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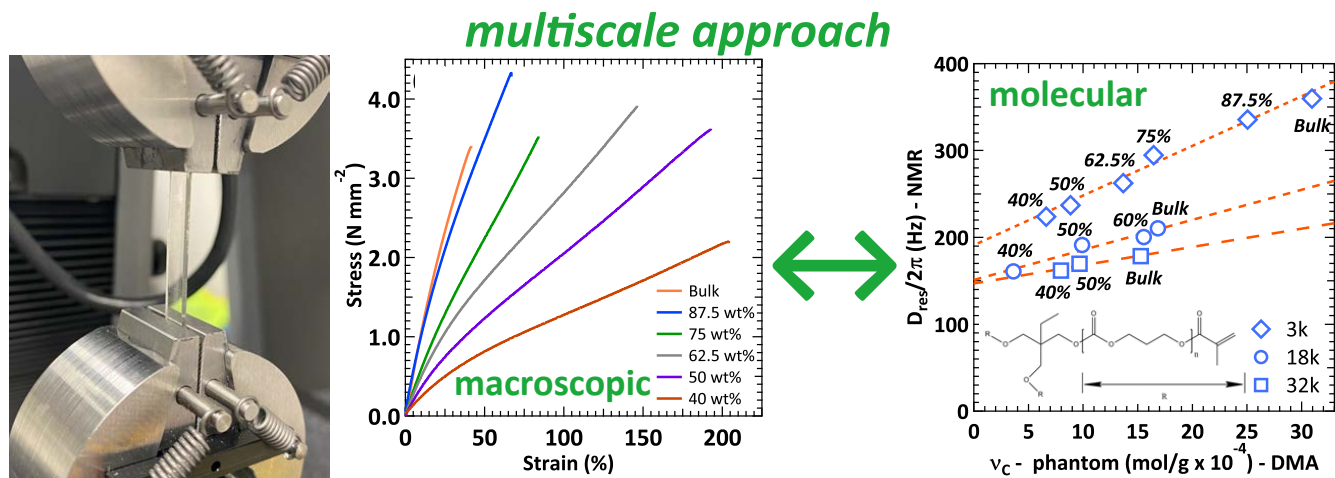
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# Graphical Abstract

## Multiscale Structural Characterization of Biocompatible Poly(trimethylene carbonate) Networks Photo-cross-linked in a Solvent

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# Highlights

## **Multiscale Structural Characterization of Biocompatible Poly(trimethylene carbonate) Networks Photo-cross-linked in a Solvent**

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- Biocompatible Poly(trimethylene carbonate) crosslinked polymers have been successfully synthesized in solvent containing various polymer concentrations.
- DMA and Time Domain  $^1H$   $DQ$  NMR data was in good correlation, relating to network crosslink density and morphology.
- The PTMC crosslink density varies linearly with polymer content in the reactive media.
- The presence of physical entanglements mechanically reinforces PTMC samples of low polymer content and/or high macromer molecular weight.
- A Multiscale approach based on Mechanical testing, DMA, and Time Domain  $^1H$   $DQ$  NMR was robustly used to characterize the influence of polymer network morphology on their macroscopic mechanical behavior.

# Multiscale Structural Characterization of Biocompatible Poly(trimethylene carbonate) Networks Photo-cross-linked in a Solvent

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## Abstract

Poly(trimethylene carbonate) (*PTMC*) polymeric networks are biocompatible materials with potential biomedical applications. By combining Dynamic Mechanical Analysis (DMA), Solid State Nuclear Magnetic Resonance (NMR), and tensile testing, it was possible to fully characterize the inner structure and its relationship with the macroscopic properties of photo-crosslinked *PTMC* materials in a solvent medium. *PTMC* prepared from macromer with various molecular weights (3 kg/mol, 18kg/mol, and 32 kg/mol) and with various polymer concentrations within the reactive media were analyzed, with the variation of thermomechanical properties and NMR signal decay characterized as a function of both aforementioned synthesis parameters. DMA and solid state Double Quantum (*DQ*)  $^1H$  NMR investigations demonstrated that the network crosslink density is directly related to the macromer molar mass and polymer concentration. More interestingly, tensile tests confirmed that mechanical behavior depended on the materials' inner structure, notably their crosslink density. Specifically for the 18 kg/mol *PTMC* networks, dangling and free chains reinforced the network, exemplified by higher Young's modulus  $E$  values. This multiscale investigation provides a promising and precise approach to tailor the macroscopic behavior of *PTMC*

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materials, by controlling their specific inner network structural morphologies during synthesis.

*Keywords:* Time-Domain  $^1H$  DQ NMR, Dynamic Mechanical Analysis, Biocompatible polymers, Thermomechanical properties, Structure-properties relationship, Poly(trimethylene carbonate)

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## 1. Introduction

Biodegradable polymer networks have great potential in biomedical applications due to their biocompatibility and biodegradability[1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15]. More specifically, tough elastomeric polymer networks are of interest as bulk materials[12, 16, 17, 18, 19, 20, 21]. Elastomeric polymer networks can be designed to exhibit mechanical properties required for their intended application, especially soft tissues[22]. Much work has been done to improve the toughness of the elastomeric networks, but it remains a challenge to obtain elastomers that are both degradable and tough[18].

To improve the toughness of elastomeric networks, the growth of micro-cracks must be hindered[23]. This has been extensively studied for both natural and synthetic elastomers, though many of these studies have been performed with non-degradable elastomers such as polyisoprene and poly(dimethyl siloxane) (PDMS)[24, 25, 26, 27, 28, 29]. The formation of micro-cracks in these networks was slowed down by introducing crystallizable domains into the polymer network, crosslinking in solvent, and crosslinking chains with bimodal chain length distributions.

Tough, biodegradable elastomeric networks have been prepared via the photo-crosslinking of methacrylate-functionalized *PTMC* oligomers (*PTMC* macromers) and acrylated poly( $\epsilon$ -caprolactone-co-D,L-lactide) (P(CL-co-DLLA)) oligomers[16, 18, 30]. Such networks are formed by adding a photo-initiator which forms radical species upon light irradiation [31]. These radical species can initiate polymerization of the (meth)acrylate end-groups of the macromers, creating poly(methacrylate) kinetic chains as the networks are formed. In the case of *PTMC*-networks, toughness increased with macromer molecular weight[16]. In addition, the preparation of bimodal *PTMC* networks resulted in a significant increase in toughness[19]. For both *PTMC*- and *P(CL-co-DLLA)* macromers, crosslinking in solution resulted in less rigid networks that demonstrated increasing elongation with decreasing macromer concentrations, resulting in superior elastomer toughness[18, 30].

It has been suggested that the presence of solvent during crosslinking results in disentangled chains prior to crosslinking formation, which in turn leads to simpler topologies in the networks[25]. The crosslinks of such networks are more likely to be both spatial and topological neighbors as

compared to networks prepared in bulk. In addition, networks prepared in solution have fewer chain-junction and inter-chain entanglements. The polymer chains in these networks are less-firmly embedded in the network structure; subsequently these networks deform more non-affinely. In order to fully understand the influence of crosslinking in solution on the macroscopic properties of elastomeric networks, the thermomechanical behavior and structural morphology of these networks need to be fully studied.

To facilitate this, a macroscopic experimental approach combining tensile testing, DSC and DMA analyses, and Time Domain  $^1H$  Double Quantum  $DQ$  NMR was undertaken in this work. Tensile tests yielded the macroscopic mechanical resistance of the studied  $PTMC$  samples, while DSC and DMA allowed the evolution of molecular mobility to be analyzed via observation of the glass transition temperature. Moreover, DMA allowed the characterization of the mechanical crosslink density of  $PTMC$  networks with varying molecular weight and solvent concentrations.[32, 33]. These studies were complemented by Time Domain  $^1H$   $DQ$  NMR. This specific technique has been principally used to characterize the structure, morphological organization and molecular mobility of polymeric networks [34, 35, 36, 37, 38, 39, 40, 41, 42, 43, 44, 45]. Double-Quantum  $DQ$   $^1H$  sequences have been proven to effectively characterize elastomeric-like polymer networks, specifically their molecular mobility, crosslink density  $\nu_C$ , and chain defect concentration  $w_{DEF}$ [46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57], in addition to the evolution of these properties with temperature [58, 59], chemical [60, 61, 62] and physical modifications[63], and thermal ageing [64, 65]. This is made possible due to the ability of this technique to discriminate dynamical and structural effects, permitting semi-local structural features of networks to be recovered from local dynamical measurements.

More extensively, Time Domain  $^1H$   $DQ$  NMR has been efficiently used in combination with DMA analyses to study the relationship between the structure of  $PTMC$  networks of various macromer molar masses, and their macroscopic thermomechanical behavior[66]. Therefore, this work is a logical evolution of this previous study. By applying the same approach to analyse the influence of polymer concentration during  $PTMC$  network formation, we aim to deepen the understanding of the influence of intrinsic polymer structure on the macroscopic behavior of  $PTMC$  networks, which in turn will provide deeper insight regarding how to chemically tailor network structure, so as to better regulate and adjust  $PTMC$  specific macroscopic properties.  $PTMC$  networks were chosen for this study, as such networks are extensively studied for a range of biomedical

applications requiring different properties, such as intervertebral disks [67], meniscus implants [17] and bone implants [20]. Its biocompatibility has been shown previously with, among others, synovium derived cells [68], mesenchymal stem cells [69], and chondrocytes [70] *in vitro* and in several studies *in vivo* [68, 71, 72].

## 2. Materials

Trimethylene carbonate (TMC) monomer was purchased from Huizhou ForYou Medical Devices Co. (China). Hydroquinone, methacrylic anhydride, tin(II) 2-ethylhexanoate ( $\text{Sn}(\text{Oct})_2$ ), Trimethylol propane (TMP), and triethylamine were purchased from Sigma (USA) and used as received. Dichloromethane and chloroform were obtained from Merck (Germany), and *d*-chloroform was purchased from VWR. Ethanol was obtained from Altia oyj (Finland). TPO-L (2,4,6-trimethylbenzoylphenyl phosphinate) was obtained from Carbosynth Limited (United Kingdom).

### 2.1. Macromer Synthesis

To obtain star-shaped methacrylate-functionalized *PTMC* oligomers (macromers), hydroxyl-terminated *PTMC* oligomers were first prepared by ring-opening polymerization reactions of TMC at 130 °C under a nitrogen atmosphere for 3 days using TMP as initiator and  $\text{Sn}(\text{Oct})_2$  as catalyst. Oligomers with different molecular weights ( $\overline{M}_n$ ) could be prepared by adjusting the monomer-to-initiator molar ratio. Second, the oligomers were dissolved in dichloromethane (2 mL/g oligomer) and functionalized with methacrylic anhydride (7.5 mol/mol oligomer) in the presence of triethylamine (7.5 mol/mol oligomer). Hydroquinone (0.1 wt% relative to the monomer) was added to the solution to act as a free-radical scavenger to prevent premature crosslinking. The macromers (*PTMC-tMA*) were precipitated in cold ethanol after 5 days and dried at 40 °C under vacuum for 1 week. The  $\overline{M}_n$  of the obtained oligomers, the monomer conversion, and the degree of functionalization were determined by  $^1\text{H-NMR}$  as described previously.[17] Briefly, the molecular weight was determined by comparing the area of the  $-\text{CH}_3$  group of the initiator peak at  $\delta = 0.92$  ppm with the area of the *PTMC* methylene peak at  $\delta = 4.24$  ppm. The conversion was calculated by comparing the area of the *PTMC* methylene peak at  $\delta = 4.24$  ppm with the area of the TMC monomer peak at  $\delta = 4.45$  ppm. The degree of functionalization was determined by comparing the  $-\text{C}=\text{CH}_2$   $^1\text{H}$  signals at  $\delta = 5.58$  ppm and  $\delta = 6.13$  ppm of the methacrylate groups with the  $-\text{CH}_3$  initiator peak.

## 2.2. Network Formation

To photo-crosslink *PTMC* in bulk (also called 100% hereafter), the macromers were dissolved in chloroform and 5 wt% TPO-L photo-initiator (relative to the macromer) was added. The solutions were cast in teflon molds and kept in the dark overnight to allow for chloroform evaporation.

To photo-crosslink *PTMC* in solution, the macromers were dissolved in propylene carbonate at 70 °C. The polymer solutions contained 87.5, 75, 62.5, 50 or 40 wt% polymer. After complete dissolution, 5 wt% TPO-L photo-initiator (relative to the macromer) was added to the solutions. The solutions were cast onto a glass plate to obtain networks with an approximate thickness of 1 mm.

The cast macromers were crosslinked at room temperature under nitrogen in a custom built cross-link box for 30 min at 395-405 nm at an intensity of 1 mW/cm<sup>2</sup>. The obtained networks were subsequently post-cured under visible light for 40 min at room temperature. The conversion of the  $-\text{C}=\text{CH}_2$  double bond was confirmed by ATR-IR (Spectrum Two, Perkin Elmer). As these networks have a functionality  $f=3$ , the theoretical chemical cross-link density  $v_{chem}$  of such networks can be calculated as  $1/3\overline{M}_n$ .

## 3. Experimental Methods

### 3.1. Swelling Characterization

The volume degree of swelling  $q$  was determined in triplicate at room temperature by swelling rectangular shaped specimens ( $5 \times 5 \times 1$  mm<sup>3</sup>) in chloroform for 24 h, which was sufficient time to reach solvent sorption equilibrium. The  $q$  was calculated from Equation 1

$$q = 1 + \left( \frac{m_{swollen} - m_{dry}}{m_{dry}} \right) \left( \frac{\rho_p}{\rho_s} \right) \quad (1)$$

where  $m_{swollen}$  is the mass of the swollen networks,  $m_{dry}$  the mass of the networks after drying, and  $\rho_p$  and  $\rho_s$  the densities of *PTMC* ( $=1.31$  g/cm<sup>3</sup>)[16] and chloroform ( $=1.48$  g/cm<sup>3</sup>), respectively.

### 3.2. DSC Analysis

The thermal properties of the obtained *PTMC* networks were determined by a TA Instruments Q2000 differential scanning calorimetry (DSC). Samples with weights between 5 and 10 mg were cooled to -90 °C and subsequently heated to 100 °C at 10 °C/min. After a second cooling cycle



to  $-90\text{ }^{\circ}\text{C}$  at  $10\text{ }^{\circ}\text{C}/\text{min}$  the samples were held at  $-90\text{ }^{\circ}\text{C}$  for 5 min and a second heating scan was performed. The glass transition temperatures ( $T_g$ ) were determined from the second heating scan. Temperature calibration was performed using indium as a calibration standard.

### 3.3. Tensile experiments

The tensile properties of the networks were determined by tensile testing using an Instron universal tensile tester 5944, equipped with a 2 kN load cell at  $24\text{ }^{\circ}\text{C}$  and 50% relative humidity. The tensile samples dimensions were  $5.3 \times 40\text{ mm}^2$ . The elongation of the samples was determined from the grip-to-grip separation which was initially 20mm. The tensile modulus ( $E$ ), maximum tensile strength ( $\sigma_{max}$ ) and elongation at break ( $\varepsilon_{break}$ ) were determined according to ASTM D882 at a test speed of 10mm/min. The yield stress ( $\sigma_{yield}$ ) and elongation at yield ( $\varepsilon_{yield}$ ) were determined from the intersection of the tangents to the stress strain curve. The toughness ( $W$ ) was determined by calculating the area under the stress-strain curve. Five specimens of each type of network were used for the tensile testing.

### 3.4. Dynamic Mechanical Analysis

Dynamic Mechanical Analyses were performed using a TA Instruments Q800 DMA operating in tensile mode. *PTMC* samples were cut into ISO 527-4b dogbone-shaped specimens with dimensions of  $18 \times 2 \times 0.7\text{ mm}^3$ . Samples were heated from  $-140^{\circ}\text{C}$  to  $150^{\circ}\text{C}$  with a heating rate of  $3^{\circ}\text{C}/\text{min}$  under a frequency of 1 Hz, a pre-strain of 0.01%, and a subsequent strain of 0.1%. The main  $\alpha$  relaxation temperature  $T_{\alpha}$  was calculated from the half-height point of  $E'$  drop corresponding to this relaxation [73]. All values are shown as averages  $\pm$  standard deviation of 5 repeats.

Furthermore, the crosslinking density  $\nu_{C-DMA}$  for each *PTMC* network was obtained by conducting a DMA strain sweep measurement at  $T_{\alpha} + 90^{\circ}\text{C}$  so as to allow a precise comparison between these results and those obtained by NMR at the same molecular mobility state. The linear regime was thus determined by plotting the storage modulus  $E'$  as a function of the strain  $\epsilon$  and the value of  $E'$  was taken from the linear regime plateau. Then, the crosslinking density  $\nu_{C-DMA}$  was calculated according to Equation 2[32].

$$\nu_{C-DMA} = \frac{E'}{\phi RT f} \quad (2)$$

where  $R$  is the ideal gas constant =  $8.314\text{ J/mol} \cdot \text{K}$ ,  $T=T_{\alpha} + 90^{\circ}\text{C}$ ,  $f$  is the network functionality which for *PTMC* samples is equal to 3, and  $\phi$  is a factor linked to the network model. For

the *affine* model[74]  $\phi = 1$ , whereas for the *phantom* model [75]  $\phi = \frac{f-2}{f}$ . In this work, two series of  $v_{C-DMA}$  values were thus calculated according to both the affine and phantom models.

### 3.5. DQ $^1H$ Solid State NMR Measurements

$^1H$  Double Quanta DQ experiments were conducted on a Bruker Avance III 400 NMR equipped with a 5mm  $^1H$  Static probe. These experiments are based on Baum - Pines pulse sequences[47, 48] optimized by Saalwächter[49, 50, 51, 52, 53, 56, 54, 57, 58, 59, 65]. Such experiments yield two components as a function of the DQ evolution time  $\tau_{DQ}$  : the DQ buildup  $I_{DQ}$  and the reference decay  $I_{REF}$ . Figure SI.1 (*Support Information*) shows an example of such signals. The full magnetization of the sample  $I_{TOT}$  corresponds to the sum of  $I_{DQ}$  and  $I_{REF}$ , and comprises the response of both the dipolar coupled network and the uncoupled mobile network defects such as dangling or free chains. These fractions are characterized by different types of relaxation behavior. Chains fully belonging to the network relax faster and non-exponentially, whereas dangling and free (*i.e.* non-elastic) chains exhibit a slower exponential relaxation.

As described previously[66], in order to properly get access to the network structure, DQ experiments must be undertaken in the temperature-independent regime, *i.e.* the probed molecular motions must be effective in the fast motion regime. Preliminary studies on *Bulk 3k PTMC* were carried out at various temperatures ranging from  $T_\alpha + 50$  C° to  $T_\alpha + 100$  C°. Such measurements showed that the temperature-independent was reached for  $T_\alpha + 90$  °C (see Equation SI.1 and Figure SI.2 (*Support Information*)). In this work, all of the samples were thus studied at  $T = T_\alpha + 90$  °C, ensuring that they were all tested at the same state of molecular mobility in the temperature-independent regime of the DQ signal.

In this temperature-independent regime, to discriminate the network structure influence on the DQ build-up, a normalization of this signal has to be carried out. This normalized signal  $I_{nDQ}$  is calculated according to Equation 3 and theoretically it must reach a plateau at the value 0.5[62, 76].

$$I_{nDQ} = \frac{I_{DQ}}{I_{REF} + I_{DQ} - I_{DEF}} \quad (3)$$

This normalization (*i.e.* Equation 3) considers the subtraction of the non-elastic chains  $I_{DEF}$  (*i.e.* dangling or free chains) contribution from the total signal. A heuristic approach readily allowing the identification and subtraction of this contribution is to fit  $I_{DEF}$  through a double exponential from the  $I_{REF} - I_{DQ}$  *v.s.*  $\tau_{DQ}$  signal [53, 62]. Extrapolating this  $I_{DEF}$  fit to  $\tau_{DQ} = 0$

gives the percentage of defects (*i.e.* non-elastic chains)  $w_{DEF}$ . An example of this approach is given in Figure SI.1 (*Support Information*).

$DQ$  NMR experiments give access to the dipolar coupling constant  $D_{res}$ , which is related to an average local dynamic segmental orientation parameter.  $D_{res}$  is in turn related to the crosslink density  $\nu_C$  through Equation 4:

$$\nu_C \propto \frac{1}{M_C} \propto k \frac{D_{res}}{D_{stat}} \quad (4)$$

where  $D_{stat}$  is the static dipolar coupling constant and  $k$  a proportionality factor related to the details of intersegmental motions at the scale of the Kuhn length. Then, the numerical value of  $D_{res}$ , related to the network structure, is obtained by fitting the corresponding  $I_{nDQ}$  signal up to 0.48 by Equation 5 (See Figure SI.3 - *Support Information*).

$$I_{nDQ} = 0.5 [1 - \exp(-D_{res}\tau_{DQ})^n] \quad (5)$$

where  $n$  is an exponent varying between 1 and 2. The closer  $n$  is to the value of 2, the more homogeneous the networks are. In this investigation, the values for  $k$  and  $D_{stat}$  were not obtained, nevertheless these factors should be identical for all samples. Thus, a quantitative comparison between all studied materials can be performed as described in Equation 4 by obtaining  $D_{res}$ .

#### 4. Results and discussion

Three-armed *PTMC* macromers were synthesized via the ring-opening polymerization of *TMC* into *PTMC* oligomers followed by functionalization with methacrylic anhydride. By adjusting the monomer-to-initiator ratio, oligomers with three different molecular weights were obtained. Table 1 shows the obtained oligomer molecular weights as confirmed by  $^1\text{H-NMR}$ . The subsequent functionalization resulted in macromers with a degree of functionalization of  $\geq 86\%$ . In Figure SI.4 (*Support Information*), a typical  $^1\text{H-NMR}$  spectra of a *PTMC* oligomer and macromer is shown. Detailed interpretations of such spectra can be found elsewhere[16]. The chemical structure of the *PTMC* macromer is shown in Figure 1.

*PTMC* networks were prepared by solvent casting and subsequent photo-crosslinking resins containing 6 different concentrations of polymer. For the two highest molecular weights, some polymer concentrations were too viscous to be cast, even though the resins were heated to 70 °C. The conversion of the  $-\text{C}=\text{CH}_2$  double bond was confirmed by ATR-IR. The methacrylate double

Table 1: Conversion and Degree of Functionalization of the Obtained *PTMC* Macromers

$M_n$ (g/mol)	Conversion (%)	Degree of Functionalization (%)
3100 ( <i>3k</i> )	97	100
18000 ( <i>18k</i> )	99	100
32400 ( <i>32k</i> )	97	86

bond ( $1640\text{ cm}^{-1}$ ) disappears when it polymerizes during the photo-crosslinking, as can be seen in Figure 2. The obtained networks were extracted and dried prior to characterization.

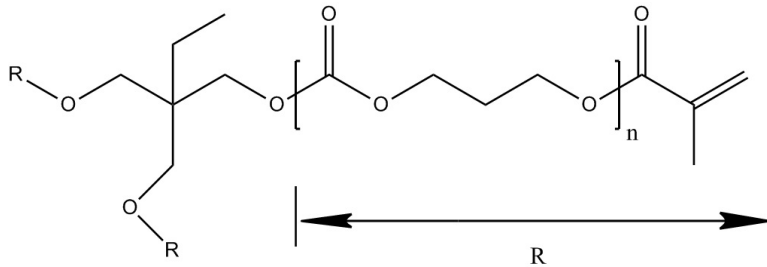


Figure 1: Chemical structure of *PTMC* macromer.

The glass transition temperatures  $T_g$  of the obtained networks were determined by DSC and are shown in Table 3. The trend in the  $T_g$  values of the networks is similar to that observed by Schüller et al[16]: with increased macromer molecular weight the,  $T_g$  decreases. While slight differences can be observed between the solvent-cast networks prepared with the same molecular weight macromers, no correlation between the obtained  $T_g$  values and polymer concentration during cross-linking is observed. In addition, Table 3 shows that the degree of swelling  $q$  increased not only with macromer molecular weight, but also with decreasing polymer content. This indicates that when cross-linking in the presence of solvent, the cross-link density seems to decrease with polymer concentration.

Zant et al. previously demonstrated that the elongation, and thus the toughness, of extracted and dried *PTMC* networks prepared from two-armed *PTMC* macromers with a molecular weight of 10 kg/mol was strongly dependant on the amount of solvent present at the time of crosslinking [18]. Decreasing the polymer concentration within the reactive media resulted in the elongation and toughness significantly increasing. We expected this to be the same for our extracted and dried networks prepared from three armed macromers, as it was previously shown that by decreasing the

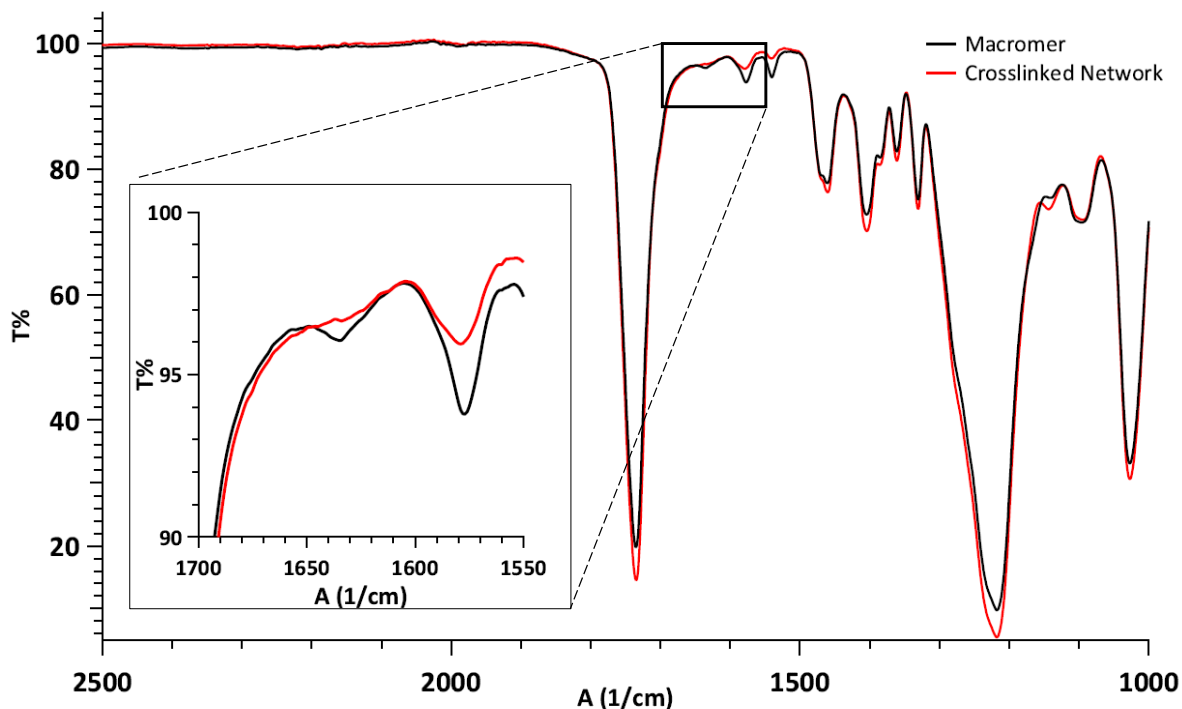


Figure 2: Typical IR spectra of a *PTMC* macromer and its subsequently obtained network. The insert shows the disappearance of the methacrylate double bond at  $1640\text{ cm}^{-1}$  which occurs due to the photo-crosslinking.

network density by means of increasing the three-armed macromer molecular weight, the elongation and toughness of the networks was increased [66, 16]. And as shown above, our networks exhibited a decrease in network density with decreasing polymer content. To confirm this, five samples per network were cut from the network films and subjected to tensile testing. An overview of the results can be found in Table 2.

Indeed, within each molecular weight, the elongation at break increases with decreasing polymer concentration. Figure 3a illustrates this trend showing typical stress-strain curves of *PTMC 3k* networks. Figure 3b further highlights this trend for all networks within this study. The toughness of the networks generally increases with decreasing macromer content, except for networks prepared from the two lowest polymer concentrations which utilized macromers with a molecular weight of *3k*; this is shown in Figure 3c. It appears that the considerable decrease of modulus for these two networks results in a decrease in toughness. The networks prepared from *PTMC 3k* are the only networks that show such a considerable decrease in modulus through the series, as exemplified in Figure 3d.

DMA analyses yielded the main transition temperature  $T_{\alpha}$  as well as the elastic modulus  $E'$  at  $T_{\alpha}+90^{\circ}\text{C}$  for each *PTMC* sample. These values are listed in Table 3. From these values

Table 2: Tensile properties of the studied *PTMC* networks.

$M_n$ (g/mol)	Polymer (wt%)	$E - modulus$ (MPa)	$\sigma_{yield}$ (N/mm <sup>2</sup> )	$\sigma_{max}$ (N/mm <sup>2</sup> )	$\epsilon_{yield}$ (%)	$\epsilon_{break}$ (%)	$W$ (N/mm <sup>2</sup> )
3k	100	12.2 ± 0.49	1.38 ± 0.23	3.19 ± 0.42	12.4 ± 2.75	42.0 ± 7.83	77.3 ± 21.7
	87.5	11.6 ± 0.54	1.47 ± 0.40	4.20 ± 0.76	13.8 ± 4.54	66.2 ± 12.6	158 ± 44.7
	75	8.50 ± 0.92	1.04 ± 0.26	3.68 ± 0.91	13.7 ± 3.76	81.4 ± 18.7	170 ± 79.2
	62.5	6.27 ± 1.37	1.03 ± 0.15	4.26 ± 1.01	17.7 ± 5.75	154 ± 26.6	370 ± 131
18k	50	4.13 ± 0.81	0.78 ± 0.19	3.27 ± 0.62	20.7 ± 9.99	181 ± 33.9	338 ± 114
	40	2.45 ± 0.63	0.63 ± 0.08	2.20 ± 0.24	30.5 ± 11.0	196 ± 14.4	237 ± 34.1
	100	2.45 ± 0.22	0.86 ± 0.14	1.47 ± 0.24	39.1 ± 7.73	137 ± 24.2	127 ± 42.7
	62.5	3.98 ± 0.58	0.92 ± 0.12	1.92 ± 0.38	23.3 ± 1.21	213 ± 63.0	270 ± 118
32k	50	3.34 ± 0.51	0.95 ± 0.13	1.64 ± 0.31	29.5 ± 4.43	230 ± 27.3	262 ± 82.3
	40	2.86 ± 0.65	0.85 ± 0.06	2.19 ± 0.15	32.1 ± 11.4	609 ± 63.5	877 ± 157
	100	3.12 ± 0.30	1.03 ± 0.14	2.53 ± 0.42	35.3 ± 5.74	504 ± 26.9	849 ± 95.5
	50	2.62 ± 0.19	1.17 ± 0.10	1.93 ± 0.22	47.7 ± 3.62	716 ± 47.8	1051 ± 119
	40	2.57 ± 0.19	1.12 ± 0.16	1.72 ± 0.28	48.4 ± 5.09	868 ± 65.4	1221 ± 240

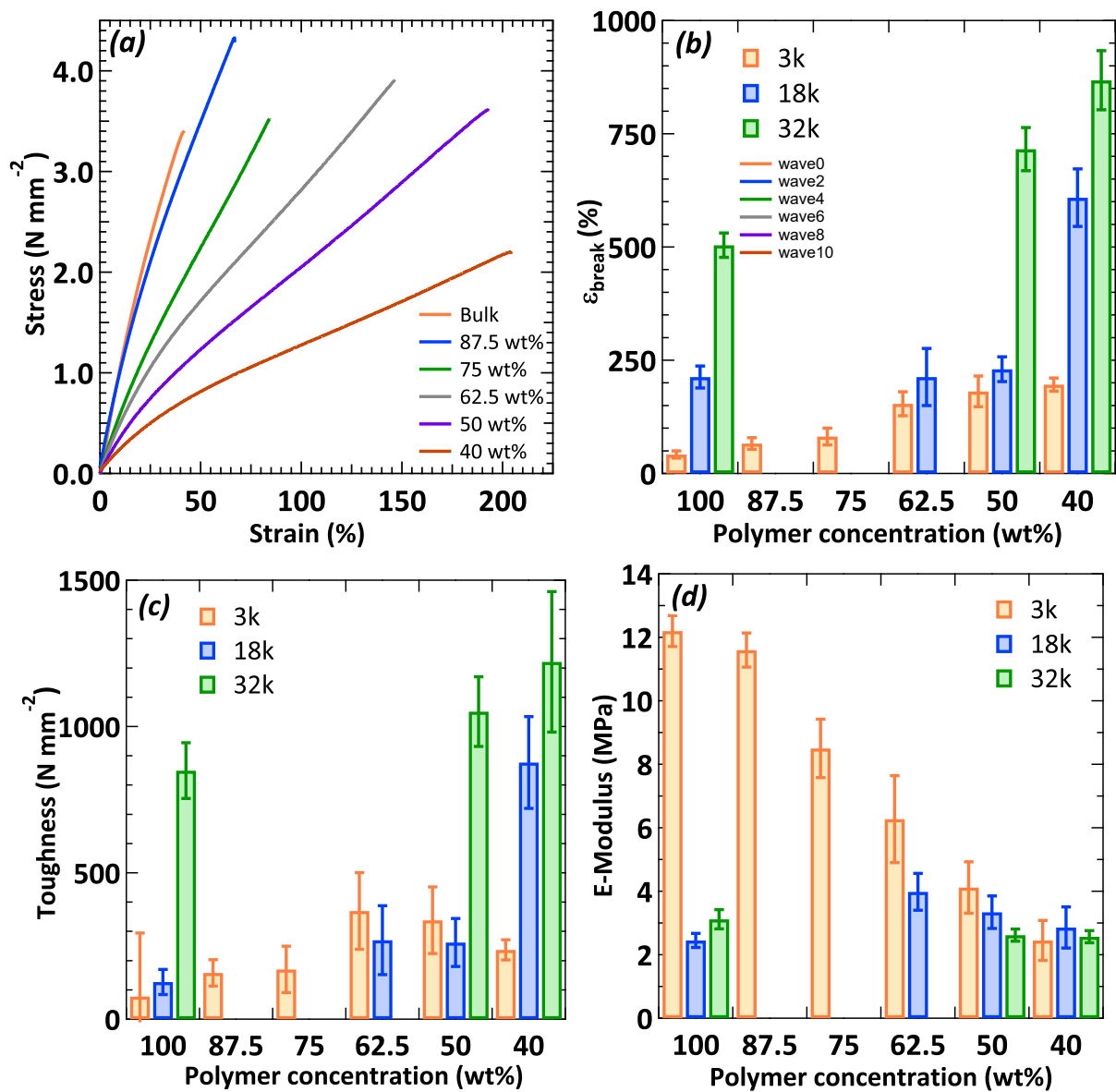


Figure 3: Overview of the tensile testing results of the extracted and dried *PTMC* networks. (a) Typical stress-strain curves of solvent-cast networks prepared from *PTMC* 3k macromer with different polymer contents. Trends in the change of (b) the strain at break, (c) the toughness, and (d) of the Young's *E* modulus as a result of changing the polymer content.

the crosslink densities  $\nu_C$  were calculated using Equation 2 according to the *affine* and *phantom* models. The computed values for all samples are also listed in Table 3.

Subsequently,  $I_{nDQ}$  signals normalized according to Equation 3 were obtained by  $DQ$   $^1H$  NMR. Figure 4a plots the obtained  $I_{nDQ}$  signals for *Bulk PTMC 3k*, *18k* and *32k* samples. It is observed that the  $I_{nDQ}$  signal becomes steeper when the macromer mass increases, which can be linked to an increase in crosslink density  $\nu_C$ . These signals were then fitted by Equation 5 in order to obtain the  $D_{res}$  value for each sample, which are listed in Table 3. Figure 4b plots the same  $I_{nDQ}$  signals as a function of the normalized  $DQ$  time  $D_{res}\tau_{DQ}$ . It is seen that all  $I_{nDQ}$  signals superpose, which would mean that the chemical and physical network structure for the three *Bulk PTMC* networks is the same. This has been previously observed for such materials[66].

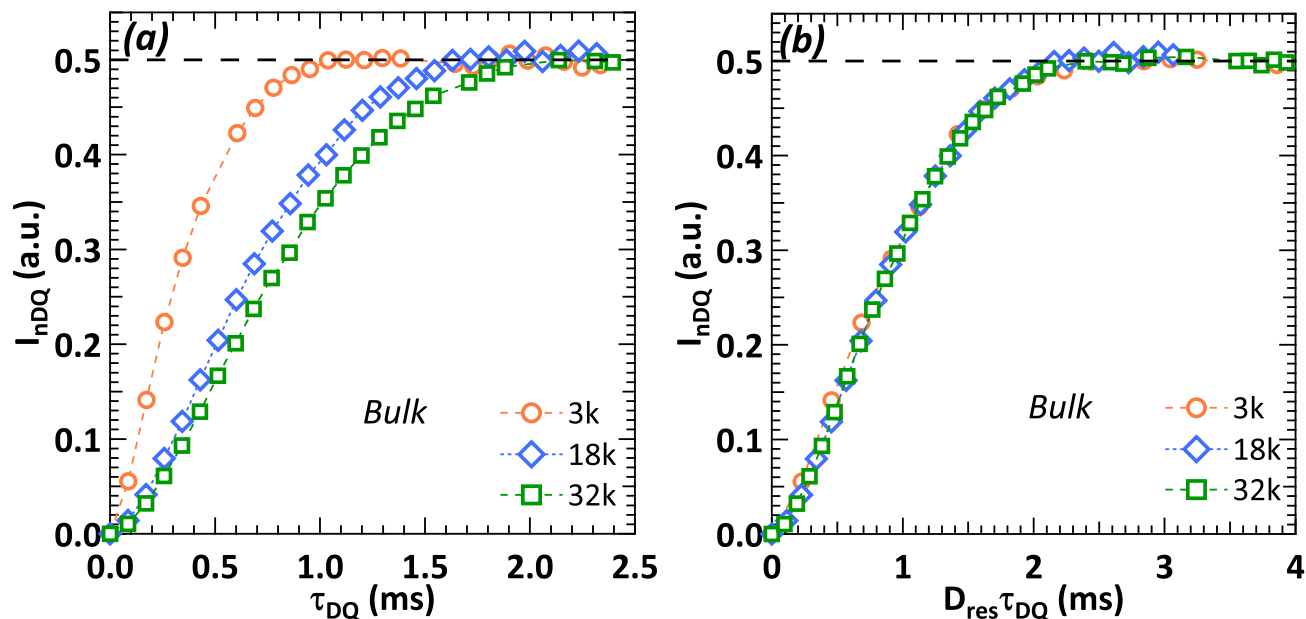


Figure 4: (a)  $I_{nDQ}$  values as a function of  $\tau_{DQ}$  for *Bulk PTMC* networks obtained by  $DQ$   $^1H$  NMR at  $T_\alpha + 90$  °C. (b) Same plot with a normalization of  $\tau_{DQ}$  by  $D_{res}$ .

The same approach was undertaken for solvent-cast *PTMC* networks. Figure 5 shows the  $I_{nDQ}$  signals obtained for the *3k*(a), *18k*(c), and *32k*(e), solvent-cast *PTMC* networks with varying polymer concentration. For each macromer molecular weight (*i.e.* *3k*, *18k*, and *32k*) it is systematically seen that when the polymer content increases, the  $I_{nDQ}$  signal becomes steeper. This is expected as propylene carbonate would act as a spacer between macromer chains, hindering the formation of denser crosslink networks. Thus, it can be concluded that the networks crosslink density strongly depends on the polymer content in the reactive medium.



Table 3: Physico-chemical, DSC, DMA, and  $^1H$  DQ NMR results obtained for the studied *PTMC* networks.

Polymer <i>PTMC</i> (wt%)	Physico-chemical		DSC		DMA			MQ $^1H$ NMR		
	$q$	$v_{C-chem}$ (mol/g) $\times 10^4$	$T_g$ ( $^{\circ}C$ )	$T_{\alpha}$ ( $^{\circ}C$ )	$E' @$ $T_{\alpha} + 90^{\circ}C$ (MPa)	$v_{C-DMA}$ affine phantom (mol/g) $\times 10^4$	$D_{res}/2\pi$ (Hz)	$w_{DEF}$ (%)	$n$	
100	$4.36 \pm 0.55$		-5.1	-11.7	9.1	10.32	30.95	359.7	4.3	1.46
87.5	$6.33 \pm 0.50$		-3.2	-12.5	7.3	8.35	25.06	335.6	1.6	1.52
75	$5.53 \pm 0.08$	8.82	-4.2	-12.9	4.8	5.49	16.46	294.7	3.4	1.58
62.5	$5.53 \pm 0.03$		-4.1	-12.4	4.0	4.56	13.67	262.3	5.3	1.56
50	$7.30 \pm 0.57$		-5.5	-12.4	2.6	2.95	8.85	236.9	5.2	1.59
40	$8.05 \pm 0.33$		-4.9	-13.2	1.9	2.21	6.63	223.7	1.2	1.80
100	$8.71 \pm 0.20$		-11.8	-20.4	4.8	5.62	16.85	210.5	1.2	1.70
62.5	$9.72 \pm 0.12$	1.80	-11.9	-22.5	4.4	5.19	15.56	200.3	2.5	1.79
50	$12.9 \pm 0.57$		-11.9	-21.8	2.8	3.30	9.91	191.2	4.6	1.76
40	$15.2 \pm 0.44$		-12.0	-22.1	1.0	1.21	3.63	160.8	6.0	1.72
100	$10.3 \pm 0.51$		-12.4	-14.8	4.4	5.09	15.28	178.5	13.8	1.70
50	$15.3 \pm 1.45$	1.04	-13.7	-20.7	2.8	3.23	9.68	169.7	4.9	1.78
40	$22.6 \pm 0.14$		-13.1	-21.7	2.3	2.66	7.97	161.8	3.9	1.77

Then,  $I_{nDQ}$  signals were fitted by Equation 5, yielding the numerical value of  $D_{res}$  for each solvent-cast sample, which are summarized in Table 3. Figures 5b, 5d, and 5f represent these  $I_{nDQ}$  signals as a function of the normalized time  $D_{res}\tau_{DQ}$ . Once more, for a given macromer molar weight, all  $I_{nDQ}$  signals superpose with each other with this normalization, implying that although the crosslink density  $\nu_C$  depends on polymer concentration during *PTMC* crosslinking, the chemical and physical morphology of the solvent-cast *PTMC* networks remains the same regardless of macromer molecular weight and initial polymer concentration.

Furthermore, for all bulk and solvent-cast *PTMC* networks, the  $D_{res}$  values obtained by *DQ*  $^1H$  NMR were plotted as a function of the crosslink density  $\nu_C$  calculated by the phantom model from DMA data as shown in Figure 6. (A similar plot with  $\nu_C$  calculated from the affine model is shown in Figure SI.5 - *Support Information*.)

Figure 6 shows that for each macromer molecular weight, a systematic linear relationship between  $D_{res}$  and  $\nu_C$  is obtained with a non-zero intercept for  $\nu_C = 0$ . This linear relationship is expected from rubber elasticity theory[77, 78] as well as from the affine[74] and phantom[75] models. Analogous results have been observed in elastomers[62] as well as in *PTMC Bulk* for networks of various molecular weights[66]. From this plot, several phenomena and conclusions can be drawn.

First of all, the non-zero intercept corresponds to physical entanglements within the networks. As mentioned in[66], *DQ*  $^1H$  NMR is able to probe these entanglements as they relax in a time domain slower or similar to those of the experiment, contrary to DMA whose characteristic measuring time is longer, during which entangled chains have relaxed. Moreover, it is observed in Figure 6 that the non-zero intercept value for *PTMC* networks diminishes with macromer molar weight (*i.e.*  $D_{res-0-3k} = 191.1Hz$ ,  $D_{res-0-18k} = 150.9Hz$ , and  $D_{res-0-32k} = 147.0Hz$ ). This may be attributed to a higher probed crosslink density induced by shorter macromer chains. Furthermore the  $D_{res-0}$  values for 18k and 32k are close to those reported in our previous work[66] ( $D_{res-0} = 135.5Hz$ ) in which a *PTMC 40k* network had the largest molecular weight. These values are relatively high when compared to the  $D_{res}$  values for *Bulk PTMC*. This would mean that the presence of physical entanglements contributes significantly to network properties, especially for low polymer content and/or high macromer molecular weight. This is especially true for the *32k PTMC* network, where the variation of  $D_{res}$  with polymer concentration is small.

Moreover, the  $D_{res}$  *vs.*  $\nu_C$  slopes diminish with increasing macromer molecular weight. The

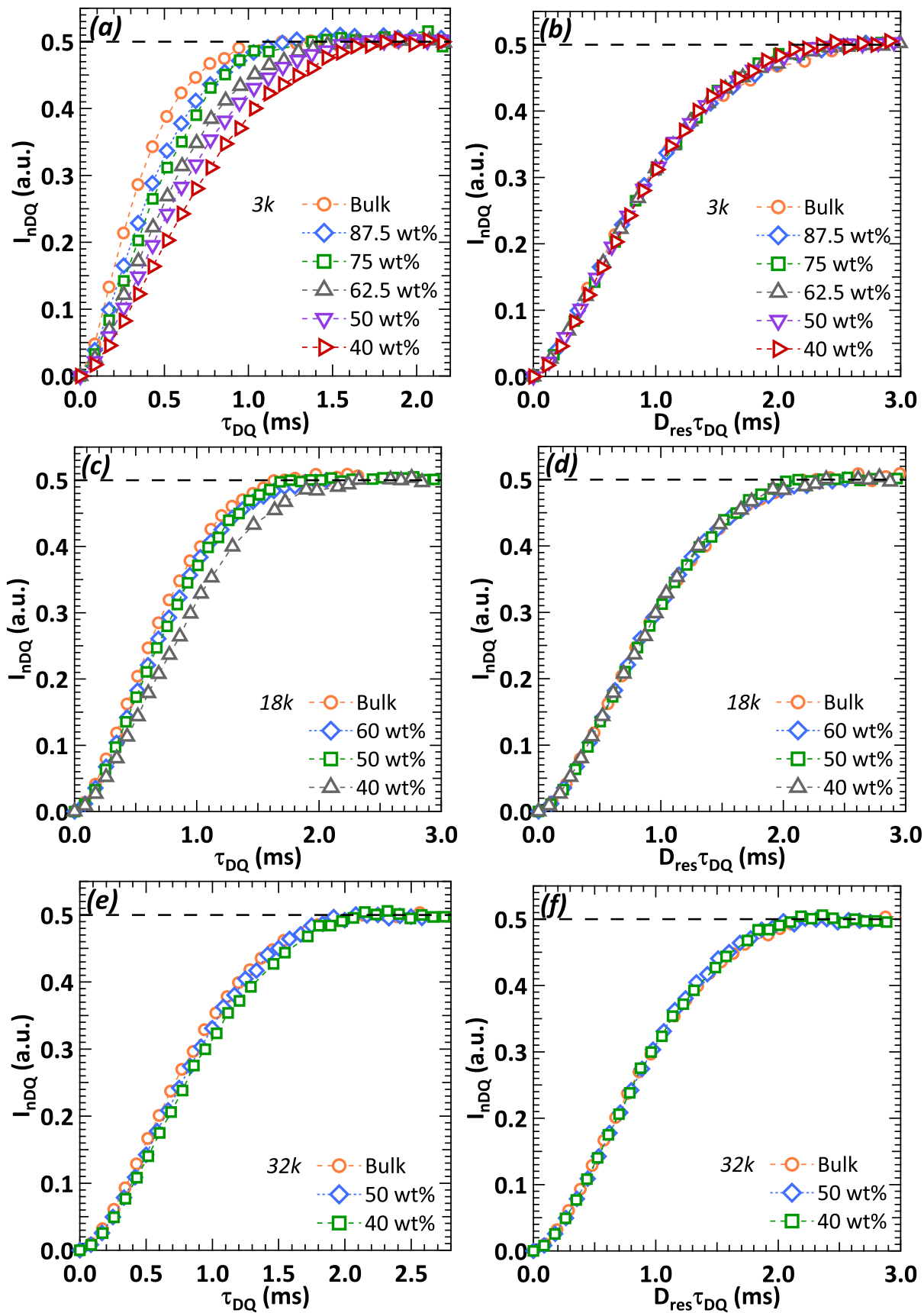


Figure 5:  $I_{nDQ}$  values as a function of  $\tau_{DQ}$  for (a) 3k (c) 18k, and (e) 32k solvent-cast *PTMC* networks obtained by  $DQ^1H$  NMR at  $T_\alpha + 90^\circ C$ . (b,d,f) Same plots with a normalization of  $\tau_{DQ}$  by  $D_{res}$ . The percentages correspond to polymer content in the reactive media.

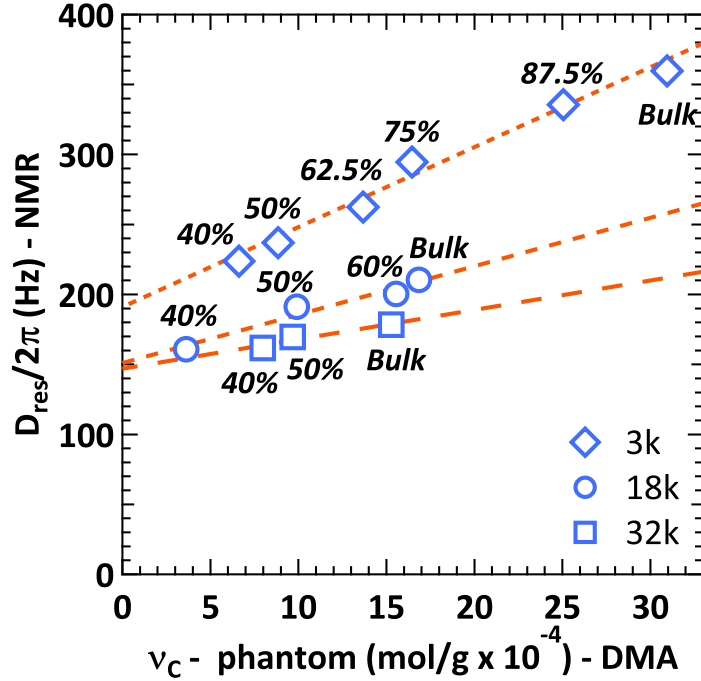


Figure 6:  $D_{res}$  values obtained by  $DQ$   $^1H$  NMR measurements as a function of DMA  $\nu_C$  crosslink densities calculated from the *phantom* model for all studied *Bulk* and *Solvent-cast PTMC* networks. Percentages correspond to polymer content in the reactive media. Dashed lines are linear fits and act as a guide for the eyes.

$DQ$   $^1H$  NMR data indicates that the influence of polymer concentration on the *chemical* crosslink density decreases as macromer molecular weight increases. This is expected because the crosslink density of the bulk networks decreases with increasing molecular weight, therefore solvent molecules would diffuse easily between two macromer reactive functionalities, limiting its steric influence.

Additionally, the  $DQ$   $^1H$  NMR analyses yielded the percentage of defects  $w_{DEF}$  and "network homogeneity factor"  $n$  (*i.e.* Equation 5). These values are compiled in Table 3. In the case of the  $n$  factor, it was systematically observed that for all macromer molecular weights, that when the polymer concentration decreases,  $n$  gets closer to the value of 2 (*i.e.* a "homogeneously crosslinked" network). This could be explained by the following hypothesis: as *propylene carbonate* acts as a spacer during the crosslinking reaction, *PTMC* macromer chains are less constrained and entangled within the reactive mixture. This would enhance the efficiency and likelihood of steric-oriented reactions between macromers, inducing a better-ordered network following photocrosslinking.

Furthermore, the evolution of the percentage of defects  $w_{DEF}$  with polymer concentration seems to depend on the macromer molecular weight. Indeed, each *PTMC* series of networks have a different evolution of  $w_{DEF}$  with polymer content, as can be seen from the values reported in

Table 3. In the case of *PTMC 3k* networks, no clear evolution of  $w_{DEF}$  is observed, with the values remaining close to each other. This could be explained by the fact that macromer chains are relatively short. Indeed, *PTMC* repeating unit molar mass is  $M_0=102$  g/mol, so the *3k* network possesses *ca.* 30 repeating units per chain (which is only *ca.* 10 repeating units per arm). Even though a better-ordered network is obtained, dangling or free chains are still obtained due to some steric hindrance as a result of the size of the macromer chain. Concerning the *PTMC 18k* networks, it is seen in Table 3 that  $w_{DEF}$  increases with decreasing polymer content. The *18k* network possesses *ca.* 150 repeating units per chain (*ca.* 50 repeating units per arm). It is probable that *propylene carbonate* induces some steric hindrance in the reactive medium, hampering crosslinking reactions, generating a higher amount of dangling and free chains. Lastly, for the *PTMC 32k* networks, Table 3 shows that  $w_{DEF}$  decreases with polymer concentration, contrary to the *PTMC 18k* networks. The *32k* network possesses *ca.* 300 repeating units per chain (*ca.* 100 repeating units per arm). In this case, it is possible that less steric hindrance is induced by the presence of *propylene carbonate*, allowing more crosslink reactions, thus yielding a lower amount of defects.

Finally, it would seem that the percentage of defects  $w_{DEF}$  has a prominent influence on *PTMC* macroscopic properties. As expected, material toughness and elongation at break depended mainly on the crosslink density as previously shown in Figure 3. However, in the case of the measured Young’s modulus  $E$ , its evolution with the *PTMC* structural morphology is more complex. It would also be expected that  $E$  would primarily depend on the crosslink density. Nonetheless, in the case of *PTMC 18k*,  $E$  increases when the polymer concentration decreases (*i.e.* when the crosslink density decreases). However, for this macromer molecular weight, the percentage of defects  $w_{DEF}$  increased when the polymer content decreased (*i.e.* Table 3). This would mean that for this given macromer molecular weight, *dangling and free chains seem to mechanically reinforce the materials*. In the case of *PTMC 3k* and *32k*,  $w_{DEF}$  remains constant or increases with polymer content, as does  $E$ . This indicates that dangling or free chains play a less important role in determining the mechanical moduli when compared to the influence of the polymer’s crosslink density.

## 5. Conclusion

This investigation has highlighted the advantages of combining a multiscale experimental approach to better understand the macroscopic mechanical properties of *PTMC* networks from their intrinsic structural morphology. By combining tensile testing, DMA analyses and  $^1H$  DQ NMR

measurements it was shown that *PTMC* crosslink density depended on the polymer content during the crosslinking reaction; the lower the polymer content, the lower the crosslink density value. This was systematically observed for the three studied macromer molar masses of *3k*, *18k* and *32k*. Moreover, it was observed by  $^1H$  DQ NMR that even though the samples had varying crosslink densities, their intrinsic chemical and physical network morphology remained the same. Furthermore, it was observed that *PTMC* becomes more ductile (*i.e.* a decrease in toughness and an increase in elongation at break) when the polymer content decreases, *ergo* the crosslink density decreases. More surprisingly, it was observed that for the *PTMC 18k*, an increase in Young's modulus  $E$  was observed when the polymer content decreased (*i.e.* decrease of crosslink density). Indeed, for the *PTMC 3k* and *32k*,  $E$  diminished when the polymer content decreased, which is expected as fewer chemical or physical crosslinks are present in the material. In the case of *PTMC 18k*, the rise of  $E$  might be due to the presence of a higher amount of dangling and free chains acting as reinforcement. A higher percentage of dangling and free chains is attributed to *propylene carbonate* hampering crosslinking reactions due to steric hindrance. With this study, the understanding of how the chemical and structural tailoring of *PTMC* networks influences macroscopic properties has been deepened. This proven approach will be applied to study the network structure and functional macroscopic properties of heterogeneous *PTMC* materials targeted for biocompatible applications.

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## 8. Supplementary Information

Supplementary Information available: Equation SI.1; DQ  $^1H$  NMR  $I_{ref}$ ,  $I_{DQ}$ ,  $I_{ref} - I_{DQ}$ , and  $I_{def}$  signals obtained for *3k Bulk* PTMC at  $T_\alpha + 90^\circ C$ . The contribution from defects  $I_{def}$  was fitted from the  $I_{ref} - I_{DQ}$  signal and its fraction  $w_{DEF}$  was calculated by extrapolating the fit to  $\tau_{DQ} = 0$ ;  $I_{nDQ}$   $^1H$  NMR signals obtained for *3k Bulk PTMC* at various temperatures and normalized according to Equation SI.1;  $I_{nDQ}$   $^1H$  NMR experimental signal obtained for *18k Bulk* PTMC at  $T_\alpha + 90^\circ C$  fitted with Equation 5; Typical  $^1H$  NMR spectra for a *PTMC* oligomer and macromer;  $D_{res}$  values obtained by *DQ*  $^1H$  NMR measurements as a function of DMA  $v_C$  crosslink densities calculated from the *affine* model for all studied *Bulk* and *Solvent-cast PTMC* networks. Percentages correspond to macromer *wt%*. Dashed lines are linear fits and act as a guide for the eyes.

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