaction. The resulting polymer@silica (PS-P2VP-PEO/silica) nanocomposite particles were separated from the solvent by centrifugation at 14,000 rpm (19,720x g) and then washed 3 times with MQ water. Finally, the polymer@silica composite was dried at 50 °C for 12 hours.

**Formation of porous hollow silica nanospheres**

Porous hollow silica nanospheres were obtained by removing the polymeric template from the polymer@silica nanocomposite by calcination in a muffle furnace. The temperature of the furnace was ramped to 550 °C at the rate of 1 °C min\(^{-1}\) and retained at 550 °C for 4 h in air. The temperature required for the decomposition of polymer was pre-determined by thermo-gravimetric analysis (TGA). TGA was carried out using a Q600 SDT thermo-gravimetric analyzer (TA Instrument, United States). Approximately 2.5 mg of the polymer@silica nanocomposite sample was heated from 50 to 600 °C in a 40 µl platinum crucible (PN: 960149.901, United States). The heating rate
was 10 °C min⁻¹ under an atmosphere of continuous high purity nitrogen gas flowing at 120 ml
min⁻¹. The decomposition temperature required for the block copolymer was also ensured by TGA
analysis as described above.

Characterization

**Micro-morphology analysis.** The external surface morphology and elemental composition of
synthesized BHSNs and polymer@silica nanocomposite were evaluated using a field emission
scanning electron microscope (FESEM, Zeiss Supra 55VP, Carl Zeiss AG) fitted with an energy
dispersive X-ray spectroscopy (EDAX) facility. For SEM analysis, all the dried powder samples
were placed on a sample holder followed by coating with Au-Pd (8.5 nm) using a Leica EM
ACE600 high vacuum coater for high-resolution imaging. The SEM images were generated with
an accelerating voltage of 4.5 kV, and multiple image magnifications of various areas of the sample
were obtained. The hollow structure of the BHSNs was investigated by using a transmission
electron microscope (TEM, Tecnai T20, FEI Tecnai (LaB6)). Before TEM analysis the BHSNs
powder sample was dispersed in ethanol followed by drying one drop of BHSNs-containing
suspension placed on a copper grid (Ted Pella Inc., CA, USA). The TEM images were carried out
at an accelerating voltage of 120 kV. Whereas the high resolution TEM images were taken by
using a JEM-2100F (JEOL, USA) transmission electron microscope at an accelerating voltage of
200 kV. The shell thickness and void space of HSNs were evaluated by image analysis using
ImageJ software (NIH, free download).

**Fourier transform infrared spectroscopy.** The functional groups present in the polymer@silica
nanocomposite before and after calcination, as well as in the polymeric powder samples, were
assessed by attenuated total reflectance Fourier transform infrared (FTIR) spectrometer (ATR-
FTIR analysis was performed in the 4000 to 400 cm⁻¹ range with a signal resolution of 4 cm⁻¹. Each sample was scanned 32 times to obtain a good signal to noise ratio.

**X-ray diffraction pattern analysis.** The X-ray diffraction (XRD) pattern of BHSNs was evaluated by a Bruker D8 Discovery X-ray diffractometer equipped with Cu-Kα radiation (λ = 0.154 nm) generated at 40 kV and 40 mA. The scanning rate was 1° (2θ) min⁻¹ within the scanning range from 1° to 10° and 10° to 80° (2θ) with a step size of 0.04°.

**Hydrodynamic diameter and zeta potential measurement.** A Zetasizer Nano ZS90 (Malvern Instruments Ltd., UK) operating with a He-Ne laser at a wavelength of 633 nm was used in this study to determine both the zeta potential and hydrodynamic diameter of sample materials. The measurement of zeta potential was achieved by a combination of Laser Doppler Velocimetry and phase analysis light scattering (PALS) with Malvern’s patented M3-PALS technique. A blank consisting of ultrapure water (Milli-Q), Millipore, USA) was run systematically before the samples to calibrate the instrument. Zeta potential was measured within the pH range of 4-11. The particles’ nanosized diameter was determined by dynamic light scattering (DLS) mode.

**RESULTS AND DISCUSSION**

**Characterization of polymeric micelle.** For synthesis of BHSNs, the micelles of PS–P2VP–PEO were prepared initially as described in the experimental section. Figure 1 shows that spherical micelles of PS–P2VP–PEO were successfully formed with a uniform size range. The average diameter of the micelles was 40 ± 3.78 nm (at pH > 5) measured from SEM images using ImageJ software (NIH). In contrast, the average diameter of micelles had a measurement of 64 ± 2.3 nm using the DLS system at low pH (~4), indicating a monodispersed size distribution (Figure 2). The
difference in average diameter was observed in these two methods due to pH responsive behavior of the functional P2VP block. This block mainly consists of aromatic amines such as pyridine. The N atom of pyridine is readily protonated at lower pH (< 5) due to the presence of lone pair electrons (see Supporting Information, Scheme S1). The electrostatic repulsive forces generated among the PVP blocks result in considerable expansion of micelles’ diameter.\textsuperscript{41} In contrast, the deprotonated P2VP block is reported as being hydrophobic; hence, it collapsed on the PS block at high pH (> 5). In an earlier study, a similar trend was observed as the average diameter of the micelles was found to be 67 nm in the DLS system when produced with a triblock copolymer PS(20.1k)–P2VP(14.2k)–PEO(26.0k), while the average diameter was 42 nm as determined by TEM image analysis.\textsuperscript{11} The sizes as well as expanded diameters of micelles also varied depending on the block sizes of the copolymer.\textsuperscript{36,42} The effect of pH on micelle size distribution is well described elsewhere.\textsuperscript{41}

Figure 1. The morphological structure of micelles synthesized using the PS–PVP–PEO block copolymer as PS(20.0k)–P2VP(15.0k)–PEO(27.0k).
Figure 2. Histogram obtained from the DLS system showing the micelles’ diameter and corresponding intensity percentage of the PS–PVP–PEO block co-polymeric micelles at pH 4.

**Confirmation of polymer@silica composite structure.** To confirm polymer@silica composite structures, and to understand the interactions of silica deposition with the block copolymer, changes in the bonding structures of pristine PS-P2VP-PEO block copolymer as well as polymer@silica composite before and after calcination were evaluated by FTIR analysis (Figure 3). The structure of the block copolymer (PS-P2VP-PEO) is mainly dominated by alkyl and alkoxy groups. For this reason the block copolymer exhibited very strong bands at 2850 cm$^{-1}$ and 1097 cm$^{-1}$ corresponding to C–H stretching vibrations of alkyl (methyl and/or methylene) groups and –C–O–C– stretching vibrations of ethoxy groups of polymer backbone, respectively (Figure 3a).$^{42-44}$ The energy bands observed at 1582 cm$^{-1}$ and 1456 cm$^{-1}$ were attributed to >C=C< stretching energies present in the aromatic ring of corresponding phenyl and pyridyl groups that were present in the PS block and P2VP block, respectively, in the polymer backbone.$^{11,42}$ The band vibrations corresponding with wavenumbers at 1344 cm$^{-1}$ and 1265 cm$^{-1}$ were due to C–H bending of alkyl
groups of the polymeric chain, whereas band vibrations observed at wavelength number 692 cm\(^{-1}\) and 758 cm\(^{-1}\) were due to C-H bending of aromatic rings.

The FTIR spectrum of the polymer@silica composite has represented complex band vibrations of block copolymer and silica. (Figure 3b). Significant changes were observed in band vibrations of block copolymer present in polymer@silica composite compared to pristine PS-P2VP-PEO block co-polymer. These changes were caused by deposition of silica over the PS core as well as chemical bonding between silica and the P2PV block. It has previously been shown that polystyrene exhibits characteristic bands around 3001 cm\(^{-1}\) for C–H stretching of aromatic rings, 2850 cm\(^{-1}\) corresponds to C–H symmetric stretching of alkyl polymeric chains, and band vibration at 1452 cm\(^{-1}\) indicates >C=C< stretching of aromatic rings. In addition, the characteristic peaks at 1200-1400 cm\(^{-1}\) and 650-780 cm\(^{-1}\) were denoted as C-H bending in the aliphatic chain and in the aromatic ring, respectively.\(^45,46\) Conversely, shifting of the characteristic band vibrational peak of >C=C< stretching was related to aromatic phenyl rings of PS to a higher wavenumber (1500-1600 cm\(^{-1}\)).\(^47\) This peak shifting was mostly observed when PS was grafted with P2VP to form a block copolymer.

The PS block of the triblock copolymer used in this study is hydrophobic in nature, thus, core of the micelles are formed on dialysis via attractive forces among PS blocks (hydrophobic and π-π interaction) which is required to minimize energetically unfavorable solvophobic interactions.\(^48\) In this condition, band vibrational signal of phenyl group present in PS blocks becomes weak. Whereas polymer@silica composite was formed due to deposition of silica over the core, i.e. PS block, the silica layer may prevent the reception weak band vibrational signal of the PS block. Consequently, all the characteristic peaks of the PS block disappeared in the FTIR spectrum of the polymer@silica composite. Similarly, the band vibrational signal of oleic acid disappeared due to
silica deposition forming nanocrystal structure.\textsuperscript{49} It is therefore apparent that the band vibration observed at 1525 cm\textsuperscript{-1} is likely to be due to >C=C< stretching of the P2VP block’s pyridine rings. The peak appearing at wavenumber 1647 cm\textsuperscript{-1} is representative of the N-H group and indicates the hydrogen bonding interaction between silica and pyridine rings. However, the broad band with peak maxima at 1525 cm\textsuperscript{-1} and 1647 cm\textsuperscript{-1} occurred may due to overlapping of >C=C< stretching of pyridine rings and C–H bending of alkyl groups as well as OH/-NH vibrations\textsuperscript{49} respectively (Figure 3b inset). The formation of an N-H bond was also confirmed from the band vibrational peak observed at 2324 cm\textsuperscript{-1} (Figure 3b inset). In their study, Huang et al.\textsuperscript{50} observed a similar peak for N-H band vibration of structural amine present in octadecylamine. Furthermore, the symmetric C-H stretching band position at 2850 cm\textsuperscript{-1} shifted to a higher wavenumber (2910 cm\textsuperscript{-1}). The shifting of band position is also indicative of asymmetric stretching of C-H in the polymeric chain of P2VP.

Moreover, the shifting of band position provides information on electron transfer to pyridine rings, which ensures their electron resonance when a lone pair electron of N atoms was involved in hydrogen bonding interaction with silanol (Si-OH) or other moieties, such as ethanol (hydrolyzed by-products of TEOS). The newly developed band vibrations at 3832 cm\textsuperscript{-1} and 3740 cm\textsuperscript{-1} indicated O-H stretching of Si-OH. However, the band position of the most intensified block copolymeric peak shifted from 1097 cm\textsuperscript{-1} to 1078 cm\textsuperscript{-1} in the FTIR spectrum of the polymer@silica composite. This can be attributed to band vibration for both Si-O-Si and –C–O–C– stretching around this wavenumber, which was confirmed from the FTIR spectrum of the calcined polymer@silica composite sample. In the calcined polymer@silica composite sample, the peak intensity was drastically reduced with further shifting in peak position from 1078 cm\textsuperscript{-1} to 1074 cm\textsuperscript{-1}. This indicated Si-O-Si stretching whereas the reduced peak intensity also highlighted –C–O–C–
stretches at 1074 cm$^{-1}$ in the polymer@silica composite. The intensified band vibration at 1074 cm$^{-1}$ was attributed to the ethoxy group’s strong spectrum signal. It can therefore be assumed that the PEO block was free from silica deposition.

Moreover, the FTIR spectrum of the calcined polymer@silica composite sample revealed that along with –C–O–C– stretching band vibration all of the polymeric bands completely disappeared in the calcined polymer@silica composite structure. This left the characteristic vibration bands of pure SiO$_2$ (Figure 3c), indicating the complete removal of polymer from the polymer@silica composite. In the calcined sample a strong vibration band at 1074 cm$^{-1}$ corresponded to Si–O–Si symmetrical stretching ($\gamma_{\text{sym}}$) whereas the shoulder growth at 1184 cm$^{-1}$ indicated Si–O–Si asymmetrical stretching ($\gamma_{\text{asym}}$).$^{51}$ Other characteristic band vibrations were observed at 782 cm$^{-1}$ and at 954 cm$^{-1}$ possibly due to symmetric Si-O stretching vibration of Si-O-Si and Si-O stretching vibration bond of Si-OH, respectively.$^{52}$ The band vibration at 3832 cm$^{-1}$ and 3740 cm$^{-1}$ of O-H stretching also disappeared in the calcined sample, suggesting that the P2VP block interacted with the Si-OH group of the deposited silica.
Figure 3. FTIR-ATR spectra of a. PS-PVP-PEO block copolymer, b. Polymer@silica nanocomposite and c. Polymer@silica nanocomposite after calcination.

**Thermal analysis.** TGA analysis of the triblock copolymer as well as polymer@silica composite before and after calcination was carried out to investigate their thermal behavior and to quantify the amount (wt.%) of block copolymer template present in the composite (Figure 4). Figure 4a illustrates that 100% weight loss was observed for the block copolymeric sample, whereas the corresponding DTG curve demonstrates that the decomposition of the block polymer took place over the temperature range of 300 to 500 °C with the maximum mass loss rate occurring at 408 °C (Figure 4i). In contrast, 43% weight loss was calculated from the TGA curve due to decomposition of the block copolymer for the polymer@silica composite sample (Figure 4b). The DTG curve of the polymer@silica sample indicated the decomposition of polymeric materials from the
nanocomposite structure also occurred within the 300-500 °C temperature range. However, the temperature of the maximum mass loss rate shifted to 432 °C (Figure 4ii). In the polymer@silica sample, the peak maximum in the DTG shifted to a higher temperature compared to the polymer, probably due to dehydroxylation through which the P2VP block interacted with silica networks. This outcome also indicated that during core-shell arrangement in polymer@silica composite, silica was deposited on the polymeric core establishing a hydrogen bonding interaction with the P2VP block, which generated structural Si-OH. The FTIR spectrum of the polymer@silica composite also exhibited the specific band vibrations of –OH group (Figure 3b). Conversely, only 0.2% weight loss was found for the calcined polymer@silica composite sample and no corresponding specific peak was observed in the corresponding DTG curve (Figure 4iii). From this observation, it is reasonable to assume that total weight loss (around 43%) observed in polymer@silica composite was due to the block copolymer.
Elemental analysis. The elemental analysis of the polymer@silica composite before and after calcination strongly supports the results obtained by TGA analysis considering the three major elements, i.e. C, O and Si (Figure 5). It was found that the polymer@silica composite contained around 50% C which was drastically reduced to less than 1% by calcination (Table 1). Hence, calcination of the polymeric template completely removed it from the composite structure. Whereas the atomic (%) ratio of O to Si is much greater than 2:1 in the polymer@silica composite, it was almost 2:1 after calcination.
Figure 5. SEM images and EDAX spectrum of polymer@silica composite before calcination (a) and after calcination (b)

Generally, the atomic ratio of O and Si in SiO₂ is 2:1; hence, one can predict from this elemental analysis that due to calcination of the polymer@silica composite the polymeric template was

21
completely removed it from the composite structure. After calcination SiO$_2$ is the only material that is present in the composite structure. The ratio of O to Si in the polymer@silica was much greater than 2:1 due to the presence of O containing PEO block in the composite structure. Furthermore, from SEM images, it was observed that before calcination, the polymer@silica composite remained in a highly aggregated state that changed to a dispersed or loosely aggregated following calcination of the composite (Figure 5). This phenomenon supports the assumption of FTIR analysis that during polymer@silica formation the PEO block was free from silica deposition and mainly acted as a barrier between the silica nanospheres. Therefore, homogeneously distributed and uniform silica nanospheres were formed. The uniform size-distribution of silica nanospheres was further confirmed by TEM analysis as well as an assessment of particle size distribution of synthesized silica nanospheres using the DLS system.

Table 1: Elemental composition of polymer@silica nanocomposite before and after calcination determined by EDAX analysis

<table>
<thead>
<tr>
<th>Material status</th>
<th>C Weight %</th>
<th>C Atomic %</th>
<th>O Weight %</th>
<th>O Atomic %</th>
<th>Si Weight %</th>
<th>Si Atomic %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Polymer@silica</td>
<td>51</td>
<td>62</td>
<td>32</td>
<td>29</td>
<td>17</td>
<td>9</td>
</tr>
<tr>
<td>(Before calcination)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Polymer@silica</td>
<td>1</td>
<td>1</td>
<td>55</td>
<td>68</td>
<td>44</td>
<td>31</td>
</tr>
<tr>
<td>(After calcination)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Micro-morphology of synthesized silica nanospheres. Complete decomposition of the polymeric template by calcination was further confirmed by TEM (Figure 6a). Figure 6a shows that all the hollow-structured silica nanospheres were spherical with a uniform size, contained
large void spaces and had homogeneous size distribution. Based on the image analysis the average particle size was estimated as $41.87 \pm 3.28$ nm, the average void space was $21.71 \pm 1.22$ nm, and the average shell thickness was $10.17 \pm 1.68$ nm. Earlier studies indicated that the size of the void spaces was correlated with the chain length of the PS block, while an increasing trend in size of void spaces was observed with increasing molecular weight of the PS block.\textsuperscript{36,44} Gohy et al.\textsuperscript{41} measured the size of the PS core of the PS(20.1k)–P2VP(14.2k)–PEO(26k) micelle as 20 nm. We also observed the micelles’ diameter as 20 nm (83.6\%) when the image was analyzed using FESEM (see Supporting Information, Figure S1). Therefore it is reasonable to conclude that the void space of the synthesized HSNs is mainly structured with the PS block. In addition, shrinkage of the core also occurred during silica condensation, and consequently an expanded core diameter may be often observed by micro-morphology and DLS analysis. Shrinkage of hollow silica particles during calcination is also a common phenomenon while they are synthesized using a polymeric soft template. Rough outer surfaces were also observed indicating the presence of mesopores in the shell wall. Along with mesopores, the presence of large single pore on shell was confirmed from Figure 6b. Furthermore, TEM image analysis indicated the average diameter of the large pore on shell as 15.95 nm (see Supporting Information Figure S2). The SEM image of BHSNs also confirmed that the synthesized silica nanospheres were spherical in shape and homogeneously distributed (Figure 6b6c). Furthermore, the SEM images also depicted single large pore openings in the silica nanospheres, indicating the successful synthesis of bowl-structured hollow silica nanospheres (BHSNs) (Figure 6b6c). However, only a small fraction of the BHSNs possess such characteristic under electron microscopy due to morphological orientation of the particles.\textsuperscript{53} The single large opening or hole was observed only in SEM image when the nanospheres deposited with keeping the large single pore or hole towards electron beam. In other
positions, it was not possible to detect via SEM. Similar limitation was also observed for TEM analysis and only the pore openings could be observed when the pore openings are faced horizontally. From the available literature, similar phenomenon was observed for different nanoparticles containing large through holes on shell.54-56 The mesoporous shell can also be proved by the nitrogen adsorption-desorption isotherm (see Supporting Information Figure S3a), which exhibits typical type-IV hysteresis loop. Additionally, the Barrett–Joyner–Halenda (BJH) pore size distribution plot (see Supporting Information Figure S3b) indicates the presence of various size mesopores.
Figure 6. TEM images showing uniform size distribution of silica nanospheres with large void spaces (a), HRTEM images showing large single pore on shell (b) and SEM images showing large pore openings (c). Red arrows indicate pore openings in the shell.
X-ray diffraction pattern of BHSNs. The powder XRD patterns of the synthesized BHSNs are presented in Figure 6. The appearance of diffraction peaks 2θ between 1 to 2 degrees indicates the presence of mesopores on the surface (Figure 7a). A broad diffraction peak at 2θ angle between 2 to 3.2, conversely, suggests the possibility of a large porous structure consisting of hollow spherical particles, further confirming SEM observations of large pore openings in the shell. The d-spacing value indicated the average pore size was around 9 nm. However, the XRD pattern at higher 2θ angle highlighted a single broad peak centered at approximately 23°, which is characteristic of the diffraction of amorphous silica (Figure 7b). This peak is possibly indicative of a mesostructure of synthesized silica. No other peak was observed throughout the 2θ angle (from 1 to 80 degrees) indicating that the synthesized material is pure silica free from impurities.
Figure 7. XRD patterns of synthesized BHSNs obtained by using the PS-P2VP-PEO block copolymer a. lower angle and b. at higher angle

**Zeta potential and surface chemistry.** To investigate the surface chemistry of BHSNs, zeta potential values were measured over a wide range of pH (4-11). During this process, the initially suspended pH of BHSNs was adjusted at different pH levels followed by equilibration at room temperature for 12 h. After equilibration the final pH was measured and the corresponding zeta values were determined. The zeta values and the final pH are presented in Figure 8a. From Figure 8a, it is apparent that the negative zeta value of BHSNs increased when the pH of the aqueous suspension increased. From XRD analysis it is clear that the structures of the synthesized BHSNs are predominantly amorphous and apparently similar to pure silica. Silica is inherently an acidic oxide; the pH of an aqueous suspension of silica is slightly acidic. When exposed to water, partial or total particle surface hydroxylation can cause the formation of silanol groups such as Si(OH)$_n$.

Such silanol groups may dissociate in pure water through the following reactions:

$$\equiv \text{Si-OH} + \text{OH}^- \rightarrow \text{Si-O}^- + \text{H}_2\text{O} \quad \text{(pH > 9.0)}$$  (eq. 1)
Following eq. 1, the Si-OH normally tends to act as a proton donor in aqueous solution and becomes negatively charged. With increasing pH the silica surfaces become more negatively charged. Under low pH conditions Si-OH tends to become protonated and the degree of protonation increases with decreasing pH of the aqueous suspension (eq. 2). Si-OH only remains in a fully protonated state at an extremely low pH value (< 2). Therefore, over a wide range of pH, SiO₂ exhibits a negative surface charge in aqueous medium. Changes in initial pH of BHSNs after 12 h also indicated their similar surface properties. The differences between initial and final pH are presented in Figure 8b. It was observed that the final pH values increased when the initial system pH was below pH ~4.5. However, the opposite results were observed in suspensions with an initial pH above 4.5. Therefore, it can be assumed that the synthesized BHSNs acted as a proton acceptor at lower pH whereas they were deprotonated at higher pH. According to Figure 8b, it can also be concluded that the degree of deprotonation of BHSNs is much larger than protonation. Deprotonation of Si-OH groups at water-silica interfaces is important for binding ions and molecules to their surfaces.⁵⁷ The surface charge behavior of BHSNs demonstrates their ability to establish significant cumbic or electrostatic interactions with positively charged molecules at higher pH.
Figure 8. Showing (a) zeta potential and (b) protonated-deprotonated behavior of BHSNs in aqueous media at different pH conditions.

**Particle size of BHSNs.** In addition to surface properties, the ability of nanomaterials to disperse is also important to achieve the desired performance. Hence, the diameter of BHSNs was further investigated using the DLS system. Considering the higher zeta values of BHSNs at higher pH, the suspension pH was adjusted to 11.2. In the DLS system monodispersed particle size
distribution was observed where the z-average particle size was 44.61 nm (Figure 9). This average particle size proved to be similar to the average particle size observed through TEM image analysis (Figure 6a). This finding indicates that the synthesized BHSNs are able to completely disperse in aqueous solutions at higher pH because of their high negative surface charge where the electrostatic repulsion force is more prominent among the nanospheres.

Figure 9. Histogram obtained from the DLS system showing the particle size distribution of BHSNs and corresponding number percentage.

Mechanism of synthesis

Under acidic (pH <5) or basic (pH>8) conditions the synthesis of silica by the sol-gel process from organo-silicates such as TEOS proceeds through hydrolysis and condensation steps. These can be described by equations 3 and 4.\textsuperscript{58,59}

Hydrolysis of TEOS:

\[
\text{Si(OC}_2\text{H}_5)_4 + 4\text{H}_2\text{O} \rightarrow \text{Si(OH)}_4 + 4\text{C}_2\text{H}_5\text{OH} \quad \text{(eq. 3)}
\]

Condensation step:
$\text{Si(OH)}_4 \rightarrow \text{SiO}_2 + 2\text{H}_2\text{O}$  
(eq. 4)

However, the processes are deemed to occur through several steps on a suitable template such as: (1) fast hydrolysis catalyzed by highly acidic (e.g., in the presence of HCl) or basic (e.g., in the presence of NH$_3$) aqueous solution; (2) condensation of Si(OH)$_4$ to form silica-coated template spheres; (3) condensation of Si(OH)$_4$ to form free silica nanoclusters; (4) capture of free silica nanoclusters onto the template; and (5) aggregation of free silica nanoclusters into unwanted and irregular aggregates of silica.$^{60}$ It is essential to prevent the irregular aggregation that is driven by fast hydrolysis of TEOS in the presence of diluted HCl in aqueous solution.$^{60}$ It is expected that slow hydrolysis along with condensation of silica over a template could avoid such aggregation, while ensuring the absence of a free catalyst in the solution.$^{61}$

So far, core-shell-corona structured micelles of ABC triblock co-polymer (PS-P2VP-PEO) have demonstrated their potential as a template for synthesis of HSNs where core-shell interfaces of the micelles played the vital role in silica growth and deposition over micelles’ core. Thus, the void spaces of synthesized BHSNs were primarily depended on the core size of the micelles. Due to the hydrophobic nature of the PS block the rigid core structure of the micelles was formed during the dialysis process; it remained unchanged throughout the sol-gel reactions required for core-shell arrangement as polymer@silica composite formation. Therefore the PS blocks were employed to construct hollow interiors of BHSNs. Earlier studies showed that the size of the void spaces was correlated with the chain length of the PS block, while an increase in the size of void spaces was observed with increasing molecular weight of the PS block, i.e. chain length of the block.$^{36,44}$ In addition, shrinkage of the core occurred during condensation process. Shrinkage of hollow silica particles during calcination is also a common phenomenon while they are being synthesized using a polymeric soft template.
The pH responsive functional P2VP block of the copolymer played a significant role in the hydrolysis and condensation of TEOS in the sol-gel process. The P2VP block was ionized at low pH (< 5) and not only captured free silica nanoclusters onto the template but also acted as an acid catalyst during hydrolysis of TEOS. This was followed by controlled condensation of Si(OH)₄ to form silica-coated template spheres. The role of the P2VP block in hydrolysis and polycondensation of TEOS is described in Supporting Information (Schemes S2 and S3). Inner smooth surface and rough outer surfaces of BHSNs (Figure 1a, Supporting Information Figure S5S6) indicated the condensation of Si(OH)₄ on the PS block, condensation of Si(OH)₄ to form template-free nanoclusters as well as the capture of free silica nanoclusters onto the template. Herein, no other catalyst was required which ensured that no free catalyst was present in the synthesis media. Subsequently it prevented the irregular aggregation of silica in the solution. Therefore the core-shell arrangement of polymer@silica was successfully formed. This result was also supported by the TEM images of BHSNs since no rigid-structured silica nanoparticles were observed (Figure 6a, 6b). As the growth and deposition of silica on the template was facilitated by the P2VP block, subsequently the TEOS: P2VP molar ratio proved critical in controlling the shell thickness of silica nanospheres. PEO is also able to interact with silanol groups (Si–OH) of silica prominently through hydrogen bonding but only in extremely acidic conditions (pH 1). Hence, we believe that the PEO block was free from silica deposition and growth of silica networks was exclusively formed interacting with P2VP block of the micelles.

In this study, along with hollow spherical nanostructured silica spheres, a single large opening or hole was observed in the shells (Figure 6b6c). The silica nanospheres have been designated bowl-structured hollow silica nanospheres (BHSNs) due to their large openings (~15 nm) in relation to their small particle size (~45 nm). Synthesis of BHSNs using ABC tri-block
copolymeric micelles as a soft template has not yet been reported but hollow-structured mesoporous silica nanospheres have already been successfully synthesized.\cite{11,35,37,42} So far, depending on large pore openings in the shells, silica nanospheres synthesized in earlier studies had different structure. For example, in earlier studies, such material was denoted as ‘silica nanobottle’ and this may be due to the relatively large particle size compared to the pore openings.\cite{63,64} However, what caused the formation of single large pores on the shells is still unclear. In an earlier study, Zhang et al.\cite{63} assumed that the hole or large pore in the spheres was formed during calcination. This was attributed to the high pressure resulting from polymeric decomposition with this playing an active role in the formation of such pore structures in silica nanospheres.\cite{63}

It is important to note that the decomposition of the polymeric template could play a key role when the shell is solid and unable to form mesopores; in most cases, the shells are broken down.\cite{4} It is also reasonable to conclude that if high pressure plays a key role in large pore formation then such a structure would be developed in each case while a hollow structure is obtained from the sacrificial polymeric templating approach through calcination. However, this has not been commonly observed.\cite{51,65} It is therefore possible that not only high-pressure generation during calcination but also other mechanisms are involved in single large pore formation. The main difference between this study and those reported in the literature for synthesizing HSNs is the use of an organosilicate precursor, i.e. TEOS instead of TMOS where similar types of ABC triblock copolymer were used. From morphological differences between synthesized HSNs using these precursors (see Supporting Information Figure S43-S65), it can be assumed that the chemical behavior of the precursor or their by-product played a significant role in the formation of large pore openings, which is consistent with the recent investigations conducted by Yi et al.\cite{64} In their
study, a silica nanobottle with an extended neck and having a ‘flask bottle’ shape was
synthesized.\textsuperscript{64} Interestingly, the particle shape and size completely being changed with the
replacement of precursor materials has been reported elsewhere.\textsuperscript{64}

Considering the observations of Yi et al.\textsuperscript{64}, we therefore postulate that a by-product of TEOS
hydrolysis, such as ethanol, played a pivotal role in the formation of bowl-structured hollow silica
nanospheres. The chemical structure of ethanol includes both hydrophilic (-OH group) and
hydrophobic (hydrocarbon chain) moieties. In this study, ethanol generated from hydrolysis of
TEOS possesses a high affinity for the hydrophobic surface of the micelles’ PS core better than
water, yet ethanol is more hydrophobic than the PEO block, i.e. corona of the micelles.\textsuperscript{66} Therefore,
one ethanol is produced as a hydrolyzed by-product of TEOS, it remains at the PS-water interfaces
of micelles in an aqueous suspension. Further deposition of silica over the PS block enhanced shell
growth and ethanol tended to exude from PS surfaces, but aggregated together on micelles’ core
due to hydrogen bonding affinity of ethanol to ethanol. During the final stage of shell formation
the accumulated ethanol extended throughout the P2VP block to form droplets. The highly
accumulated ethanol may also interact with P2VP by hydrogen-bonding and the accumulated
ethanol droplets also facilitate gel formation of the PEO block of the micelles\textsuperscript{67} which may be
responsible for particles’ aggregation, as observed in Figure 5a.

In the FTIR spectrum, an intensified peak growth was observed in the polymer@silica
nanocomposite compared to the block copolymer at 1078 cm\textsuperscript{-1},\textsuperscript{68} whereas the presence of -OH
band vibrations indicated the possible existence of ethanol (Figure 2a). Consequently, accumulated
ethanol molecules prevented the deposition of silica over the PS block and bowl-structured hollow
porous silica nanospheres were observed after calcination of the polymer@silica composite. For
this reason the ethanol/water ratio is an important factor for the deposition of silica over PS.\textsuperscript{69} The postulated mechanisms for the synthesis of such a structure are illustrated in Scheme 1.

Scheme 1: Postulated mechanisms of bowl-structured hollow silica nanosphere synthesis

CONCLUSIONS

In this study, HSNs with large through-hole or pore openings in the shell were successfully synthesized using core-shell-corona structured micelles of a self-assembled ABC tri-block copolymer (PS-P2VP-PEO) as a soft template. Due to the large pore openings in the shells, the synthesized HSNs were termed as bowl-structured hollow porous silica nanospheres (BHSNs). In this process, core-shell interfaces of the micelles played the key role in deposition and growth of silica (SiO\textsubscript{2}) shells on to micelles’ core. Herein, protonated the P2VP block not only acted as an acid catalyst during hydrolysis of the precursor, namely, TEOS but also initiated the condensation
of a hydrolyzed precursor by acting as a proton-transferring agent. P2VP also played a vital role via binding or capturing the silica nanoclusters onto the PS block and preventing their unwanted growth in the solution. The presence of catalytic moieties on the template’s surface led to the formation of silica shell growth on the micelles’ core surface. As a result, core-shell nanocomposites structure of polymer@SiO₂ was formed via a simple sol-gel method where the PS block of the triblock copolymer acted as the core. Consequently, a hollow structure was obtained by selective removal of the polymeric cores through the calcination process.

The morphology, structure, and composition of the BHSNs produced were determined and characterized by TEM, SEM, FTIR spectroscopy, TGA, XRD, zeta potential analyzer and by a DLS system. TEM images revealed a spherical hollow structure with an average particle size of 41.87 ± 3.28 nm, similar to average particle sizes observed in DLS analysis (44.61 nm). The average diameter of void spaces was 21.71 ± 1.22 nm and the shell thickness was 10.17 ± 1.68 nm. The average pore size was determined as 15.95 nm. SEM images further confirmed their bowl-structure with a large single pore, or opening, in shells. The large pores in shells were formed due to the prevention of silica growth on the core by the action of accumulated hydrolyzed by-products of TEOS, i.e. ethanol on PS core and slow hydrolysis process. In this case, accumulation of ethanol on micelles core was initiated via hydrophobic interaction caused by the hydrophobic nature of PS and interaction among the ethanol molecules. This established the hydrogen-bonding interaction among them as well as with the P2VP block that may have created gelation of PEO. XRD analysis also indicated a disordered mesoporous structure, i.e. the synthesized BHSNs were amorphous. The zeta values indicated that the synthesized BHSNs possess similar surface properties to amorphous silica and a highly negative charge at higher pH. It is expected that the BHSNs
produced may have advantages over HSNs for applications as a carrier for active molecules during loading via the simple immersion method due to their large pore openings.

ASSOCIATED CONTENT

Supporting Information. The following file is available free of charge. It contains further details of chemical structure of the PS-P2VP-PEO block copolymer, micelles core size measurement procedure, the catalytic effect of P2VP block during hydrolysis and condensation of TEOS, pore size measurement of synthesized BHSNs, surface area and surface pore size analysis and morphological differences of HSNs synthesized using TMOS and TEOS (PDF)

AUTHOR INFORMATION

Corresponding Author

*Ravi Naidu
E-mail: Ravi.Naidu@newcastle.edu.au; Ravi.Naidu@crccare.com; Tel: +61-02-4913 8705

Notes
The authors declare no competing financial interest.

ACKNOWLEDGMENT

The first author is grateful, firstly, to the University of Newcastle for the University of Newcastle Postgraduate Research Scholarship (UNIPRS), and to the Cooperative Research Centre for Contamination Assessment and Remediation of the Environment (CRC CARE) for scholarship funding. The first author is also grateful to University of Technology Sydney (UTS) for providing support during characterization of the samples.
ABBREVIATIONS
BHSNs, bowl-structured hollow porous silica nanospheres; CRFs, Controlled-release formulations; DLS, Dynamic light scattering; EDAX, Energy dispersive X-ray spectroscopy; FESEM, Field emission scanning electron microscopy; FTIR, Fourier transform infrared; HSNs, hollow porous silica nanospheres; MQ, Milli-Q; PEO, Polyethylene oxide; PS, Polystyrene; P2VP, Poly-2-vinyl pyridine; SCF, Supercritical Fluid; SEM, Scanning electron microscopy; TEM, Transmission electron microscopy; TEOS, Tetraethylorthosilicate; TGA, Thermo-gravimetric analysis; TMOS, Tetramethylorthosilicate; XRD, X-ray diffraction.

REFERENCES


(21) Ge, X.; Ge, X.; Wang, M.; Liu, H.; Fang, B.; Li, Z.; Shi, X.; Yang, C.; Li, G. One-Pot Synthesis of Colloidal Nanobowls and Hybrid Multipod-like Nanoparticles by Radiation


(27) Deng, Y.; Wei, J.; Sun, Z.; Zhao, D. Large-pore ordered mesoporous materials templated from non-Pluronic amphiphilic block copolymers. *Chemical Society Reviews* **2013**, *42* (9), 4054.


Karayianni, M.; Pispas, S. In *Fluorescence Studies of Polymer Containing Systems*; Procházka, K., Ed.; Springer International Publishing: Cham, 2016, DOI:10.1007/978-3-319-26788-3_2 10.1007/978-3-319-26788-3_2.


Dialysis

Hydrolysis & condensation

Drying followed by calcination at 550 °C

BHSNs

PS-P2VP-PEO block copolymer