A concept map of death-related anxieties in patients with advanced cancer

Sigrun Vehling, Carmine Malfitano, Joanna Shnall, Sarah Watt, Tania Panday, Aubrey Chiu, Anne Rydall, Camilla Zimmermann, Sarah Hales, Gary Rodin, and Christopher Lo

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Abstract

Objectives: Fear of death and dying is common in patients with advanced cancer, but can be difficult to address in clinical conversations. We aimed to show that the experience of death anxiety may be deconstructed into a network of specific concerns, and provide a map of their interconnections to aid clinical exploration.

Methods: We studied a sample of 382 patients with advanced cancer recruited from outpatient clinics at the Princess Margaret Cancer Centre, Toronto, Canada. Patients completed the 15-item Death and Dying Distress Scale (DADDS). We used item ratings to estimate a regularized partial correlation network of death and dying-related concerns. We calculated node closeness-centrality, clustering, and global network characteristics.

Results: Death-related anxieties were highly frequent, each associated with at least moderate distress in 22% to 55% of patients. Distress about “Running out of time” was a central concern in the network. The network was organized into two areas: one of more practical fears concerning the process of dying, and another of more psychosocial or existential concerns including relational problems, uncertainty about the future and missed opportunities. Both areas were yet closely connected via important bridges, e.g., fear of suffering and a prolonged death was linked to fear of burdening others.

Conclusions: Individuals with advanced cancer may have multiple interconnected death-related concerns, forming patterns that vary with individual priorities and fears. The bridging links between more practical and more psychosocial concerns emphasize that the alleviation of death anxiety may require interventions that combine symptom management, advance care planning, and psychological treatment approaches.

Key words: Death anxiety, death distress, advanced cancer, psycho-oncology, network analysis, end of life
Introduction

Extended survival and the growing emphasis on palliative care have highlighted the importance of conversations about death and dying in advanced incurable cancer. Death-related concerns and fears are inherent in discussions about advance care planning and preferences for prognostic disclosure, and may also trigger the search for meaning in face of impending mortality [1, 2]. For patients nearing the end of life, the possibility to discuss death and dying in a safe place can facilitate adaptation and prevent severe psychological and existential distress [3]. However, the evaluation of clinical conversations and supportive interventions that aim to alleviate death anxiety are limited by our understanding of this concept [4].

In advanced cancer, the understanding of death anxiety has been limited by the view that talking about this difficult topic may be demoralizing for patients. While this perspective has been related to modern medicine’s emphasis on cure [5], it is also evident in traditional theories of death anxiety. These have often assumed that death awareness is an intolerable human experience and that self-distancing and denial of death are necessary to maintain psychological health [6, 7]. However, many patients with advanced cancer prefer open discussions about death-related concerns and timely preparation for the end of life [8, 9]. This observation suggests that exploration of patients’ death-related concerns and fears is essential to clinical interventions in this population [10–13]. Empirical evidence that may guide such exploration remains scarce [1].

Although death anxiety has been viewed as an overarching construct, it may also be deconstructed into particular fears and concerns about death and dying. Accordingly, the Death and Dying Distress Scale (DADDS [14]) captures concerns about the loss of time and missed opportunities in life, uncertainty about the future, fears about the process of dying, and the impact of one’s death on others. Although factor analyses of the DADDS have found a single-factor structure [14, 15], qualitative studies and conceptual reviews point to a complex system of interacting thoughts, fears, and worries associated with death and dying in advanced cancer [4, 16–18]. Also, while studies have suggested a
distinction of the fear of dying from the fear of death itself, the association between these domains is not well understood [17, 19, 20].

The present study is focused on particular death-related fears and concerns and their links among patients with advanced cancer. We investigated the network structure of death-related concerns by a detailed analysis of the connections between them. This network approach thus “unpacks” the relatively abstract construct of death anxiety by closely examining the interrelation of its constituent elements. Related existential distress constructs such as demoralization, loss of dignity, and end-of-life preparation have been studied in advanced life-threatening disease [21–23]. However, relationships between existential concerns on the item level have rarely been examined [24].

We analyzed the network structure of death-related anxieties in patients with advanced cancer and we aimed to: (a) identify the presence of central concerns that connect together a large number of other concerns; (b) characterize the overall configuration of the network by describing the areas formed by closely interconnected concerns; and (c) interpret the role of bridge concerns that link those areas. The product of this analysis is a map of death-related concerns and their interconnections, which may be helpful in guiding the exploration and treatment of death anxiety in patients with advanced cancer near the end of life.

Methods

Participants and procedures

Patients with advanced cancer were recruited from outpatient oncology clinics at the Princess Margaret Cancer Centre, University Health Network (UHN) in Toronto, Canada. Advanced cancer was defined by diagnosis of a stage IV solid tumor or stage III lung or ovarian tumor, or any stage pancreatic cancer (due to aggressiveness of this disease), all with an expected survival of 12-18 months. Patients were invited to participate in a randomized controlled trial of a psychotherapeutic
intervention called CALM (Managing Cancer And Living Meaningfully) to alleviate distress (for study protocol, see [25]). After providing written informed consent, participants completed baseline self-reported measures, including the assessment of death-related concerns or anxieties, which are the focus of the present study. Exclusion criteria were age younger than 18 years, severe cognitive or physical impairment, language barrier, and current receipt of other psychosocial interventions from the Centre’s Department of Supportive Care. Patients provided written informed consent and completed baseline measures before randomization. Three hundred and eighty-two out of 2601 eligible patients with advanced cancer (15%) completed baseline death anxiety questionnaires. This study was approved by the UHN Research Ethics Board.

Measures

Demographic data were collected by a standardized questionnaire. Disease-related characteristics were obtained from medical charts. We used the 15-item Death and Dying Distress Scale (DADDS) [14, 15] to assess death anxiety. The scale was specifically developed to capture concerns related to death and dying in patients with advanced cancer. It encompasses psychosocial and existential concerns about missed opportunities and the loss of time, past regrets and uncertainty in the future, and the impact or burden of death on others. Further, it assesses more practical concerns about the process of dying, including fears about dying alone or with suffering, or worries about its timing. Items are answered on a 6-point Likert scale from 0 (no distress) to 5 (extreme distress). Total scores may range from 0 to 75 and scores ≥45 refer to the presence of at least moderate death anxiety. By inquiring about the specific death-related concerns of most relevance to patients with advanced cancer, the DADDS is distinct from death anxiety scales developed for use in non-advanced cancer populations for whom mortality is not imminent [15]. It focuses on the assessment of disturbing thoughts that require alleviation, and does not assess reaction to symbolic reminders of death nor death attitudes, for which the direction of desired clinical change is less clear (e.g., belief in an
afterlife, death acceptance). This reflects a perspective on death anxiety as becoming more focused on the fear of death among patients with advanced disease, given the imminence of mortality and the many foreseeable challenges associated with dying [6].

**Statistical analysis**

We used R version 3.3.0 [26] and package qgraph, version 1.3.3 [27]. Descriptive statistics including means, standard deviations, and frequencies were calculated for the total scale as well as individual DADDS items. Pearson correlations were calculated between each pair of items given non-skewed distributions. Missing values occurred in only 2 of 5685 responses and these were mean-imputed.

**Box 1: Introduction to the network approach** We applied a network analysis perspective to death anxiety [28]. A network analysis examines the intercorrelations between concerns or symptoms in a graphical form. In a network graph, stronger links due to higher correlations between concerns are indicated by shorter and thicker lines (called “edges”) connecting them. Network structures thus graphically illustrate which concerns tend to co-occur. They focus on the shape and interconnectivity of symptoms – unlike factor analysis in which is closely related symptoms are grouped to define discrete syndromes. Network structures are suggestive of how effective alleviation of a particular symptom (called a “node”) may require addressing closely related symptoms in the network. In that regard, bridge symptoms that link otherwise separate areas of a network are an important feature and can indicate how intervention effects may transfer across network domains.
We conducted a network analysis (see Box 1 for an introduction) of the 15 DADDS items to estimate and visualize the interrelationships between these concerns. A network analysis begins by calculating the partial correlations between each pair of items, that is, pairwise associations that have been adjusted for the effects of the other items in the network. This partial correlation matrix is visualized in a graph where items are displayed as nodes in the network and the partial correlations become the edges connecting these nodes. Thicker edges indicate higher partial correlations between nodes. Highly correlated nodes are placed closer together through the force-directed Fruchterman-Reingold algorithm implemented in qgraph [27]. The sensitivity of the partial correlation matrix was optimized using a graphical lasso (least absolute shrinkage and selection operator) regularization technique. The glasso shrinks very small partial correlations to zero to maintain a sparse partial correlation matrix. The tuning parameter $\alpha$ returning the optimal degree of shrinkage is determined through an algorithm that minimizes the extended Bayesian information Criterion (EBIC). The hypertuning parameter $\gamma$, defining the degree of simplicity preferred by the EBIC, was set to the default medium value of 0.25.

We further calculated descriptive indices to characterize the network’s node-specific and global properties. At the local level, a node’s closeness-centrality describes its total distance to all other nodes in the network. Higher values indicate greater centrality of the node within the network. A node’s closeness is calculated by the inverse sum of its distances to all other nodes. The clustering coefficient describes the number of nonzero connections relative to the maximum number of possible connections between the neighbors of each node. Clustering can be interpreted as a measure of a node’s redundancy. Values may range from 0 (no clustering) to 1 (high clustering). Barrat’s coefficient was used to adjust for both edge width and number when calculating clustering coefficients. At the global level, the average shortest path length (ASPL) measures the average shortest number of edges between each pair of nodes. For example, a network with ASPL=2 means that on average, the shortest link between each pair of nodes is two edges. If ASPL=1, the network is highly concentrated and, on average, nodes are linked by only one edge. We further calculated Opsahl’s weighted version of the ASPL, which adjusts for edge width. The transitivity of the network
is a measure of its global clustering. It describes the average probability that two nodes, which are linked to the same neighbor, are neighbors themselves. Values may range between 0 and 1. The smallworldness index (SWI) relates a network’s transitivity to its unweighted ASPL. A smallworld network is characterized by high transitivity (i.e., high clustering) and low ASPL. Smallworldness is indicated by an SWI ≥3.

Results

Patient characteristics and frequency of death anxiety

382 patients completed baseline death anxiety questionnaires. Table 1 presents demographic and disease-related characteristics of this sample. Figure 1 shows the frequency of patients reporting each death-related concern (cut-off ≥ 3 indicating at least moderate distress). The most frequent concerns were: the impact of one’s death on loved ones (65.4%); that one’s death and dying may be prolonged or drawn out (55.3%); and that it may happen with a lot of pain and suffering (54.8%). Of note, even the least frequent concern (fear of dying alone) was reported by almost a quarter of participants. Further descriptive statistics of concerns are reported in the online supplement, Table 2. Patients reported having an average of 6.5 (SD=4.7) out of 15 possible death-related concerns. The mean total death anxiety score was 31.6 (SD=18.0). Moderate death anxiety was present in 27.4% of the patients (cut-off ≥ 45).

Table 1: Demographic and disease related sample characteristics (N=382).
<table>
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<th>Education</th>
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<td>52.9</td>
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<td></td>
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<td></td>
<td>Lung</td>
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<td></td>
<td>Breast</td>
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<td>8.6</td>
</tr>
<tr>
<td></td>
<td>Other</td>
<td>42</td>
<td>11.0</td>
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<tr>
<td><strong>No. of physical symptoms, mean (SD)</strong></td>
<td>12.31 (4.9)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

*Possible range: 0-28, according to the Memorial Symptom Assessment Scale [29]*

Please insert **Figure 1** about here

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**Network of death-related anxieties**

Figure 2 presents the regularized partial correlation network of death-related concerns. The network has at least four major features in terms of structural configuration, concern areas, and bridge connections that facilitate “traveling” across areas in the map. The first important structural feature is that “Running out of time” (item 7) is located at the center of the network. Centrality is a different indication of item importance than frequency. A central item is one that retains the most relationship to all other concerns, after adjusting for all associations. Although it was not the most frequent concern, this node had the highest centrality value meaning that it was the most conceptually connected concern in the network (see Figure 3). The remaining concerns formed a circular chain around the “Running out of time” node. This circular arrangement is a reflection of the distribution of centrality values shown in Figure 3, i.e., less central concerns are placed more distant from the center.
The second feature in Figure 2 describes a possible organization of the total network into two halves or areas, which coincides with differences in item content. The left area included all five items inquiring about practical fears about the dying process (grey-colored items 11 to 15). The right area included all, in a narrower sense, psychosocial and existential concerns related to the anticipation of death (items 1 to 10). The third feature of the network then showed how these concern areas of the map were connected via important bridges. Three main bridges (network edges) connected the left more “practical” and right more “psychosocial/existential” area (item 13 to 2, 15 to 7, and 12 to 8). The nodes at the ends of these edges can be viewed as key or bridge nodes that are travelled through to move across the network areas. This means that, for example, starting from fear of a prolonged death (item 12), exploration toward the right psychosocial area is possible via the closely related concern of being a burden to others (item 8).

A fourth noteworthy feature of the network is evident from examining the outer ring of death-related concerns surrounding “Running out of time”. It describes a further possible organization of the items of this ring into four areas or quadrants. Starting at the upper right, the first quadrant was composed of items associated with past regrets and unfinished business, i.e. items 2, 1, 3, and 6. The second quadrant was composed of items relating to the future and uncertainties near the end of life, i.e. items 5, 4, and 10. The third quadrant referred to items describing a prolonged and painful death and
dependency needs in this circumstance, i.e. items 9, 8, 12, and 14. The fourth quadrant included items describing aspects of an unprepared death, specifically one that will happen soon, unexpectedly, and in the absence of family and friends, i.e. items 15, 11, and 13. As shown in Figure 3, clustering coefficients were high for concerns 3 and 14, indicating that these were most redundant in the network given closely interconnected neighbors.

Global network characteristics indicated a “mediumworld” structure of death anxiety. This means that concerns were neither extremely clustered (i.e., redundant) nor extremely distant (i.e., conceptually unrelated). The total number of nonzero edges in the network was 64 (61% of possible edges). The average shortest path length between two nodes in the network was 1.4. That is, approximately half of node pairs were directly connected via one edge, while the other half were connected via two edges. The average weighted shortest path length was 12.5, which was approximately equal to the weighted distance between concerns 7 and 12. The transitivity (global clustering coefficient) was 0.60, indicating that 60% of nodes that shared a neighbor were themselves neighbors. The smallworldness index was 1.05; the network did hence not show smallworld properties.

**Discussion**

Although death anxiety has been viewed as an overarching concept, it may be deconstructed into a network of more specific fears or concerns, which in turn may be complexly interrelated. Using a network approach, this study graphically depicted the interconnections between 15 death-related concerns in a large sample of patients with advanced cancer. By visualizing the associations between concerns, the network may serve as a guide for clinicians and researchers to explore the structure of death-related anxieties in patients with advanced disease [30]. The interconnections between nodes suggest potential pathways along which the discussion of death-related fears may proceed. Although such conversations may begin anywhere, the centrality of the concern that time is running out suggests that starting the conversation here might allow for maximum flexibility in how the
discussion unfolds. The “mediumworldness” of the network and similar centrality of remaining concerns indicate that each issue is non-redundant and worthy of exploration.

Although physical and practical concerns about dying clustered somewhat separately from psychosocial concerns, these network areas were also connected by clear pathways or bridges, suggesting that treatments to alleviate death anxiety may require addressing physical and psychosocial issues jointly. Although psychotherapeutic treatments [31] and palliative care interventions [32] have been developed to alleviate suffering and promote preparation for the end of life, their integration is less common. It is possible that integrated approaches will have more powerful effects than delivery of either alone.

The network showed that fears of a prolonged death (item 12) tend to be associated with concerns about how this would burden others (item 8) and its negative impact on loved ones (item 9). This result is consistent with earlier studies documenting an association between worries about the future and the experience or anticipation of stress imposed on others, which relates to perceived burdensomeness [33, 34]. Another important set of linkages emerged between the fear of dying alone (item 13), having not said important things to others (item 2), and regret about things that were not done (item 1). This suggests that a sense of unfinished business for individuals may often involve unresolved or problematic relationships. Interventions that aim to support patients in achieving a sense of social connectedness and life completion may require understanding how individuals view their social legacy and the possibilities for reconnection or letting go of the things that cannot be changed in life [12, 13, 35, 36]. Discussion of the practical issues of end-of-life preparation may contribute to interventions that focus to help patients in defining what is important to them in the time that is left [3].

The presence of a single central node is a unique feature of the present network [37, 38]. The experience of time running out may be central to the psychology of incurable disease, which may fundamentally be a problem of time [39]. Nonetheless, patients with advanced cancer may present
with different individual configurations or profiles of death-related concerns. For some, a foreshortened future can disrupt the sense of meaning attached to previously held values and goals in life, causing existential regret [19, 39, 40]. Others may be more overwhelmed by the uncertainty and losses that lie ahead, including concerns about how others will respond to unfolding events[18], as suggested by the links between being concerned about “my own death and dying”, “not knowing about what happens at the end of life”, and the “impact on loved ones” in the network.

We note the following limitations and future research directions. The present network shows how concerns are linked on average in the sample, although a range of individual death anxiety profiles may be expected. This overview of conceptual associations between concerns may provide a fruitful departure for further investigations that focus on how individuals may differ in their patterns of death anxiety. Network structures of death-related concerns may further be compared across subgroups with high vs. low physical symptom burden, time since diagnosis, and desire for hastened death. Further, time series will enable insight into single-person networks. In addition, networks including a broader range of psychological and existential distress symptoms could further explore the structure of distress in advanced cancer. We are aware that the 15-item-DADDS may not cover all dimensions of death anxiety in advanced cancer. However, its shortness and conceptual parsimony avoid overlap with other distress concepts. While this applies to all studies using self-report measures, semantic overlap between items is a source of correlation that must be taken into account in interpretation, in addition to the association among concepts. Lastly, our results are most representative of outpatients with advanced cancer with an expected survival of 12-18 months interested in receiving psychotherapeutic intervention. The structure of concerns may change with disease progression and differ in patients closer to death.

In conclusion, our study offers an overview of the richness and complexity of associations involved in the structure of death-related distress in individuals with advanced cancer by a detailed investigation of the links between specific fears and worries. The map of concerns and their connectivity that has
been generated, may help to guide clinical conversations and the development of supportive interventions to alleviate these symptoms.

Acknowledgements

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Conflicts of interest

The authors declare no conflict of interest.

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Contributors

Conception and design: SV, GR, CL. Data analysis: SV, CL. Interpretation and manuscript writing: SV, CM, JS, SW, TP, AC, AR, CZ, SH, GR, CL. Final approval: SV, CM, JS, SW, TP, AC, AR, CZ, SH, GR, CL. Guarantor: SV, GR, CL.

Ethics approval
University Health Network Research Ethics Board.
References


**Figure Captions**

**Figure 1:** Frequency of death-related concerns (N=382). Moderate distress is defined by item ratings \( \geq 3 \). The question stem for items 1 to 10 was: “Over the past two weeks, how distressed did you feel about…”. The question stem for items 11 to 15 was: “Over the past two weeks, how distressed did you feel that your own death and dying may…”.

**Figure 2:** Network of death-related concerns. Nodes represent the death-related items; edges represent partial correlations between items. Thicker edges mean higher partial correlations. The question stem for items 1 to 10 (white) was: “Over the past two weeks, how distressed did you feel about…”. The question stem for items 11 to 15 (grey) was: “Over the past two weeks, how distressed did you feel that your own death and dying may…”. Partial correlations < 0.05 are not shown. All edge weights are listed in the online supplement. See Figure 1 for complete items.

**Figure 3:** Closeness-centrality and clustering coefficients for death-related concerns. Closeness-centrality indicates a node’s connectivity within the network. It describes the node’s total distance to all other nodes in the network. Higher values indicate greater centrality. The clustering coefficient indicates a node’s redundancy relative to other nodes. It describes the percentage of nonzero connections that exist between its neighbor nodes. Higher values indicate greater clustering.
9. The impact of my death on my loved ones 65.4%
12. Be prolonged or drawn out 55.3%
14. Happen with a lot of pain or suffering 54.8%
8. Being a burden to others 51.2%
10. My own death and dying 49.6%
5. Not having a future 49.1%
7. Running out of time 47.8%
4. Not knowing what happens near the end of life 46.4%
1. Not having done all the things I wanted to do 45.1%
6. The missed opportunities in my life 40.9%
3. Not having achieved my life goals and ambitions 36.7%
15. Happen very soon 34.3%
2. Not having said all I wanted to say to the people I care about 27.2%
11. Happen suddenly or unexpectedly 25.3%
13. Happen when I am alone 22.4%
### Table 2: Descriptive statistics for death-related concerns

<table>
<thead>
<tr>
<th>Death-related concerns</th>
<th>Mean</th>
<th>SD</th>
<th>Skew</th>
<th>Kurtosis</th>
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<tr>
<td>9. The impact of my death on my loved ones.</td>
<td>3.1</td>
<td>1.7</td>
<td>-0.5</td>
<td>-1.0</td>
</tr>
<tr>
<td>14. Happen with a lot of pain or suffering.</td>
<td>2.5</td>
<td>1.8</td>
<td>-0.1</td>
<td>-1.4</td>
</tr>
<tr>
<td>8. Being a burden to others.</td>
<td>2.4</td>
<td>1.7</td>
<td>-0.0</td>
<td>-1.2</td>
</tr>
<tr>
<td>10. My own death and dying.</td>
<td>2.4</td>
<td>1.7</td>
<td>0.0</td>
<td>-1.2</td>
</tr>
<tr>
<td>12. Be prolonged or drawn out.</td>
<td>2.4</td>
<td>1.7</td>
<td>-0.1</td>
<td>-1.2</td>
</tr>
<tr>
<td>5. Not having a future.</td>
<td>2.3</td>
<td>1.7</td>
<td>0.0</td>
<td>-1.3</td>
</tr>
<tr>
<td>7. Running out of time.</td>
<td>2.3</td>
<td>1.7</td>
<td>0.7</td>
<td>-1.2</td>
</tr>
<tr>
<td>1. Not having done all the things I wanted to do.</td>
<td>2.2</td>
<td>1.5</td>
<td>0.1</td>
<td>-0.9</td>
</tr>
<tr>
<td>4. Not knowing what happens near the end of life.</td>
<td>2.2</td>
<td>1.7</td>
<td>0.1</td>
<td>-1.2</td>
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<tr>
<td>6. The missed opportunities in my life.</td>
<td>2.0</td>
<td>1.7</td>
<td>0.3</td>
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<tr>
<td>3. Not having achieved my life goals and ambitions.</td>
<td>1.9</td>
<td>1.6</td>
<td>0.4</td>
<td>-1.0</td>
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<tr>
<td>15. Happen very soon.</td>
<td>1.8</td>
<td>1.7</td>
<td>0.5</td>
<td>-1.0</td>
</tr>
<tr>
<td>2. Not having said all I wanted to say to the people I care about.</td>
<td>1.6</td>
<td>1.4</td>
<td>0.6</td>
<td>-0.6</td>
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<tr>
<td>11. Happen suddenly or unexpectedly.</td>
<td>1.4</td>
<td>1.5</td>
<td>0.8</td>
<td>-0.4</td>
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<tr>
<td>13. Happen when I am alone.</td>
<td>1.2</td>
<td>1.5</td>
<td>1.1</td>
<td>-0.1</td>
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</table>

*Total mean score of death anxiety*  
31.6  

*Number of death-related concerns*  
6.5

*aNumber of concerns rated ≥ 3, indicating at least moderate distress*
Table 3: Correlations among death-related concerns

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Note: All correlations were significant (p<.001).

1. Not having done all the things I wanted to do
2. Not having said all I wanted to say to the people I care about
3. Not having achieved my life goals and ambitions
4. Not knowing what happens near the end of life
5. Not having a future
6. The missed opportunities in my life
7. Running out of time
8. Being a burden to others
9. The impact of my death on my loved ones
10. My own death and dying
11. Happen suddenly or unexpectedly
12. Be prolonged or drawn out
13. Happen when I am alone
14. Happen with a lot of pain or suffering
15. Happen very soon
Table 4: Regularized partial correlations among death-related concerns (edge weights of the network shown in Figure 1)

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Note: Greater edge weights result in thicker edges in the network shown in Figure 1.

1. Not having done all the things I wanted to do
2. Not having said all I wanted to say to the people I care about
3. Not having achieved my life goals and ambitions
4. Not knowing what happens near the end of life
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