Conference on ‘Malnutrition matters’

Symposium 2: The skeleton in the closet: malnutrition in the community

Encouraging appropriate, evidence-based use of oral nutritional supplements

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With the considerable cost of disease-related malnutrition to individuals and to society (estimated to be >£13 x 10^9 for the UK, 2007 prices), there is a need for effective and evidence-based ways of preventing and treating this condition. The wide range of oral nutritional supplements that may be prescribed for the dietary management of malnutrition and other conditions account for only about 1% (about £99 x 10^6, 2007 data) of the prescribing budget in England. Systematic reviews and meta-analyses consistently suggest that ready-made, multi-nutrient liquids which may be prescribed can improve energy and nutritional intake, body weight and have a variety of clinical and functional benefits in a number of patient groups. Meta-analyses have repeatedly shown that oral nutritional supplements produce significant reductions in complications (e.g. infections) and mortality, and a recent meta-analysis shows a reduction in hospital admissions (OR 0.56 (95% CI 0.41, 0.77), six randomised controlled trials). Such benefits suggest that the appropriate use of oral nutritional supplements should form an integral part of the management of malnutrition, particularly as there is currently a lack of evidence for alternative oral nutrition strategies (e.g. food fortification and counselling). As with all therapies, compliance to oral nutritional supplements needs to be maximised and the use monitored. To make sure that those at risk of malnutrition are identified and treated appropriately, there is a need to embed national and local policies into routine clinical practice. In doing so, the economic burden of this costly condition can be curtailed. As recently suggested by the National Institute for Health and Clinical Excellence, substantial cost savings could be made if screening and treatment of malnourished patients was undertaken.


Healthcare systems with budgetary constraints focus on the use of cost-effective evidence-based treatments, particularly during economic downturns. Therefore, in the current economic climate, it is becoming increasingly important to identify conditions promptly and to treat them in the most clinically and cost-effective way. This applies to the treatment of disease-related malnutrition, a condition that is prevalent in hospital inpatients (10–60% of hospital admissions are at risk of malnutrition), outpatients (15–30%) and in individuals living in the community (including free-living older people (aged 65 years and over), those visiting general practice and those living in care homes or sheltered housing, 14–44%) (see Fig. 1 for prevalence of disease-related malnutrition identified with the ‘Malnutrition Universal Screening Tool’ (‘MUST’), see www.bapen.org.uk for screening criteria)(1–10). At any given point in time, more than 3 million people in the UK are malnourished with most (approximately 93%) living in the community(11). Disease-related malnutrition is detrimental physiologically and clinically, impairing the quality of life and delaying recovery from illness(12). Data suggest that disease-related malnutrition doubles the risk of

Abbreviations: ESPEN, European Society for Clinical Nutrition and Metabolism; ‘MUST’, ‘Malnutrition Universal Screening Tool’; NICE, National Institute for Health and Clinical Excellence; ONS, oral nutritional supplement; RCT, randomised controlled trial.

*Corresponding author: Dr Rebecca J. Stratton, fax +44 2380 794945, email R.J.Stratton@soton.ac.uk
mortality in hospital patients and triples mortality in older patients (aged 65 years and above) in hospital and after discharge\(^{(5,13)}\). Disease-related malnutrition increases the use of health care (hospitalisations, General Practitioner visits and support required after hospital discharge\(^{(1,3,12–16)}\)) and latest estimates suggest that this condition costs the UK more than £13 \(\times 10^3\) annually (2007 data; Fig. 2)\(^{(17)}\).

Despite the enormous public expenditure on malnutrition, the expenditure on treatments which may be prescribed for the management of this condition remains low. In 2007, the total cost of prescriptions in England was £8·372 \(\times 10^6\) (£83·72·7 \(\times 10^5\)). Of this, 1·88% (£157 \(\times 10^3\)) was for oral nutritional supplements (ONS) and approximately £56 \(\times 10^3\) (approximately 0·7%) was for enteral tube feeds\(^{(18)}\). The costs of other forms of nutritional support, including the costs of dietetic time to manage malnourished patients and the costs of other interventions (food snacks, food fortification) remain unclear and need to be elucidated.

Considering the enormous costs of disease-related malnutrition, a condition that is largely treatable, prompt identification with screening, followed by the most appropriate, effective, evidence-based treatment is recommended\(^{(1,19,20)}\). The National Institute for Health and Clinical Excellence (NICE), recently released cost saving guidance, within the top four of which was nutrition support (in the form of oral nutrition support, tube feeding and parenteral nutrition). Specifically, NICE suggested that improving the systematic screening, assessment and treatment of malnourished patients (NICE CG32 guideline) could lead to an estimated cost saving of £28·472 per 100,000. NICE suggests that ‘If this guidance (CG32) was fully implemented and resulted in better nourished patients then this would lead to reduced complications such as secondary chest infections, pressure ulcers, wound abscesses and cardiac failure. Conservative estimates of reduced admissions and reduced length of stay for admitted patients, as well as reduced demand for General Practitioner and outpatient appointments indicate that significant savings are possible\(^{(21)}\).

A pragmatic programme of screening implementation (using ‘MUST’) in care homes in Peterborough Primary Care Trust highlights some such benefits\(^{(22)}\). A programme involving education and training on malnutrition, screening and treatment using the framework of ‘MUST’, locally agreed care plans and monitoring, improved the documentation of nutritional status, the proportion of residents screened and the use of appropriate care plans. After the implementation of the screening programme, significant reductions in the number and duration of hospital admissions were observed, associated with a significant cost saving of about £600 per resident over 3 months\(^{(22)}\). This could be equivalent to approximately £1 \(\times 10^5\) annually if extrapolated to the entire population of care home residents in the UK, although the costs of training and screening would need to be taken into account in order to estimate the overall net cost saving. Similar improvements of nutritional care and outcome have been observed in other settings where screening has been implemented. Rypekama et al.\(^{(23)}\) showed improvements in nutritional status and fewer complications in older (>65 years) inpatients with a multi-disciplinary intervention involving a screening programme and Stratton and coworkers found an increase in the documentation of nutritional information, use of care plans and shorter hospital stay in neurology wards after ‘MUST’ implementation (RJ Stratton, M Collins and M Elia, unpublished results). In another study, early nutritional screening and the treatment of malnourished patients reduced the length of hospital stay in malnourished patients who had low handgrip strength (i.e. frail patients). The authors concluded that to shorten the mean length of hospital stay by 1 d for all malnourished patients, a mean investment of 76 euros (£69) in nutritional screening and treatment would be needed (only about one-quarter of the cost of one bed day)\(^{(24)}\).

Implementing a screening programme is an effective way to identify those who do and do not need nutritional support, such as help with meals, feeding or shopping or artificial nutrition. Most patients who are malnourished or
at risk of malnutrition can be managed using the oral route, often with a variety of dietary approaches including dietary modification (fortification, extra snacks, etc.), counselling by a dietitian and/or ONS\textsuperscript{25}. However, with the rise of evidence-based practice, there is a need to demonstrate the effectiveness of these different strategies while making sure that they are used appropriately and their efficacy is maximised.

Although the focus of this review is the use of ONS, it is important to note that there is little formal evidence to support many of the other strategies used in the management of malnutrition, such as food snacks, fortification and dietary counselling\textsuperscript{26}. Although the clinical and cost effectiveness of such strategies appear to be undocumented, they are widely advocated across health care settings, especially in primary care, in an attempt to reduce the use and costs of products which may be prescribed. Policies that focus only on reducing costs by withholding or stopping treatments are considered unsatisfactory, since they do not assess the clinical and economic consequences of this reduction and do not represent cost effectiveness\textsuperscript{27}. Malnutrition that is untreated or ineffectively treated is likely to lead to a poorer quality of life, more complications, greater health care use and greater total health care expenditure. There is an additional financial implication for patients if prescribed forms of nutritional support are withdrawn or withheld, as the individual becomes responsible for acquiring and paying for the extra food or over the counter products that are recommended. An additional risk is that patients may then seek unsafe or unsuitable alternative forms of over the counter supplements, some of which may be unbalanced and potentially dangerous for their condition. In such patients, the end result could be increased costs to deal with unforeseen complications. Therefore, the implementation of such policies needs to be carefully considered and the effects monitored frequently to assess their clinical and cost effectiveness in both the short and long term.

**Oral nutritional supplements**

Many individuals while acutely or chronically ill are unable to consume the energy and/or nutrients they need from food alone, for a wide variety of reasons\textsuperscript{12}. These are broadly divided into two main causes: a variety of disease-related factors and the inadequate availability and quality of food. Disease-related factors are many and include anorexia, nausea and vomiting, changes in taste and smell, painful or dry mouth, disorders of swallowing, breathlessness, difficulties in chewing, fatigue and other specific side-effects of surgery, drugs or inflammatory conditions. Disease-related psychosocial problems may also impair food intake, including depression, anxiety, social isolation and unappealing meal environments. For individuals at home, inadequate availability of food may be due to inadequate resources (finances and cooking facilities), poor access to shops or physical difficulties affecting food preparation. Within institutions, the quality, timing and presentation of food may limit intake. ONS offer a useful and effective strategy to meet this deficit in nutrient intake. ONS are typically multi-nutrient containing a mix of macronutrients (protein, carbohydrate and fat) and micronutrients (vitamins, minerals and trace elements). Ready-made, nutritionally complete supplements are also energy dense (mostly containing 6–3 kJ (1.5 kcal)/ml to 10–1 kJ (2.4 kcal)/ml, 1.26 MJ (about 300 kcal) per serving (125–220 ml)) and provide a good source of protein (10–20 g per 1.26 MJ (about 300 kcal) serving) and a balance of micronutrients. Most are liquids that are available ready-made and so are convenient for individuals to use at home when ill and for health care professionals to administer in busy community and hospital settings, such as care homes and hospital wards. Powder supplements (to be reconstituted with whole milk before consumption) are also available on prescription or over the counter, although currently there is limited evidence to support their use\textsuperscript{12,20}. The current evidence base supporting supplements is almost entirely restricted to ready-made types\textsuperscript{12,20} and highlights their effectiveness for use in patients with or at a risk of malnutrition, particularly older individuals, those who are acutely ill and perioperative patients. In the UK, there are a number of prescribed indications for the use of ONS which extend beyond disease-related malnutrition (and vary according to the supplement type; see Table 1 for a full list).

**Evidence-base for oral nutritional supplements**

A systematic review, which may or may not include a meta-analysis, has traditionally been considered as the best way of assessing the evidence-base for an intervention, such as ONS (type I in the hierarchy of evidence\textsuperscript{28}). This is particularly so when the review is undertaken by those with a good understanding of the clinical use of the treatment, in combination with statistical expertise. In the last decade, an increasing number of systematic reviews have been undertaken to examine the effectiveness of oral nutrition support strategies in the management of malnutrition. The majority of these systematic reviews have focused on the use of ONS, the strategy for which there is the greatest number of individual trials available, usually in comparison with routine care (no nutritional support)\textsuperscript{12,20}. Such reviews have had a major influence on the guideline development by national and international organisations, including non-governmental and governmental bodies. Indeed, there are already a substantial number of guidelines and standards referring to the use of ONS. Within Europe, these include guidelines from British organisations, such as the British Association for Parenteral and Enteral Nutrition, NICE and the Scottish Intercollegiate Guidelines Network (SIGN) as well as the European Society for Clinical Nutrition and Metabolism (ESPEN) (summarised in Table 2).

**Clinical outcomes and oral nutritional supplements**

Systematic reviews and meta-analyses\textsuperscript{12,20,29–51} consistently suggest that ONS produce a range of clinical benefits including reduced mortality and fewer complications, such as wound and chest infections, pressure and leg ulcers. These are described in more detail in a ‘review of reviews’\textsuperscript{12} and a summary of these meta-analyses is shown in Table 3. In most trials, the daily reported intake of ONS was typically between 1.05 and 2.52 MJ
Consider ONS as part of the care plan for the treatment of malnutrition:

- Liquid ONS can be used if improvements in energy, protein, and micronutrient intakes are required. Liquid supplements tend not to suppress appetite or voluntary food intake and in some patient groups may stimulate food intake. A supplement containing a mix of macronutrients and micronutrients is important. Supplements with higher energy density may improve compliance and nutritional intake.
- For patients requiring longer term oral nutritional support, often in the community, it is likely that a variety of types of ONS and concomitant use of other dietary strategies would be beneficial to maintain improvements in nutritional intake.
- Liquid ONS can be used to attenuate weight loss in the acutely ill patient or aid weight gain in chronically ill patients. Improvements in weight (>2 kg), especially in the underweight, are associated with improvements in function in the chronically ill.
- High-protein ONS may be beneficial in preventing pressure ulcers in high risk groups and in improving clinical outcome in fractured neck of femur patients.
- Liquid, ready-made, multi-nutrient supplements (about 250–600 kcal (1-05–2.52 MJ) daily) can improve clinical outcome, reducing mortality and complications (infections and pressure ulcers), and healthcare use (reducing hospital admissions and length of stays).

Other oral nutritional support strategies that can form part of the care plan for the management of malnutrition include food snacks, food fortification and dietary counselling (although there is currently little evidence to support their efficacy). Although dietetic support may be ideal for all patients identified with malnutrition, limited resources in many countries mean that this is an impractical strategy. However, if time and resources permit, or if there is clinical concern and specialised food or artificial nutritional support may be required, referral should be made to a dietitian, or nutrition support team for detailed nutritional assessment. This should consider intakes and requirements for energy, protein and micronutrients. Any specific identifiable nutrient deficiencies (trace elements, minerals and vitamins) should be corrected where possible.

The goal of any nutritional treatment, including ONS, should be identified for an individual patient at the onset of treatment. Regular and frequent monitoring is essential to assess the effectiveness of ONS, including the acceptability and compliance with ONS, changes in clinical and nutritional status, whether ONS are still required and need for other forms of nutritional support (e.g. changes to diet, tube feeding, etc.).

Table 1. Recommendations for use of oral nutritional supplements in clinical practice

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<thead>
<tr>
<th>Use oral nutritional supplements (ONS) as part of the management of disease-related malnutrition and for the other listed prescribed indications (pre-operative preparation of malnourished patient, inflammatory bowel disease, short bowel syndrome, intractable malabsorption, post-total gastrectomy, dysphagia, bowel fistulae, growth failure and hypoproteinaemia) (varies according to supplement, see British National Formulary). For disease-related malnutrition, routine screening can be used to identify at-risk individuals across health care settings. A valid, evidence-based tool such as ‘MUST’ should be used. Implement appropriate nutritional treatment as part of a care plan as soon as possible. Consider ONS as part of the care plan for the treatment of malnutrition:</th>
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of complications with ONS (12,20,29–31,33,35), including infections (wound, chest, urinary, etc.), incomplete wound healing, pressure ulcers and total complications (see Table 3 for meta-analysis results). One systematic review showed a significant reduction in the development of one specific complication (pressure ulcers) with ONS vs. routine care (OR 0.75 (95% CI 0.62, 0.89), four RCT, n 1224) (Table 3) (29). In this meta-analysis, most studies used a liquid, high protein ONS for between 2 and 26 weeks, across hospital and long-term care. The other systematic reviews and meta-analyses mostly reported significant reductions in a range of complications with ONS use in hospital, community or combinations of health care settings (12,20,31,33,36). In most cases, liquid, ready-made, multi-nutrient ONS were used, with reported intakes of 250–600 kcal (1.05–2.52 MJ)/d. In one systematic review including only high-protein ONS (containing >20% energy from protein), marked reductions in complications were also observed (36). Systematic reviews of trials in patients in the peri-operative period have consistently indicated fewer complications with ONS use before, during and after hospitalisation, including total complications, infectious complications, major complications and intra-abdominal/thoracic complications (12,32,33,37). Similarly, in hip fracture patients, a Cochrane review (36) suggests significantly fewer patients with an unfavourable outcome (mortality and complications) (relative risk 0.52 (95% CI 0.32, 0.84), three RCT, n 139) with ONS vs. routine care.

The reduction in complications with ONS does not appear to differ between studies in which the mean BMI is <20 from those in which it is >20 kg/m² (12). Indeed, reductions in complications are observed in surgical patients (e.g. gastrointestinal surgery (32,33,37)), who are not obviously thin but in whom poor nutritional intake in the post-operative period may have contributed to the development of complications.

Functional outcomes and oral nutritional supplements.

Due to the wide range of functional outcomes measured in trials of ONS across different patient groups, it can be difficult to synthesise the evidence into a single meta-analysis of RCT. One systematic review that considered the impact of ONS on function in detail was Stratton et al. (12). Within this systematic review, a number of individual studies in both hospital and community patients were found to show significant improvements in functional measures with ONS in groups such as older people, patients with liver disease and those undergoing surgery. The functional improvements included muscle strength, quality of life, immune function, walking distances and activities of daily life. This review also reported that in chronically ill patients in the community, functional benefits were more likely to occur in underweight individuals (BMI <20 kg/m²) who gained weight (>2 kg) with ONS. Another review that attempted to integrate a functional outcome (muscle strength) from only five trials in older people into a meta-analysis reported no significant effects of ONS (31). Further research is warranted to determine the effects of ONS on important functional measures, especially quality of life, and other ‘Patient reported outcome measures’, which are now incorporated into policies within the National Health Service in England.

Nutritional outcomes and oral nutritional supplements

Nutritional status. Systematic reviews and meta-analyses that assess nutritional status consistently indicate significant improvements with ONS (12,31,39). The most common indicator of nutritional status assessed has been body weight. All meta-analyses that examined this outcome appeared to report a significant improvement with ONS relative to routine care in all settings (hospital, long-term care and community) (12,31). Meta-analyses by Milne et al. (31) indicated significant improvements in the percentage weight change with ONS relative to routine care in older patients in long-term care (weighted mean difference 2.51 (95% CI 1.73, 3.20%), in the community (2.25 (95% CI 1.72, 2.70%) and in the hospital (1.75 (95% CI 1.12, 2.30%)) with a wide range of conditions, although it was unclear whether they were malnourished or not (31). Another meta-analysis indicated a significant improvement in weight with ONS relative to dietary advice (weighted mean difference 1.09 (95% CI 0.29, 1.90) kg) (36). In general, ONS use aided weight gain in chronically ill patients in the community setting and attenuated weight loss in studies in acutely ill patients in the hospital (12). Improvements in weight with ONS were seen more frequently in community trials in which the mean BMI was <20 kg/m² (∼0.5 kg/m²) than in other RCT (e.g. gastric bypass surgery (37)). The composition of weight gain achieved with ONS (e.g. lean tissue and fat mass) was neither assessed nor unclear in many studies, although a few RCT indicated significant improvements in lean tissue or body fat. In most trials, upper-arm anthropometry was used, possibly as an easier and more practical method to use clinically than other more sophisticated body composition techniques (30). The review by Milne et al. (31) showed a significant increase in the mid-arm muscle circumference with ONS use in older hospital patients (weighted mean difference 1.41% (95% CI 0.46, 2.35%), six RCT). Irrespective of the composition of weight change, the increases in weight observed with ONS were linked to functional benefits, particularly in older people and in patients with chronic obstructive pulmonary disease (12). In studies in the acute setting, improvements in clinical outcome were often associated with very small changes in body weight (<1–2 kg) as the periods of supplementation were sometimes as brief as 1 week. It is possible that the improvements in outcome observed with ONS, particularly in the acutely ill, occur via mechanisms that are largely independent of changes in body mass. One or more nutrients (macronutrients or micronutrients) provided by liquid, multi-nutrient ONS, could influence immune/inflammatory responses, and hence clinical outcome irrespective of changes in body weight or lean tissue mass (31).

Nutritional intake. Systematic reviews that address the impact of ONS on nutritional intake consistently show improvements in total energy intakes in acutely and chronically ill patients in hospital and community settings (12,20,39). Some indicate significant improvements in the intakes of protein and micronutrients (12,20,20). Although often not assessed in trials, it is likely that total nutrient intakes (including micronutrient intakes)
Table 2. A summary of guidelines from British Association for Parenteral and Enteral Nutrition, National Institute for Health and Clinical Excellence, Scottish Intercollegiate Guidelines Network and European Society for Clinical Nutrition and Metabolism referring to oral nutritional supplement (ONS) use

<table>
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<tr>
<th>Body</th>
<th>Patient group</th>
<th>Title</th>
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<tr>
<td>British Association for Parenteral and Enteral Nutrition</td>
<td>Malnutrition Advisory Group(14) Patients in the community</td>
<td>Guidelines for detection and management of malnutrition November 2000</td>
<td>5.3.2. Treatment typically begins with food, but may progress to the use of supplements. In some patients it may begin with both food and supplements. 5.3.2.2. If ordinary food is ineffective in improving nutritional status and ineffective in achieving the goals set at the beginning of treatment, nutritional supplements (mixed micronutrient and macronutrient supplements in solid or liquid form) can be of value. This is because they are readily available, easy to consume between meals, require little or no preparation and are largely additive to food intake in undernourished subjects (A)</td>
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<td>National Institute for Health and Clinical Excellence (NICE) NICE (CG32)(20) All patients in hospital and in the community</td>
<td>Nutrition Support for Adults Oral Nutrition Support, Enteral Tube feeding and Parenteral Nutrition</td>
<td>Healthcare professionals should consider oral nutrition support* to improve nutritional intake for people who can swallow safely and are malnourished or at risk of malnutrition (A) Healthcare professionals should ensure that the overall nutrient intake of oral nutrition support* offered contains a balanced mixture of protein, energy, fibre, electrolytes, vitamins and minerals. (D(GPP)) Oral nutrition support* should be stopped when the patient is established on adequate oral intake from normal food. (D(GPP)) Peri-operative oral nutrition support* should be considered for surgical patients who can swallow safely and are malnourished. (B) Nutritional support/supplementation for the treatment of patients with pressure ulcers should be based on – nutritional assessment (using a recognised tool e.g. ‘MUST’) – general health status – patient preference – expert input supporting decision making (dietician or specialists) (D) If the BMI is low, patients should also be given nutritional supplements to increase their total energy intake, and be encouraged to take exercise to augment the effects of nutritional supplementation. (D)</td>
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<td>NICE and Royal College of Nursing(57) Patients with pressure ulcers (primary and secondary care) NICE(58) Chronic obstructive pulmonary disease</td>
<td>The management of pressure ulcers in primary and secondary care. A clinical practice guideline. September 2005 Chronic obstructive pulmonary disease. Management of chronic obstructive pulmonary disease in adults in primary and secondary care. Clinical guideline 12</td>
<td>Nutritional support should be initiated for people with stroke who are at risk of malnutrition. This may include ONS, specialist dietary advice and/or tube feeding</td>
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<td>NICE(59) Stroke patients</td>
<td>National Clinical Guideline for Diagnosis and Initial Management of Acute Stroke and Transient Ischaemic Attack</td>
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<td>Scottish Intercollegiate Guidelines Network (SIGN) SIGN (60) Older people – hip fractures</td>
<td>No. 111 Management of hip fracture in older people</td>
<td>Supplemeting the diet of hip fracture patients in rehabilitation with high energy protein preparations containing minerals and vitamins should be considered (A) Oral multi-nutrient feeds provide protein, energy, some vitamins and minerals, and may reduce complications while in hospital, although they have no effect on mortality. The presence of protein in an oral feed may reduce the number of days spent in rehabilitation (1++) One randomised trial observing stroke patients showed that oral nutrition supplementation is an effective method of improving nutritional status and clinical outcome (1++)</td>
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<td>SIGN(61) Stroke and dysphagia</td>
<td>No. 78 Management of patients with stroke: identification and management of dysphagia. A national clinical guideline. September 2004</td>
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<tr>
<td>SIGN(62) Post-operative</td>
<td>No. 77 Postoperative management in adults. A practical guide to postoperative care for clinical staff. August 2004</td>
<td>Patients who are malnourished either at the time of, or shortly following major abdominal or vascular surgery have a more rapid recovery of nutritional status, physical function and quality of life, if given nutritional advice and prescribed routine oral supplements in the immediate postoperative period and following two months (1++)</td>
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<td>European Society for Clinical Nutrition and Metabolism (ESPEN)</td>
<td>Pancreatic disease</td>
<td>ESPEN guidelines on enteral nutrition: pancreas</td>
<td>1.10 (Acute pancreatitis) Oral feeding (normal food and/or ONS) can be progressively attempted once gastric outlet obstruction has resolved, provided it does not result in pain and complications are under control (C).</td>
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<td>Gastrointestinal diseases (Crohn’s disease, ulcerative colitis, and short bowel syndrome)</td>
<td>ESPEN guidelines on enteral nutrition: gastroenterology</td>
<td>2.4 (Chronic pancreatitis) 10–15% of all patients require ONS (C) 3.6 (Crohn’s disease) In case of persistent intestinal inflammation (e.g. steroid dependent patients) use ONS (B) 3.1/3.2 Use ONS in addition to normal food to improve nutritional status and to eliminate consequences of undernutrition such as growth retardation (A) 4.1 Using ONS, a supplementary intake of up to 600 kcal (2.52 MJ)/d can be achieved in addition to normal food (A)</td>
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<td>HIV and other chronic infectious diseases</td>
<td>ESPEN guidelines on enteral nutrition: wasting in HIV and other chronic infectious diseases</td>
<td>2.4 (HIV) Diarrhoea does not prevent a positive effect of ONS on nutritional status (A) 2.2 (HIV) Nutritional counselling with ONS, or counselling alone, are equally effective at the beginning of nutritional support and/or for preserving nutritional status (B) 2.2 (HIV) In settings where qualified nutritional counselling cannot be provided, ONS may be indicated in addition to normal food but this should be limited in time (C) 4.0 (Infectious diseases) Nutritional support should be given to patients with undernutrition resulting from infectious diseases – prefer ONS (B)</td>
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<td>Surgery</td>
<td>ESPEN guidelines on enteral nutrition: surgery including organ transplantation</td>
<td>1.3 (Heart failure) Enteral nutrition (by means of ONS or tube feeding) is recommended in cardiac cachexia to stop or reverse weight loss (C) 2.5 (Chronic obstructive pulmonary disease) Frequent small amounts of ONS are preferred to avoid postprandial dyspnoea and satiety and to improve compliance (B)</td>
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<td></td>
<td>Chronic heart failure and chronic obstructive pulmonary disease</td>
<td>ESPEN guidelines on enteral nutrition: cardiology and pulmonology</td>
<td>6.0 Before transplantation, in undernutrition, use additional ONS (C)</td>
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<td></td>
<td>Geriatrics</td>
<td>ESPEN guidelines on enteral nutrition: geriatrics</td>
<td>4.1 Encourage patients who do not meet their energy needs from normal food to take ONS during the pre-operative period (C) 6.0 Before transplantation, in undernutrition, use additional ONS (C)</td>
</tr>
</tbody>
</table>

*Levels of evidence: 1++, high-quality meta-analyses, systematic reviews of randomised controlled trial (RCT) or RCT with a very low risk of bias; 1+, well-conducted meta-analyses, systematic reviews or RCT with a low risk of bias; 2++, high-quality systematic reviews of case-control or cohort studies, high-quality case-control or cohort studies with a very low risk of confounding bias; 2+, well-conducted case-control or cohort studies with a low risk of confounding bias. 3, case reports; 4, expert opinion.

†Grades of recommendations: A, at least one meta-analysis, systematic review or RCT rated as 1++, directly applicable to the target population or a body of evidence consisting principally of studies rated as 1+ directly applicable to the target population and demonstrating an overall consistency of results; B, a body of evidence including studies rated as 1+, well-conducted meta-analyses, systematic reviews or RCT with a low risk of bias; 2++, high-quality systematic reviews of case-control or cohort studies, high-quality case-control or cohort studies with a very low risk of confounding bias; 2+, well-conducted case-control or cohort studies with a low risk of confounding bias. 3, case reports; 4, expert opinion.

‡Oral nutritional support; includes ONS, dietary advice, food fortification, etc.
are improved with ONS. This is partly because liquid supplements contain a range of macronutrients and micronutrients and also because liquid ONS do not appear to substantially suppress voluntary food intake or appetite\(^{12,41}\). This may be particularly important in older patients and those suffering from anorexia, in whom liquid ONS have been shown to improve energy, protein and micronutrient intakes to a greater degree than isonenergetic food snacks in both the hospital and community environment\(^{42,43}\). Similarly, the Cochrane review of Baldwin et al. found significantly greater energy intakes were achieved with ONS than dietary advice (weighted mean difference 91 (95% CI 23, 159) kcal (0.38 MJ (95% CI 0.10, 0.67 MJ)), four RCT (n 138)\(^{39}\).

See Stratton et al.\(^{12}\) and Stratton\(^{44}\) for a more detailed review.

### Table 3. Effect of oral nutritional supplements (ONS) on mortality and complication rates (analyses of trials in a mix of patient groups; adapted from Stratton & Elia\(^{32}\))

<table>
<thead>
<tr>
<th>Systematic review</th>
<th>Intervention v. routine care</th>
<th>Mortality*</th>
<th>Complication*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stratton et al.(^{12}) (studies in a wider range of patient groups, hospital and community settings)</td>
<td>Mostly multi-nutrient, liquid ONS, 250–600 kcal (1·05–2·52 MJ)/d for &lt;1 week to 2 years</td>
<td>OR 0·62 (95% CI 0·49, 0·76), 17 RCT, n 2096</td>
<td>OR 0·29 (95% CI 0·18, 0·47), 10 RCT, n 494</td>
</tr>
<tr>
<td>NICE(^{29}) (studies in malnourished patients with a wide range of conditions in hospital and community settings)</td>
<td>Multi-nutrient proprietary ONS</td>
<td>RR 0·81 (95% CI 0·68, 0·97), 18 RCT, n 2564</td>
<td>RR 0·71 (95% CI 0·61, 0·82), 7 RCT, n 1001</td>
</tr>
<tr>
<td>Milne et al.(^{31}) (all studies with a mean age &gt;65 years, individuals with a wide range of conditions)</td>
<td>Mostly multi-nutrient, liquid ONS, some powders, 175–1000 kcal (0·74–4·2 MJ)/d for 10 d to 18 months</td>
<td>OR 0·86 (95% CI 0·74, 1·00), 25 RCT, n 6852</td>
<td>Unwell: OR 0·86 (95% CI 0·74, 1·00), 22 RCT, n 6630</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Aged &gt;75 years</td>
<td>OR 0·64 (95% CI 0·49, 0·85), 18 RCT, n 1611</td>
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<tr>
<td>Stratton et al.(^{29}) (studies in all patients at risk of developing pressure ulcers, hospital and community settings)</td>
<td>All multi-nutrient, liquid ONS, mostly high protein, 250–600 kcal (1·05–2·52 MJ)/d for 2 to 26 weeks</td>
<td>Geriatrics(^5)</td>
<td>OR 0·79 (95% CI 0·62, 0·89)(^{†}), 4 RCT, n 1224</td>
</tr>
<tr>
<td>Koretz et al.(^{40}) (^†) (studies in geriatrics and in perioperative patients – variety of conditions and both hospital and community settings)</td>
<td>Mostly multi-nutrient, liquid ONS</td>
<td>–</td>
<td>–</td>
</tr>
<tr>
<td>Cawood et al.(^{36}) (^†) (studies in patients with a wide range of conditions and in both hospital and community settings)</td>
<td>Ready-made, multi-nutrient ONS. High protein (&gt;20% energy from protein)</td>
<td>–</td>
<td>–</td>
</tr>
</tbody>
</table>

RR, relative risk; RCT, randomised controlled trials.

*An OR or RR < 1 indicates a lower incidence of mortality or complications in the supplemented patients than control patients. If the confidence interval does not pass 1·00, the result is statistically significant.

†In this meta-analysis the only complication assessed was pressure ulcers. In all other trials, a variety of complications were assessed (including wound infections, chest infections, pressure ulcers, etc.).

‡Added since publication of the ‘review of reviews’\(^{32}\).

§Data presented as absolute risk difference.
Orthopaedic surgery). The evidence-base for the use of nutritional support (ONS and tube feeding) to prevent pressure ulcers has been systematically examined (20). This review found that nutritional support had important clinical and statistically significant effects on reducing the development of pressure ulcers in at-risk populations. Based on this systematic review, a simple cost analysis indicated a reduction in costs with the use of ONS to prevent pressure ulcers (45). For all stages of pressure ulcers, there was a net cost saving in favour of ONS, which was significant for stage III (effect size 0.12 (95% CI 0.00, 0.11; P = 0.04)) and stage IV ulcers (0.12 (95% CI 0.01, 0.11; P = 0.04)). This corresponded to a net cost saving of £5 (stage I) to £460 per patient (stage IV) (2004 prices) (45).

Hospital admissions/readmissions. Studies are now starting to assess the effects of ONS use, either entirely in the community or partly in the community following initiation in hospitals, on hospital admissions and readmissions. Recently, two large RCT have highlighted significant reductions in hospital readmissions following the use of ONS in community settings, in older people with a wide range of conditions and in individuals with benign gastrointestinal disease (46, 47). Other trials have also shown lower (not significantly so) hospital admissions/readmissions with the use of ONS vs. routine care (48–51) (Fig. 3). When combined in a meta-analysis, the results highlight a highly significant reduction in hospital admissions with ONS (OR 0.56 (95% CI 0.41, 0.77), six RCT). The cost savings associated with such reduction in health care use are important, and the associated improvements to the patients’ quality of life are also likely to be hugely beneficial, but require verification.

Encouraging appropriate use and maximising compliance

ONS, like any type of treatment, need to be used appropriately, and as effectively as possible. In addition to using ONS for their prescribed indications (Table 1) and in line with the evidence-base for their use, there are other matters to consider. Nutrition support should be tailored to the individual as much as is feasible and an individual’s preference for the type of treatment they require should be taken into account, including the type of supplement they prefer, the time and way in which they receive and consume it (or other interventions that they may prefer). Of course, ensuring compliance to supplements is vital if improvements in nutritional intake and outcomes are required. Maximising the acceptability of supplements to patients may include using a variety of flavours, textures, temperatures, serving formats and consistencies. However, other patients are happy consuming the same supplement (type and flavour) for long periods of time. The energy density, volume and macronutrient profile of the supplement, the time and way in which it is administered and the duration of supplementation are factors that may affect compliance (12, 52). Recently, studies have suggested that increasing the energy density of liquid feeds improves both nutritional intake and compliance with supplementation, over the use of standard energy dense feeds in hospital and community patients (53–55). Increasing the energy density of ONS minimises the volume needed to be consumed by a malnourished patient, and so encourages compliance and nutritional intake (53–55). It is also likely that encouragement and education about the reasons for taking supplements would be beneficial. Overall, a greater understanding is still needed to maximise compliance with supplementation, especially over long periods of time, so that benefits can be achieved and sustained.

Like other treatments in clinical practice, it is recommended that the use of ONS (and other forms of nutritional support, such as dietary counselling) is appropriate and monitored. This typically involves reviewing the indications for the use of ONS, setting appropriate goals at the onset and reviewing these goals at intervals (the goals may include improvements in nutritional intake, weight and clinical/functional outcomes). The optimal dose and timing of supplementation for a patient is also important and should be reviewed regularly. It is necessary to review whether the termination of ONS is needed and whether the addition of other forms of nutritional support is needed. A recent publication supported by a number of national organisations (British Association for Parenteral and Enteral Nutrition, Parenteral and Enteral Nutrition Group, British Dietetic Association and National Nurses Nutrition Group) has highlighted some examples of good practice for the appropriate use of ONS in older people, while also summarising the evidence to support their use (56). The NICE guidelines also highlight the importance of monitoring the use of all forms of nutritional support (20).

Conclusion

Appropriate, evidence-based use of ONS should be an essential part of the management of disease-related malnutrition, particularly in the current economic environment we are facing. Malnutrition needs to be identified and treated effectively, since failure to do so can lead to disease complications and delay recovery from illness, at
enormous cost to the healthcare service. Prompt identification and treatment is required to attenuate the detrimental impact of malnutrition on health and the quality of life. The appropriate use of ONS should be an integral part of the management of disease-related malnutrition. Perhaps surprisingly, the use of ONS (mainly ready-made, multi-nutrient ONS) is the only form of oral nutritional support with a substantial evidence-base supporting its use. The evidence from systematic reviews and meta-analyses supports the use of ONS in a wide variety of conditions, but the evidence is strongest in the acutely ill, in older patients and in malnourished patients, in whom a range of clinical benefits, including a significant reduction in mortality and complications, such as infections and pressure ulcers, have been demonstrated. Appropriate use of ONS is likely to lead to economic benefits associated with reduced health care use, including a reduction in the number of hospital admissions and duration of hospital stay.

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References