specific PCR products, confirmed by Southern blot analysis. Control (–) and an Hae + eight patients with SCLC (three samples from patient 3) including a positive (mitotic, small basophilic tumour cells.

Figure 1  Histological features of (a) Merkel cell carcinoma (MCC; H&E staining, ×400) and (b) small cell lung cancer (SCLC; H&E staining, ×400). Both tumours are composed of highly mitotic, small basophilic tumour cells. (c) Electropherogram of a 1.5% agarose gel for MCV138 in eight patients with SCLC (three samples from patient 3) including a positive (+), and negative control (–) and an Haelll size ladder (L). (d) Corresponding Southern blot. Samples 3b and 3c display specific PCR products, confirmed by Southern blot analysis.

Assessment of gender differences in health status with the Leicester Cough Questionnaire (LCQ)

Kelsall et al were surprised that there were no differences in Leicester Cough Questionnaire (LCQ)1 scores in men and women with chronic cough and suggest that this was because the LCQ did not capture gender-specific differences in health status.2 We disagree. We have previously reported significantly worse health status in a larger group of females with chronic cough (see table 1).3,4 Furthermore, Polley et al investigated the gender differences in health status in 147 patients with chronic cough using the LCQ and another health status measure (Cough specific quality of life questionnaire) and found a clinically significant impairment in health status in females with both questionnaires.5 So why did Kelsall and colleagues not detect gender differences in health status? The lack of a significant difference in day-time cough frequency between genders may have been one factor. Health status questionnaires tend to focus more on the adverse impact of cough during the day. Interestingly, female patients in our study had significantly longer duration of cough than males; this may have contributed to the gender differences (table 1). We have previously reported a poor to moderate relationship between cough symptoms, health status, cough reflex sensitivity and cough frequency.6 Cough frequency is just one of several factors likely to be important in determining the well-being of patients. This highlights the importance of using a combination of subjective and objective tools such as automated cough monitors to assess cough severity.7

REFERENCES

Health status questionnaires are designed to quantify quality of life numerically using the least number of questions. They are not a substitute for taking a good history. During the validation of the LCO, health-related issues particularly pertinent to females were evaluated. Several items, including stress incontinence, were excluded because only a minority of patients reported them. There was also evidence that alterations in health status due to symptoms such as stress incontinence were adequately captured by items that relate particularly to the psychosocial impact. It is important to note that health status questionnaires should be designed for use in the wider population rather than targeted to a specific subset of patients.

In conclusion, the LCO is a brief, well-validated and widely used health status questionnaire for patients with cough and can be used to detect gender differences.

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REFERENCES


Are patients on treatment for pulmonary TB who stop expectorating sputum genuinely culture negative?

In patients receiving treatment for pulmonary tuberculosis (TB), change in sputum culture from positive to negative is the principal outcome measure of a therapeutic response in both clinical practice and drug trials. Patients will often stop producing sputum early in the course of treatment. We have tested the assumption that “no sputum” means that the patient is “culture negative”, as this has never been confirmed experimentally.

We prospectively followed 16 patients with newly diagnosed pulmonary TB. Sputum samples were collected at diagnosis, during weeks 1 and 2, at months 1, 2, and 4, and on completing treatment. Those patients who were not producing sputum spontaneously had specimens collected by induction. This was performed in a purpose-built negative pressure isolation chamber (Elwyn E Roberts Isolators, Shropshire, UK) where patients inhaled 3% hypertonic saline via an ultrasonic nebuliser for 20 min. Samples were homogenised with Sputasol (Oxoid, Basingstoke, UK). A dilution series

Table 1 Gender differences in health status in patients with chronic cough assessed with the Leicester Cough Questionnaire (LCO)

<table>
<thead>
<tr>
<th></th>
<th>Females (%)</th>
<th>Males (%)</th>
<th>p Value (t test)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Number</td>
<td>112 (63)</td>
<td>66 (27)</td>
<td></td>
</tr>
<tr>
<td>Age, years (SEM)</td>
<td>56 (1)</td>
<td>54 (2)</td>
<td>0.5</td>
</tr>
<tr>
<td>Cough duration, years (SEM)</td>
<td>5.5 (0.9)</td>
<td>3.3 (0.6)</td>
<td>0.04</td>
</tr>
<tr>
<td>LCO total (range 3–21) (SEM)</td>
<td>13.5 (0.4)</td>
<td>14.9 (0.5)</td>
<td>0.01</td>
</tr>
</tbody>
</table>


Airway epithelial cells as guardians of immune homeostasis?

We read with interest the paper by Wang et al and accompanying editorial by Smyth showing that healthy murine airway epithelial cells (AECs) are potent inhibitors of dendritic cell (DC)-induced T cell activation. AECs infected with respiratory syncytial virus (RSV) lost this regulatory function, allowing activation of T cell responses and airway inflammation. These in vitro observations match with the high concentrations of pro-inflammatory mediators and cells found clinically in the bronchoalveolar lavage fluid of infants with RSV bronchiolitis.

The paper by Wang et al adds to a growing body of evidence that AECs are involved in maintaining airway immune homeostasis. Mayer et al previously showed that primary murine and immortalised human AECs induce an anti-inflammatory microenvironment inhibiting DC maturation and reducing T cell proliferation through constitutive secretion of transforming growth factor-β. Wang et al comment that further studies in human primary AECs are required to validate the findings in a clinical setting. Smyth also highlights the importance of research to investigate the function of AECs in health.

Primary AECs cultured from bronchoscopic brushings taken from clinically stable lung allograft recipients free from chronic allograft dysfunction represent a useful model to study AEC function in a healthy, steady state, albeit alloimmune environment. In a recent paper we have shown that epithelial cell-conditioned medium from stable lung allografts drives the production of macrophage-like cells from monocytes rather than DCs. It is unclear whether this effect only occurs in the airway of lung transplant recipients or if it reflects a general role for AECs in the homeostasis of DC populations in the lung. Nonetheless, our findings provide complementary human evidence to the murine observations of Wang et al and indicate that, in a steady state, AECs may be important in local immune homeostasis and promote an anti-inflammatory and pro-phagocytic airway milieu.

An emerging hypothesis that encompasses these observations is therefore that, in the healthy state, AECs regulate local immune homeostasis in the epithelium and promote anti-inflammatory conditions in the airway. In response to epithelial damage such as RSV infection, danger signals are released into the microenvironment by AECs which drive the production and maturation of professional antigen-presenting DCs, promoting T cell activation and airway inflammation.

To explore this hypothesis more fully we suggest that future studies should include primary human tissue in both health and disease, and that this strategy can complement and extend animal studies.
Assessment of gender differences in health status with the Leicester Cough Questionnaire (LCQ)
S S Birring and I D Pavord

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