High Throughput Synthesis of Uniform Biocompatible Polymer Beads with High Quantum Dot Loading Using Microfluidic Jet-Mode Breakup

The MIT Faculty has made this article openly available. Please share how this access benefits you. Your story matters.

<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>As Published</td>
<td><a href="http://dx.doi.org/10.1021/la4041198">http://dx.doi.org/10.1021/la4041198</a></td>
</tr>
<tr>
<td>Publisher</td>
<td>American Chemical Society (ACS)</td>
</tr>
<tr>
<td>Version</td>
<td>Author's final manuscript</td>
</tr>
<tr>
<td>Accessed</td>
<td>Thu Jun 15 22:30:02 EDT 2017</td>
</tr>
<tr>
<td>Citable Link</td>
<td><a href="http://hdl.handle.net/1721.1/92772">http://hdl.handle.net/1721.1/92772</a></td>
</tr>
<tr>
<td>Terms of Use</td>
<td>Article is made available in accordance with the publisher's policy and may be subject to US copyright law. Please refer to the publisher's site for terms of use.</td>
</tr>
<tr>
<td>Detailed Terms</td>
<td></td>
</tr>
</tbody>
</table>
High Throughput Synthesis of Uniform Biocompatible Polymer Beads with High Quantum Dot Loading Using Microfluidic Jet-Mode Breakup

Seung-Kon Lee‡§, Jinyoung Baek‡ and Klavs F. Jensen‡*

‡ Department of Chemical Engineering. Massachusetts Institute of Technology, 77 Massachusetts Avenue, 66-342, Cambridge, MA 02139 (USA)

§ Radioisotope Research Division. Korea Atomic Energy Research Institute, 989-111 Daedeok-daero, Yuseong-gu, Daejeon 305-353 (Republic of Korea)

KEYWORDS: microfluidics, jet breakup, drop breakup, hydrogels, quantum dots
Abstract

Uniform polymer microbeads with highly loaded quantum dots (QDs) are produced using high-throughput coherent jet-breakup of a biocompatible polyethyleneglycol diacrylate (PEGDA) prepolymer resin, followed by in-line photo-polymerization. A spiraling and gradually widening channel enables maximum absorption of radiated UV light for the in-line photopolymerization without coalescence and clogging issues. Although the dripping-mode in general provides superior uniformity to the jet-mode, our nozzle design with tapered geometry brings controlled jet-breakup leading to 3% of uniform particle size distribution, comparable to dripping-mode performance. We achieve a maximum production rate of 2.32 kHz, 38 times faster than the dripping mode, at a same polymer flow rate. In addition, the jet-mode scheme provides better versatility with 3 times wider range of size control, as well as the compatibility with viscous fluids that could cause pressure build-up in the microsystem. As a demonstration, a QD-doped prepolymer resin is introduced to create uniform biocompatible polymer beads with 10 wt% CdSe/ZnSe QD loading. In spite of this high loading, the resulting polymer beads exhibits narrow band width of 28 nm to be used for the ultrasensitive bio-imaging, optical coding and sensing sufficiently with single bead.
1. Introduction

There has been intense and longstanding interest in the use of biocompatible fluorescent microparticles for a variety of applications such as bio-imaging, drug delivery, biological assays, clinical diagnostics, optical encoding, lasers and display applications. Especially, particles with embedded quantum dots (QDs) have been received attention due to their useful optical properties. Batch synthesis has been the method of choice for preparing QD-doped microparticles due to practical advantages in productivity and accessibility, despite limitations such as poor loading, low throughput and material loss. In the batch loading processes, a significant portion of loading materials is wasted during the multistep process of synthesis, purification, loading, and washing. This is especially problematic when expensive QDs or biomaterials are used.

On the other hand, recent microfluidic droplet formation schemes (typically operating in the so-called dripping mode) allow production of uniform particles without the aforementioned issues. However in dripping-mode, the controllable size range is limited because the emulsion sizes are more dependent on the nozzle diameter and interfacial properties rather than the flow conditions. To get smaller emulsion sizes, dimensions of the nozzles at the breakup junction must be decreased, causing fabrication difficulties and significant pressure drop in the system. Use of viscous polymers or colloidal suspensions further increases pressure drops, which can cause device failures. A number of studies have presented ultrafast production of uniform or non-uniform water droplets in 1–10 kHz range (See Table S1), but in-line synthesis of uniform polymer beads through breakup of viscous polymers has not been reported in such a high rate. Therefore, challenges still remain for improving the productivity, widening the controlled size range and achieving smaller size without severe pressure drop.
Weber number ($We$) than corresponding to the dripping regime, the ejected liquid forms a cylindrical jet that breaks into tiny droplets by capillary instability.$^{25-29}$ Formation of emulsions and particles using jet-breakup has been studied over several decades due to its significant practical advantage in mass production over the dripping scheme.$^{25-29}$ However, difficulties in control of jet-mode uniformity have limited wide application to particle synthesis.$^{29,30}$

In this study, we demonstrate production of uniform emulsions using high throughput harmonic breakup of a jet in a high shear co-flowing system with a well-designed nozzle. Since the jet diameter is relatively small and controllable for a given nozzle diameter, the jet-mode simultaneously provides superior control over a range of diameters, moderate pressure drops, and higher production rates, compared with the dripping-mode breakup. With this scheme, we achieve uniform fluorescent microbeads through in-line photopolymerization of biocompatible prepolymer resin containing about 10 wt\% CdSe/ZnSe QD nanocrystals. In spite of this high loading, photoluminescent spectra from the microbeads present narrow emission with band width of 28 nm without broadening or distortion of the peak shape. Unlike the batch synthesis, there are no additional washing steps required for the removal of unreacted residues causing dissipation of materials.$^{4,9,31}$

2. Experimental

**Preparation of microfluidic devices:** The microfluidic chips were fabricated creating 100 μm thick and 160 μm thick SU-8 patterns by negative photolithography followed by polydimethyl siloxane (PDMS) embossing (PDMS 184-A and B, Dow Corning). The resulting PDMS replicas were bonded to 25 mm by 75 mm slide glass (Corning) covered with a thin PDMS film following oxygen plasma treatment of both surfaces.
Flow rate control: Inner and outer fluids were introduced into the microfluidic device through Polytetrafluoroethylene (PTFE) tubing and the flow rates were controlled using syringe pumps (PHD Ultra, Harvard Apparatus) and gas tight syringes (SGE). Pressure drop of the core fluid was measured from the pressurized feeding system. The pressure range was from 0.5–3 atm when the overall flow rates vary from 3–100 μL/min. Hexadecane (99%, Sigma-Aldrich) with 4 wt% of surfactants (EM 90, ABIL) was used for the outer continuous phase.

System for UV exposure: A UV lamp (FluoroArc, 100W, Carl Zeiss) mounted on an inverted microscope (Axiovert 200, Carl Zeiss) was used for the in-line photo-polymerization of the prepolymer emulsions. The power of the UV light source was adjusted to 50 W for optimized production of the microparticles.

Laser and high-speed photography: A double frame high speed camera (Imager Intense, La Vision) and two ND:YAG laser (532 nm, 30 mW, 10 Hz, 7 ns, Big Sky Laser) were used for laser photography analysis of jet-mode breakup. A fluorescent dye (Rhodamine 6G, Sigma-Aldrich) was used as a contrast agent for polyethyleneglycol diacrylate phase (PEGDA, MW 258, Sigma-Aldrich).

Interfacial tension measurement: Interfacial tension of PEGDA and hexadecane was measured using ring tensiometry (K100, Kruss) with 0.5 inch platinum ring.

3. Result and discussion

3.1. Experimental scheme

The high-throughput microfluidic particle preparation system (Figure 1) is composed of two functional parts; a jetting zone for the jet-breakup of the core fluid and a UV exposure zone for photo-polymerization of the formed droplets. In the jetting zone, hydrophilic
polyethyleneglycol diacrylate (PEGDA) is ejected from the central nozzle with shearing flows of hydrophobic hexadecane from both sides of the junction (Figure 1a and 1b). The capillary instability eventually ruptures the formed jet into uniform droplets (Figure 1c). A spiral channel with gradually increasing width maximizes UV exposure for the photo-polymerization of the prepolymer droplets while reducing coalescence (Figure 1d). The jet-mode requires short residence time and thus affords insufficient UV-exposure for the photo-polymerization, which is further challenged when large amount of QDs, fluorescent dyes, or the other light absorbing ingredients are loaded. The gradually increasing spiral channel provides continuous reduction of flow rate and correspondingly increased residence time to allow the photo-polymerization sufficient time. The diameter of the spiral channel was matched the circular UV beam from the UV microscope objective to achieve maximum exposure throughout the whole channel (left of Figure 1d).

3.2. Characterization of the high-throughput jet-breakup

High-speed imaging enables visualization and characterization of the jet-breakup process. In the channel (width \(d_c\)), a smooth jet of length \(L_j\) and diameter \(d_j\) forms at the nozzle (diameter \(d_n\)) as shown in Figure 2a. Along the jet-thread, the capillary instability induces a sinusoidal disturbance that grows and eventually causes the jet to break into a train of emulsion droplets with diameter \(d_e\) (see also the high speed movie clip, Video S1, in the Supporting Information). Dual laser imaging provides further in-depth characterization of the jet-breakup phenomena (Figure 2b).

Double exposure images were made by twin Nd:YAG laser (532 nm, 10 Hz, 7 ns pulse duration) with 500–1000 μs interval. At outer and inner flow rates of \(Q_{out} = 20 \mu\text{L min}^{-1}\) and \(Q_{in} = 0.5 \mu\text{L min}^{-1}\), respectively, the droplet speed is measured as 9.68 cm s\(^{-1}\) from the temporal
phase shift, $\varphi_e$, in Figure 2b. The spatial wavelength of the emulsion droplets, $\lambda_e$, corresponds to a production rate of 381 Hz and generation tempo of 2.626 ms. A maximum production rate of 2,320 Hz was achieved at $Q_{out} = 30 \mu L \, min^{-1}$ and $Q_{in} = 0.5 \mu L \, min^{-1}$ with an emulsion diameter of 13 µm. The temporal phase shift and spatial wavelength of the jet instability also provide information about the characteristic time scale for the development of instabilities, fluctuations of the jet length, and ultimately the jet breakup.

At the junction area, a tapered geometry with a 15° angle helps create stable shearing of the core prepolymer flow with straight and smooth features as shown in Figure 2c. (See also Figure S1) In the jet formation, it is important to produce an isolated jet confined in 3D. Wall contact would otherwise cause unstable flows with a resulting poor size distribution. Even though the jet starts from high aspect-ratio rectangular shaped nozzles (width: 20–40 µm, height: 100–160 µm), the interfacial tension drives the formation of an axisymmetric cylindrical jet confined by surrounding hexadecane flow. The cross-sectional views in Figure 2d illustrate the development of the jet.

The jet-mode behavior can be characterized in terms of the Weber number of inner flow, $We_{in}$, and the capillary number of outer flow, $Ca_{out}$. $We$ is a measure of the relative importance of the fluid's inertia compared to its surface tension, and it is defined as $We = \rho v^2L \gamma^{-1}$, where $\rho$ is density, $v$ the average velocity, $L$ is the nozzle diameter ($d_n$), and $\gamma$ is the interfacial tension. $Ca$ represents the relative effect of viscous forces versus surface tension and a defined as $Ca = \mu v \gamma^{-1}$, where $\mu$ is the fluid viscosity. With increasing $We_{in}$ and $Ca_{out}$, the mode transformed from dripping to jetting (Figure 3a). The droplet size expands with the inner flow rate (larger $We_{in}$). As the size of the droplet reaches the channel walls, the droplets slow down and begin to coalescence with each other. As we investigated the influence of the flow mode on
the particle size and production rate, jet-mode breakup exhibited a much higher production rate than the dripping-mode with much smaller particle size. (See Figure S2) As the mode changed from jetting to dripping under this specific condition, the jet-mode production rate was 208 times higher than that achieved in dripping mode with 1/6 of diameter. Here, $Q_{in}$ was fixed at 0.5 μL min$^{-1}$ while $Q_{out}$ was suddenly decreased from 30 μL min$^{-1}$ to 3 μL min$^{-1}$.

The jet-mode breakup also offers advantages compared to the dripping mode in control of particle size range (See also Figure S3). In the jet-mode, the particle diameter is determined by various parameters including nozzle size and the dimensionless numbers; $We$, $Ca$.$^{34}$ Whereas in the dripping mode, the drop size is mostly dependent on the interfacial properties and the nozzle diameter.$^{32}$ The size of the emulsions in jet-mode could be varied from 13 μm to over 100 μm over the flow rate range, but it ranged only from 52 μm to 81 μm in dripping mode (See Figure S3).

For an undisturbed jet, the ratio of emulsion size and jet diameter $d_o/d_j$ falls within the range $1.87–1.91$.$^{25,28,30}$ These values can be achieved only under a stable laminar flow condition.$^{18}$ This flow condition is difficult to achieve in conventional dispensing systems since most are operated under turbulent conditions.$^{35}$ On the other hand, a microfluidic system provides an ideal low Reynolds number environment with its small dimensions and precisely crafted geometry. The measured reduced diameters from our microfluidic system are in the expected range (Figure 3b) over a wide range of flow rates indicating a broad operating window of stable jet behavior. As a result, produced emulsion shows uniform size distribution as shown in Fig. 3c and 3d. The average diameter is 31 μm with 3 % ($±1$ μm) standard deviation. This narrow size distribution is comparable to the excellent uniformity characteristic of dripping mode while the production rate is two orders of magnitude higher at same flow rate.
3.3. Preparation of the quantum dot embedded microbeads

Uniform PEGDA emulsions made by jet-breakup were photochemically polymerized in the UV exposure zone with spiral channel. As an example, Figure 4a shows an optical micrograph of highly uniform emulsions flowing along the spiral channel without any stability issues (see also the high speed movie clip, Video S2, in the Supporting Information). The polymerized PEGDA particles were directly collected from the device outlet (Figure 4b). The scanning electron microscopy (SEM) image in Figure 4c confirms the uniformity and well-defined globular shape of the crosslinked PEGDA particles.

QD-doped microbeads have received considerable attention due to their useful optical properties for many potential applications. However, most of the developed incorporation methods are based on low yield batch process. For example, Han et al. reported diffusive transport of QDs from the solvent phase into the dispersed polymer beads and Insin et al. presented deposition of the QDs on the microbeads to prepare QD-doped particles. Since the diffusive transport of nanoparticles in a viscous prepolymer resin is slow, the diffusion based QD-embedding process requires long processing time with large amount of materials. Moreover, the efficiency decreases as the size of loading material becomes larger. On the other hand, deposition method allows adsorption of QDs merely on the particle surface, making it hard to create highly loaded particles even with multistep layer-by-layer deposition process. Moreover, in both cases, a significant portion of loading materials is wasted during the multistep process of synthesis, purification, loading, and washing. This is especially not desirable when expensive loading materials, such as QDs, are used.

In this study, we synthesized CdSe/ZnS coreshell QDs (504 nm and 560 nm emission) by continuous and batch processes to incorporate into the polymer beads. Especially, large QDs
(12 nm diameter) with thick ZnS shells were prepared and used to reduce oxidation and luminescence quenching in the polymer (See Figure S4 and S5). Such a particle with large diameter is hard to incorporate into the microparticles through diffusion process. An advantage of using this microfluidic jetting platform is the versatility of material loading within the biocompatible polymer particles. It is easy to put any combination of hydrophilic ingredient into the PEGDA monomer by simply mixing them together prior to injection, but hydrophobic materials can also be incorporated into the PEGDA after surface modification with PEG-based ligands.

Fluorescent images (Figure 4d) show uniform QD-doped fluorescent particles with 9.9 wt% of loading and 504 nm of peak emission. In general, prepolymer resin with highly loaded QDs shows significant red-shift, peak-broadening, distortion or secondary peak formation caused by agglomeration of colloidal QD nanoparticles. (See Figure S5) In this study, surface of the QDs were modified with PEG-thiols compatible and dispersible with PEGDA prepolymer. Additionally, CdSe nanocrystals coated with thick ZnS shells were introduced to prevent aforementioned issue by confining electrons even with high QD loading. A comparison of photoluminescence from QDs dispersed in chloroform, prepolymer resin and crosslinked PEGDA (Figure 4e) shows that the QD optical properties remain essentially unchanged even at high concentration of QDs, only 1.3 nm red-shift of peak emission and 0.62 nm difference of FWHM is observed (See Figure S5 for the other QDs).

We measured the solid contents of the QD solution two different ways; optical absorbance and density measurement. From the location of the first absorption peak and particle diameter, we estimated number of cores and their volumetric percentage. However, this calculation method consistently showed much higher QD loadings than the density based
estimates because it assumes highly diluted concentration of QDs. Since very high concentration of QD solutions with significant self-absorption were used, we report only values based on the density measurement. Solid contents of the particles used in the microfluidics experiments were ranged from 2.2‒3.5 vol% (7.1‒11.9 wt%). To our knowledge, this is the largest value reported for QD loaded polymer particles to date. Beyond 5.6 vol%, viscosity of the loaded prepolymer resin increased significantly and the resin completely gelled above 9 vol% (20 wt%). Compared with diluted resin, these thick resins show significant self-absorption. (See Figure S4)

Conclusions

In conclusion, we have realized coherent jet-mode followed by stable drop breakup in a microfluidic system with a nozzle designed for the rapid production of biocompatible prepolymer emulsions loaded with fluorescent QDs. A spiraling and gradually widening channel enabled maximum absorption of radiated UV light for the in-line photopolymerization without coalescence and clogging issues. Compared to the often used dripping mode, the developed microfluidic jet-breakup scheme provided advantages in an expanded range operating conditions from 13 µm to 110 µm with over 38 times higher productivity at 2.32 kHz while still retaining a narrow size distribution of 3 %. By using large CdSe/ZnS QDs with PEG-terminated ligand well dispersible in PEGDA, uniform biocompatible microparticles with more than 10 wt% of very high QD loadings were achieved. The final QD load polymer particles had a 28 nm of narrow emission band width at 504 nm and 560 nm without significant peak broadening or distortion.
Figure 1. Gallery of the high performance microparticle generator. The design is optimized for coherent and high rate jet-breakup and subsequent photo-polymerization to produce uniform fluorescent microbeads in continuous manner. (a) Schematic of the jetting zone with 40×100 μm nozzle in the center for hydrophilic PEGDA ejection. Hexadecane is introduced through the cross channels. (b) Overall chip design (left, scale bar = 3 mm) and nozzle design (right, scale bar = 200 μm) from CAD drawing. (c) Optical micrograph of jet-breakup corresponds to the
schematic in (a). Scale bar is 100 μm. (d) UV exposure zone with gradually widening spiral flow channel for the in situ polymerization of PEGDA. Left image shows spot of UV exposure on the overall spiral channel. Scale bar is 1000 μm. Right image shows highly uniform emulsions flowing along the spiral channel without any stability issues (at $Q_{out} = 50 \, \mu\text{L} \, \text{min}^{-1}$, $Q_{in} = 0.5 \, \mu\text{L} \, \text{min}^{-1}$).
Figure 2. High-speed images and dual-pulsed laser photography images for the characterization of the jet-mode at $Q_{out} = 20$ μL min$^{-1}$, $Q_{in} = 0.5$ μL min$^{-1}$. (a) High-speed image showing characteristic features of jet-mode breakup mechanism as a series of events; ejection, elongation, instability build-up and breakup. Jet parameters are emulsions diameter, $d_e$, jet diameter, $d_j$, channel diameter, $d_c$, nozzle diameter $d_n$, disturbance wavelength, $\lambda_j$ and jet length, $L_j$. (b) Dual-pulsed laser photography provides information about the spatial interval ($\lambda_e$, $\lambda_j$) and temporal shift ($\varphi_e$, $\varphi_j$) of emulsion trains and jet (red first image, green second image). (c) Magnified image of nozzle area designed for the shaping of a stable jet. By shaping the edge of cross-channel (red triangle), the shearing flow gets smaller adjacent angle improving the stability of the jet. (d) Schematic corresponding to the blue box of (c) showing how the jet is formed and remain isolated from channel walls.
Figure 3. Behavior of jet with fluid parameters. (a) Flow phase diagram plotted with $We_{in}$ versus $Ca_{out}$. At 0.01–0.1 of $Ca_{out}$, the flow transitions from dripping (triangle) to jetting (black dot). The jet-mode region expands with increasing $We_{in}$, but above a certain value of $We_{in}$ (~$10^{-5}$), an unstable breakup pattern (crossed circle) appears. (b) Plot of reduced dimensionless diameters ($d_e/d_j$) over the various relative volumetric flow rates ($Q_{in}/Q_{out}$). The data converge around the expected value from theoretical studies, 1.87, indicating that the jet-mode is stable over a wide range of flow conditions. (c) Optical micrographs of highly uniform emulsions flowing along the microchannel (at $Q_{out} = 30$ μL min$^{-1}$, $Q_{in} = 0.5$ μL min$^{-1}$). (d) Size distribution of the produced microparticles as narrow as 3% (31±1 μm).
Figure 4. Images and characteristics of the polymerized microparticles. (a) Uniform 24 μm PEGDA particles flowing along the spiral channel. (at $Q_{out} = 50 \mu$L min$^{-1}$, $Q_{in} = 0.5 \mu$L min$^{-1}$)

(b) Transmission optical micrograph of the uniform PEGDA microbeads. (c) SEM image of PEGDA microbeads verifying highly uniform sizes. (d) Pseudo-color fluorescent image of highly monodisperse PEGDA polymer beads with 9.9 wt% of highly loaded CdSe/ZnS coreshell nanocrystals. The inset shows a real color image from the same QD-doped particles. (e) Photoluminescence spectra from QDs which are dispersed in chloroform (black), PEGDA monomer (red) and polymerized PEGDA (green), respectively. The QD loaded polymer beads have strong emission at 504 nm with 28 nm narrow FWHM (full-width half-maximum). The insert shows that only a small 1.3 nm red-shift and 2 % widening in FWHM in the solvent.
exchange from chloroform to PEGDA. Moreover, no shift or widening of the peak is observed during photo-polymerization.

AUTHOR INFORMATION

Corresponding Author

*Department of Chemical Engineering. Massachusetts Institute of Technology, 77 Massachusetts Avenue, 66-342, Cambridge, MA 02139 (USA), Fax: +1-617-258-8992; Tel: +1-617-253-4589; E-mail: kfjensen@mit.edu

Author Contributions

The manuscript was written through contributions of all authors. All authors have given approval to the final version of the manuscript.

ACKNOWLEDGMENT

The authors thank the US Army Research Office through the Institute for Soldier Nanotechnology (DAAD-19-02-0002) for support. We thank Dr. S.-H. Kim at KAIST for the tensiometry experiments. SKL would like to dedicate this work to Prof. Seung-Man Yang, a great scientist and a mentor, who passed away on September 26th, 2013.

SUPPORTING INFORMATION

Supporting Information and video clips are available free of charge via the Internet at http://pubs.acs.org.
REFERENCES


Supporting Information

High Throughput Synthesis of Uniform Biocompatible Polymer Beads with High Quantum Dot Loading Using Microfluidic Jet-Mode Breakup

by Seung-Kon Lee, Jinyoung Baek and Klavs F. Jensen*

E-mail: kfjensen@mit.edu

Table S1 Comparison of high throughput microfluidic emulsion production system

<table>
<thead>
<tr>
<th>Developer</th>
<th>Device material</th>
<th>Maximum production rate [kHz]</th>
<th>Core flow rate at for maximum production rate [µL/min]</th>
<th>Operation mode</th>
<th>Uniformity</th>
<th>Production material</th>
</tr>
</thead>
<tbody>
<tr>
<td>J.-Y. Kim et al. [a]</td>
<td>Thermoset polyester</td>
<td>1.73</td>
<td>20</td>
<td>Dripping</td>
<td>Good</td>
<td>Water emulsion</td>
</tr>
<tr>
<td>J. J. Agrestia et al. [b]</td>
<td>PDMS</td>
<td>2</td>
<td>6.67</td>
<td>Dripping</td>
<td>Good</td>
<td>Water emulsion</td>
</tr>
<tr>
<td>Shelley L. Anna et al. [c]</td>
<td>PDMS</td>
<td>10</td>
<td>-</td>
<td>Dripping/Jetting</td>
<td>Varies [d]</td>
<td>Water emulsions</td>
</tr>
<tr>
<td>C. Holtze et al. [e]</td>
<td>PDMS</td>
<td>10</td>
<td>1.67</td>
<td>Dripping</td>
<td>Good</td>
<td>Water emulsions</td>
</tr>
<tr>
<td>Adam R. Abate et al. [f]</td>
<td>PDMS</td>
<td>6</td>
<td>1.67</td>
<td>Jetting [g]</td>
<td>Good</td>
<td>Water emulsions</td>
</tr>
<tr>
<td>S. –K Lee et al.[h]</td>
<td>PDMS</td>
<td>2.32</td>
<td>0.5</td>
<td>Jetting</td>
<td>3%</td>
<td>PEGDA prepolymer emulsion/ Polymer particles</td>
</tr>
</tbody>
</table>

[d] Varies according to the flow rate condition
[g] Bubble triggered jet-breakup
[h] Present work
Figure S1 Effect of proper design for the formation of stable jet-flow. a) By applying a shaping zone around the nozzle, a more stable jet-stream is produced. b) Without the shaping zone, the side flows destabilize the jet.

Figure S2 Fluorescent image of the crosslinked PEGDA microbeads encoded with QD nanoparticles taken during the phase transition from jetting (right side) to dripping (left side). Inset (left) shows the location of the image was taken. Magnified inset image (right) illustrates size and productivity of microbeads generated in jet-mode when compared with those produced in dripping mode (left). Scale bar is 100 μm.
Figure S3 Variation of emulsion diameter with the relative flow rate change. The emulsion size can be controlled from 13 μm to over 100 μm of emulsion size. After polymerization, the smallest size becomes 10 μm with a 20–30% of shrinkage. The scatter is attributed to interactions with the wall. When the emulsion size (or jet diameter) is small, the disturbance is reduced. In dripping mode (magenta hexagons denoted with arrows), size of the produced droplets range from 52 μm to 81 μm. The controllable range with jet-mode breakup is ~3 times wider than that of dripping mode.
Figure S4 Photograph of QD loaded PEGDA prepolymer resin placed on the UV lamp and the TEM image of QDs. a) Fluorescent prepolymer resin doped with CdSe/ZnS QDs with 504 nm emission peak. From the left to right, concentration increased from 1, 3.5 to 9 vol%. Over 5 vol% (20 wt %) the resin starts to gel because of colloidal crystallization driven by high inter particle interaction. The reduced emission with increasing loading results from self-absorption from highly concentrated QDs among the prepolymer resin. b) Transmission electron microscopy (TEM) image thick shell CdSe/ZnS showing uniform size with an average diameter of 12 nm.

Figure S5 Photoluminescence spectra from QD-doped PEGDA polymers with and without thick protective shells and PEG-based surface modification. (a) CdSe/ZnS coreshell QDs with thick ZnS shells and PEG-thiol ligand. It shows smooth and narrow emission at 554.4 nm with a band width of 28 nm. (b) CdSe/ZnS coreshell QDs with thin ZnS shells without surface modification. Due to the agglomeration and phase separation of QDs in PEGDA resin, it shows significant redshift to 573 nm, wider band width of 55 nm and secondary peak formation at 610 nm. Signal to noise ratio is also poor due to the low solubility of QDs in PEGDA without compatible surface modification.