Clinical Investigations

Spectrophotometric intracutaneous analysis: a new technique for imaging pigmented skin lesions

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Summary

Background Spectrophotometric intracutaneous analysis (SIA) is a new technique for imaging pigmented skin lesions and for diagnosing melanoma. The SIAscope produces eight narrow-band spectrally filtered images of the skin over an area of 24×24 mm with radiation ranging from 400 to 1000 nm.

Objectives To present the early results of a clinical trial with SIA.

Methods Spectrophotometric inputs from the skin were analysed using complex algorithms to return high-resolution information regarding total melanin content of the epidermis and papillary dermis, collagen and haemoglobin content as well as the presence of melanin in the papillary dermis.

Results Simple, highly reproducible and reliable features were identified, e.g. the presence of dermal melanin, collagen holes and 'erythematous blush' with blood displacement. These simple features were found to be highly specific $(80\cdot1\%)$ and sensitive $(82\cdot7\%)$ for melanoma in a dataset of 348 pigmented lesions (52 melanomas) and compared very favourably with dermatoscopy when analysed using receiver-operator characteristic curves.

Conclusions This first clinical trial with SIAscopy has yielded very promising results and delivers new, useful information to the clinician diagnosing pigmented skin lesions.

Key words: dermatoscopy, diagnosis, imaging, melanoma, SIAscopy, spectrophotometric intracutaneous analysis

The overwhelming majority of the literature documents the relentless world-wide progress of melanoma up the incidence league tables of all cancers in both incidence and mortality. The highest incidence is to be found in the Antipodes where a recent report from New Zealand indicates that Auckland has overtaken Queensland, Australia with a staggering 78 per 100 000 (crude rate). By comparison, the overall incidence of melanoma in the U.K. in 1995 was reported at 5000 cases per annum (written communication from Department of Medical Statistics, Cancer Research Campaign, London). Similarly, mortality statistics throughout the world show a changing trend.

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Crude mortality measurements show that this is still rising, ³ although the rate of increase is trailing off. Statistics such as these have led to the dramatic rise being termed as an 'epidemic' and 'an emerging medical catastrophe'. ⁴ Of all the prognostic factors, both clinical and histopathological, Breslow thickness of the primary lesion is by far the most important. ⁵ Prognosis is so intrinsically linked with this measurement that apparent differences in survival between patient groups are often negated or greatly reduced when they are stratified for tumour thickness. In addition, vertical growth phase melanomas have a worse prognosis that is dependent on other factors. ⁶ In essence, early diagnosis and excision are the key to the likely survival of the patient.

The clinical diagnosis of melanoma is acknowledged as challenging by most authors and has provoked

much debate.⁷⁻⁹ Diagnostic accuracy has been stated as being between 50% and 80%. Grin et al.8 demonstrated a small improvement in accuracy with time (60% vs. 64%), and Morton and MacKie⁹ demonstrated a significant improvement in accuracy with an examiner of 10 years' experience or more (60% vs. 80%). As a result of the difficult task that clinicians can face in achieving an early diagnosis of a heterogeneous disease entity, there arises the need for tools that can improve diagnostic accuracy. Systems that have been developed as adjuncts in the diagnosis of melanoma include highfrequency ultrasound, optical coherence tomography, confocal microscopy and magnetic resonance imaging. 10-13 Some or all of the following limit these techniques: availability, expense, imaging resolution and lack of useful diagnostic detail. However, dermatoscopy (epiluminescence microscopy, dermoscopy, oilemersion microscopy, skin-surface microscopy) is a useful technique that uses oil and a hand-held microscope, which allows visualization of morphological characteristics at the dermoepidermal junction. Clinicians then attempt to diagnose the presence of melanoma by analysing the lesion by colour, pattern and specific morphological features. In 1990, a consensus paper was published to standardize the terminology used to describe features seen in the surface microscopy of pigmented skin lesions and correlated them with their underlying histology. 14 Since then, a whole wealth of literature, papers and textbooks has been published regarding this technique. A critical and systematic review of this literature¹⁵ found that where the specificity and sensitivity of the clinician was high for clinical diagnosis then the dermatoscope had little to offer in addition. 16,17 However, using the dermatoscope with equivocal lesions increased the sensitivity for diagnosis between 10% and 27%. 18,19

The identification of dermatoscopy features is based on a visual interpretation of colours within a dermatoscopic view. These colours depend upon reflection and absorption by chromophores within the superficial skin according to their depth and concentration and also upon the wavelength of incident light interacting with them. However, we have shown 20–22 that it is possible for the same colour to arise from different combinations of microscopic structures. This complication may explain the high subjectivity in identifying dermatoscopic features, with the possibility of diagnostic error as a result. As an example, bluegrey areas are defined as 'superficial fibrosis with melanophages and/or malignant cells in the papillary dermis'. This feature has a specificity of 97% for

invasive melanoma and has been described as 'the single most important surface microscopic finding' in making a diagnosis.²³ However, the same appearance can also be produced by a reduction in the thickness of the papillary dermis or an overlying layer of hyperkeratosis.²⁰ Another complicating factor of dermatoscopy is that the visual appearance of a skin lesion through a dermatoscope is the result of the layering of microscopic structures. As an analogy, it is akin to looking down on the surface of a rain forest from a helicopter and trying to identify a particular type of fern through the layers of different trees and bushes. It is clear that dermatoscopy can perform a valuable role in the investigation of pigmented skin lesions by increasing the information available for consideration in what is often a difficult diagnostic task. This extra information is particularly useful because it allows, by visual interpretation of the dermatoscopy features, an understanding of the in vivo microarchitecture of the skin lesion. However, it is the necessarily subjective identification of these, often subtle, features that leads dermatoscopy to require thorough training and experience. As a result, dermatoscopy requires a steep learning curve. Studies have shown that trainees may produce a decrease in sensitivity for the diagnosis of melanoma at the outset of learning when compared with performance on clinical grounds alone.¹⁹

It was to address the operator-dependent and subjective nature of dermatoscopy in extracting information regarding the microstructure of a skin lesion that the SIAscope was developed. Spectrophotometric intracutaneous analysis (SIA) arose from research undertaken at the University of Birmingham in the U.K. where the colours arising from skin lesions were being investigated with the aim of identifying objective diagnostic information. 20-22 It became apparent that analysis of purely visual colour information was insufficient to extract reliable diagnostic information relating to the lesion's internal structure and that to extract such information it was necessary to extend the analysis into the infrared. The result of this research is a technique capable of rapidly imaging a skin lesion in vivo and allowing examination of the distribution, position and quantity of a range of chromophores, including melanin, blood and collagen within the papillary dermis and the position of melanin relative to the dermoepidermal junction. The clinical device employing this technique is termed a SIAscope and returns SIAgraphs showing these constituents for interpretation.

The SIAscope (Fig. 1). operates by probing the skin spectrally over an area of 24×24 mm or 12×12 mm



Figure 1. A SIAscope.

with radiation ranging from 400 to 1000 nm. For this trial the larger, 24-mm handset was used. Eight narrow-band spectrally filtered images of the skin are obtained, calibrated and used as inputs to a series of complicated computer algorithms that extract information regarding the microarchitecture of the skin. The wavelengths of the wavebands are chosen to interact preferentially with constituents of the skin.²⁰⁻²² The SIA algorithm first utilizes the infrared wavebands to ascertain the quantity of collagen within the papillary dermis for every point over the skin lesion. This is the crucial step for this technique and provides a necessary transformation on the wavebands allowing accurate extraction of total melanin and blood. The total melanin, collagen and blood SIAgraphs can now be displayed. The effect of these chromophores on the wavebands is then removed, allowing a key piece of diagnostic information, the presence of melanin below the dermoepidermal junction, to be identified. This is possible because the spectral remittance of melanin changes with regard to its position in the superficial anatomical layers of the skin, namely the epidermis and the papillary dermis. Using SIAscopy the accuracy of this identification is extremely high and forms the dermal melanin SIAgraph.²¹

In this paper, we aim to introduce the technique of SIA and to present the early results of a clinical trial under way in the U.K.

Materials and methods

The study was designed to enable the identification of features from SIAgraphs that may assist in the early identification of melanoma when compared with the histopathological examination from excisional biopsy of pigmented skin lesions. Patients referred for excisional biopsy of a pigmented skin lesion at either the plastic surgery or dermatology departments at Addenbrooke's and West Norwich Hospitals were recruited to the study and informed consent was obtained. The patients were photographed twice using conventional 35-mm slide film (Kodachrome ISO 64): once using a standard ×2 macro lens (Canon) and once using a dermatoscopy lens (Heine Dermaphot: ×10 magnification). Patients were asked questions about the lesion—change in size, shape and colour; change in sensation; bleeding and inflammation—and the maximum diameter of the lesion was measured in millimetres. This information and standard demographic data (age, sex, name and hospital number) were recorded and stored in a protected computerized database. SIAscopy was performed using the SIAscope (Astron Clinica, Cambridge, U.K.) and the returned SIAgraphs were also stored on a protected computerized database. The system takes approximately 5 s to acquire the images and a further 10 s to process the data and return them as visual information on a 15-inch monitor display using dedicated software (Astron Clinica). Excisional biopsy was then performed and the results from histopathological examination (used as the reference or gold standard) were obtained and matched with the data. Two senior clinicians, one of whom specializes in dermatopathology, made the histopathological diagnosis of melanoma. A proforma was used as a guideline for the final report. Information from this report includes diagnosis, microscopic detail and, in the case of melanoma, subtype, Breslow thickness, Clark's level, regression and growth phase.

Working definitions of possible features that could correlate with underlying significant histopathology were devised for the purpose of analysis (Table 1). Examples of these features can be seen in Figures 2 and 3. A dataset of 348 lesions collected over a period of 12 months (February 2000–February 2001) was analysed for the presence or absence of each of these features independently by two of the authors (S.C., M.M.) using standard SIAscope viewing facilities. Both experimenters were blinded as to the histopathological diagnosis of the lesion. Statistical analysis was performed on the dataset using Sigma-

Table 1. SIAscopy features defined

Feature	Definition	SIAgraph view
Melanin globules	A circumscribed aggregation or flocculation of melanin within the lesion	Total melanin
Blood globules	A circumscribed aggregation or flocculation of blood within the lesion	Blood
Blood displacement	A confluent, non-pixelated area demonstrating an absence of blood within the lesion	Blood
Erythematous blush	Blood	
	normal skin for 3/4 of the circumference of the lesion	
Dermal melanin	The presence of dermal melanin within the lesion not due to hairs	Dermal melanin
Dermal melanin globules	A circumscribed aggregation or flocculation of dermal melanin within the lesion	Dermal melanin
Collagen holes	An area within the lesion coinciding with melanin (dermal or total) demonstrating	Collagen
	an absence of collagen not due to hair follicles or sebaceous/sweat ducts	
Biaxial symmetry	Symmetry of shape and pattern in two perpendicular planes within the constraints of biological variability	Total melanin
Asymmetry	A lack of symmetry of shape and pattern in any plane	Total melanin

Stat version 2.03 and SPSS for Windows version 9.0 (SPSS Inc., Chicago, IL, U.S.A.) software on a personal computer to obtain specificity and sensitivity data for each of the features. Logistic regression analysis was performed to produce this information for combinations of the features. Reproducibility and reliability data were also produced in the form of the kappa (κ) statistic for each of the features. This statistic provides a measure of agreement beyond any agreement that may have occurred by chance, and takes a value between 0 (no agreement above that which would have occurred by chance) and 1 (perfect agreement) with an improving scale of agreement represented by intermediate values. 24

Results

The dataset comprised 348 lesions (311 patients: 200 female and 111 male), including 52 melanomas (one in 6.70 lesions). Of the melanomas, six were in situ, 11 had a Breslow thickness < 0.76 mm and in total 28 lesions had a Breslow thickness ≤ 1.0 mm (range in situ to 6.0 mm). The median and mean Breslow thicknesses were 1.0 mm and 1.52 mm, respectively, and the subtypes included 41 superficial spreading melanomas, nine nodular and two acral lentiginous melanomas. Most of the non-melanoma lesions were benign naevi, but overall the sample demonstrated a case mix that a clinician would usually be presented with in a dermatology clinic (Table 2).

Each feature was measured for its reliability (interobserver) and reproducibility (intraobserver) by using the κ statistic. Most features were highly reproducible and reliable (Figs 4 and 5), indicating good working definitions. Two features, namely blood globules and melanin globules, were found to be unreliable and irreproducible and so were discarded from further

statistical analysis. As expected, no single feature was both highly specific and sensitive: if a feature was highly sensitive, e.g. the presence of dermal melanin, it was at the expense of poor specificity (Table 3). Combinations of features were then analysed using multiple logistic regression analysis. Consistently useful and specific features were the presence of dermal melanin and blood displacement with erythematous blush, and the inclusion of the presence of collagen holes as a further feature increased the sensitivity. It was then decided to investigate the effect of adding clinical information to try and improve the specificity and sensitivity of the test and to approximate the timehonoured clinical progression of history, examination and investigation of the patient. It was found that for lesions of 6 mm and greater with dermal melanin and blood blush/displacement there was a sensitivity of 82.7% and a specificity of 80.1%. The positive predictive values of the tests listed in Table 3 were disappointing but the negative predictive value was consistently very high. The positive and negative likelihood ratios were also consistently good, meaning that the tests are likely to modify pretest odds significantly and therefore give useful diagnostic information.²⁵ For example, given no other information, the chances of a patient having a melanoma (the prevalence) in our sample was 1 in 7 (15%) but, on performing SIAscopy and finding an absence of dermal melanin the odds drastically shifted to 83:1 against (1.2%).

Using logistic regression analysis, receiver-operator characteristic (ROC) curves were constructed for combinations of features (Fig. 6). ROC curves are particularly useful when considering the performance of a diagnostic test, as they allow the user to assess its operating dynamics. By selecting a certain point on the ROC curve the clinician can see what the false-positive rate will be (*x*-axis) at any given sensitivity (*y*-axis).

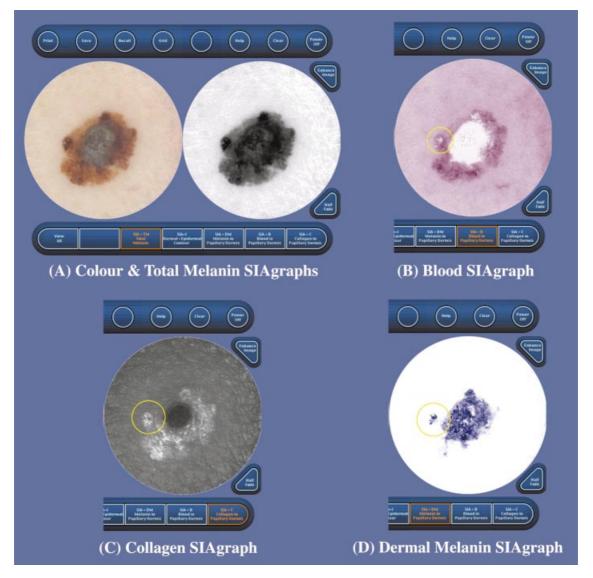


Figure 2. SIAgraphs of a superficial spreading melanoma (vertical growth phase, 2·5 mm Breslow thickness, Clark's level IV). (A) Colour SIAgraph (left) shows dermatoscopic view, especially blue–grey veil. Total melanin SIAgraph (black = melanin) enhances dermatoscopic criteria such as branched streaking and pseudopodia (right). (B) Blood SIAgraph (red = blood). The blood displacement is seen as a white area in the vertical growth phase nodule in the centre and also less obviously at the periphery at 9 o'clock (circled). The erythematous blush is quite striking in this lesion. (C) Collagen SIAgraph (white = collagen). The collagen hole is very large in the centre of this lesion. There is also a small collagen hole at the periphery at 9 o'clock (circled). (D) Dermal melanin SIAgraph (blue = dermal melanin present). There are large quantities of dermal melanin irregularly distributed in this lesion. There is also a separate focus at the periphery at 9 o'clock (circled).

Thus, the ideal test describes a curve that goes from the origin to the top-left corner to the top-right corner, and the ambivalent (useless) test describes a diagonal line from the origin to the top-right corner (dotted line). In simple terms, the closer an ROC curve is to the top-left corner, the better the test can be considered to be. From the data it was possible to construct an ROC curve for the revised seven-point checklist;²⁶ for comparison, the ROC curve for clinicians formally trained in dermatoscopy is also included.²⁷ It can be seen from this graph

that for different combinations of SIAscopy features the diagnostic performances are consistently much better than the revised seven-point checklist in detecting melanoma, and compare very well with dermatoscopy.

Discussion

This study assesses the clinical application of SIAscopy in the assessment of pigmented skin lesions. New diagnostic features that attempt to reveal the underly-

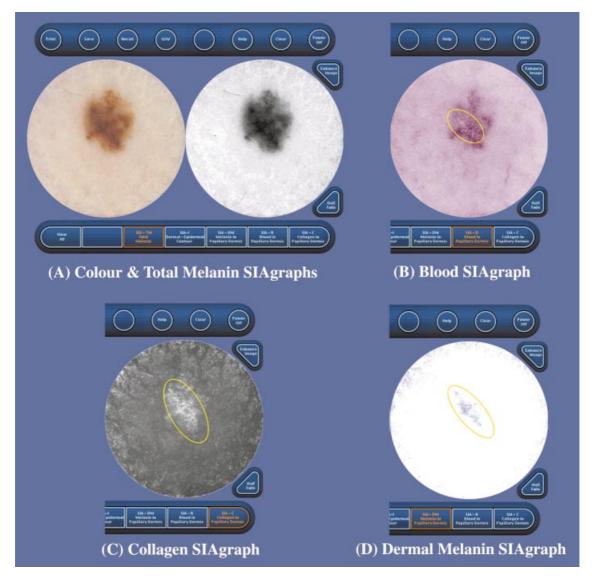


Figure 3. SIAgraphs of a superficial spreading melanoma (vertical growth phase, 0·3 mm Breslow thickness, Clark's level II). This early lesion demonstrates how SIAscopy may assist in the diagnosis of melanoma. The colour and total melanin SIAgraphs (A) are unremarkable. However, the blood SIAgraph (B) shows a subtle blood displacement with erythematous blush (circled). The collagen SIAgraph (C) shows no holes as this is only a Clark's level II melanoma, although there are large quantities of irregular collagen (circled) consistent with fibrosis. The dermal melanin SIAgraph (D) shows dermal melanin irregularly distributed across a large area of the lesion (circled).

ing histopathology have been identified by interpreting SIAgraphs that return information about the concentration and position of collagen, blood and melanin within the epidermis and papillary dermis. Significant findings were the presence of blood displacement with erythematous blush, collagen holes and dermal melanin (Figs 2 and 3). An invasive nodule or nest of melanoma cells will displace blood from the papillary dermis, while early dermal invasion will promote a strong immunocytic response characterized by local vasodilatation and fibrosis. We hypothesize that this

pathophysiological process is revealed by SIAgraphs as erythematous blush with blood displacement. The papillary dermis is the reflective layer of the skin and any light that passes through this does not return. A melanoma that is Clark's level III or greater will, by definition, have areas where the papillary dermis is replaced by invasion. These areas can be seen in collagen SIAgraphs as collagen holes (loosely termed as 'black holes' by our research group) where the melanoma nests absorb or transmit light in the papillary dermis. The presence of dermal melanin in

Table 2. Details of the dataset

Diagnosis	Number
Melanoma	52
Superficial spreading	41
Nodular	9
Acral lentiginous	2
Common naevi (compound,	185
junctional and intradermal)	
Dysplastic naevi	7
Blue naevi	12
Spitz naevi	7
Seborrhoeic keratoses	29
Lentigo	9
Basal cell carcinoma	21
Dermatofibroma	8
Mixed naevi	2
Haemangioma	2
Others	14
Total	348

the papillary dermis, as expected, is a useful diagnostic sign, with a sensitivity of $96\cdot2\%$, but the specificity was low at $56\cdot8\%$. In compound and intradermal naevi, the dermal melanocytes begin to atrophy or 'mature' and cease to produce melanin⁵ and this fact will help in distinguishing benign naevi from melanoma. However, pigmentary incontinence and melanophages are found in both and are confounding factors adding to the high false-positive rate as they will be observed in both sets of patients showing the presence of dermal melanin.

The results of this study indicate that the addition of *in vivo* information pertaining to the microscopic structure and architecture of a skin lesion can provide

useful information in the diagnosis of melanoma. The presence or absence of features in the SIAgraphs of pigmented skin lesions and the specificity and sensitivity data described compare very favourably with clinical examination and dermatoscopy. It is particularly worthwhile emphasizing that these early features described for SIAscopy of pigmented skin lesions are very simple to learn and use and this is reflected in the reliability and reproducibility of the κ statistics. For instance, the mental process of detecting dermal melanin runs along the lines of 'is there any blue on the screen where the lesion is?' Similarly, the process of detecting blood displacement runs along the lines of 'is there white on the screen where the lesion is?' In contrast, due to the difficulty in interpreting subtle features, dermatoscopy requires formal training and experience before diagnostic accuracy improves. Furthermore, formally trained clinicians, experts and pioneers in the field often disagree on dermatoscopic findings in lesions, indicating a high subjectivity in interpreting features seen, most notably the highly significant finding of a blue/grey veil. 28 Our specificity and sensitivity data of simple SIAscopic criteria compare very favourably against those published on clinicians after formal training in dermatoscopy²⁷ in the diagnosis of cutaneous melanoma, especially when one analyses the ROC curves for the two techniques. In addition, combinations of features can greatly modify the pretest odds for melanoma and this attribute is particularly desirable in any diagnostic test.²⁵ We

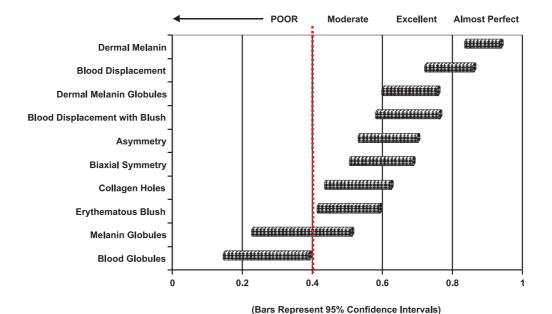


Figure 4. Reliability (interobserver agreement) of SIAscopy features.

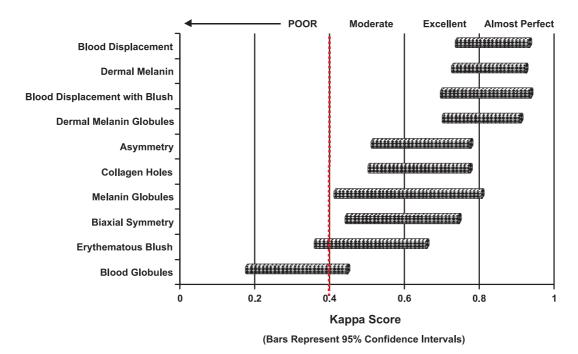


Figure 5. Reproducibility (intraobserver agreement) of SIAscopy features.

Table 3. Specificity and sensitivity analysis of SIAscopy features

Feature	Sensitivity (%)	Specificity (%)	PPV (%)	NPV (%)	Positive LR	Negative LR
Single features						
Collagen holes	78.8 (66-87.8)	74.0 (68.7–78.7)	34.7	95.2	3.03 (2.39-3.85)	0.29 (0.17-0.49)
Blood displacement	75.0 (57.3-87)	70.3 (63–76.6)	30.7	94.1	2.52 (1.99-3.19)	0.36 (0.22-0.57)
Erythematous blush	75.0 (61.8-84.8)	65.5 (60-70.7)	27.7	93.7	2.18 (1.74-2.72)	0.38 (0.24-0.62)
Blood displacement	63.5 (49.9-75.2)	84.8 (80.3-88.4)	42.3	93	4.17 (2.97-5.86)	0.43 (0.3-0.62)
with blush						
Dermal melanin	96.2 (87-98.9)	56.8 (51.1-62.3)	28.1	98.8	2.22 (1.93-2.56)	0.07 (0.02-0.26)
Dermal melanin globules	88.5 (77-94.6)	66.6 (61-71.7)	31.7	97	2.64 (2.19-3.19)	0.17 (0.08-0.37)
Asymmetry	76.9 (63.9–86.3)	62.2 (56.5–67.5)	26.3	93.9	2.03 (1.65-2.5)	0.37 (0.22-0.61)
Biaxial symmetry	3.8 (1.1-1.3)	70.3 (64.8–75.2)	2.2	80.6	0.13 (0.03-0.51)	1.37 (1.25-1.5)
Biaxial symmetry	96.2 (87-98.9)	29.7 (24.8-35.2)	19.4	97.8	1.37 (1.25-1.5)	0.13 (0.03-0.51)
not present						
Combined features						
Dermal melanin +	76.9 (63.9–86.3)	78 (73-82.4)	38.1	95.1	3.5 (2.67-4.52)	0.3 (0.18-0.47)
collagen holes						
Dermal melanin +	96.2 (87-98.9)	56.8 (51.1-62.3)	28.1	98.8	2.22 (1.93-2.56)	0.07 (0.02-0.26)
displacement with blush						
Dermal melanin +	92.3 (81.8-97)	65.9 (60.3-71)	32.2	98	2.71(2.25-3.23)	0.12 (0.05-0.28)
displacement with						
blush + biaxial						
symmetry not present						
Dermal melanin +	82.7 (70.3-90.6)	80.1 (75.1-84.2)	42.2	96.3	2.75 (2.13-3.52)	0.25 (0.13-0.43)
displacement with blush +	•					
diameter ≥6 mm						

Sensitivity = true positive/(true positive + false negative); specificity = true negative/(true negative + false positive); positive predictive value (PPV) = true positive/(true positive + false positive); negative predictive value (NPV) = true negative/(true negative + false negative); positive likelihood ratio (LR) = sensitivity percentage/(100 - specificity percentage); negative LR = (100 - sensitivity percentage)/specificity percentage. Values in parentheses are the 95% confidence interval.

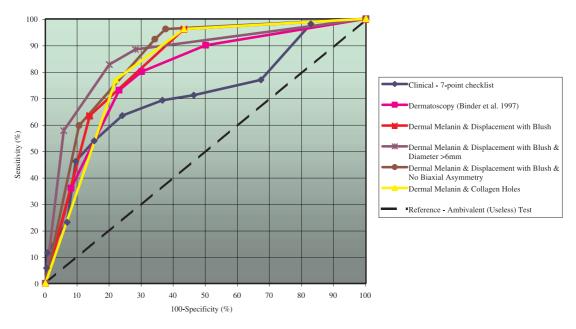


Figure 6. Receiver-operator characteristic (ROC) curves for combinations of SIAscopy features compared with clinical assessment and dermatoscopy (see text for explanation; dermatoscopy curve reproduced from Binder *et al.*²⁷ with permission from the publishers, Harcourt Health Sciences).

surmise that such an increase is due to the ability to identify specific processes occurring within a skin lesion confluent with melanoma such as the displacement of collagen by tumour cells and the vasodilatation and possibly angioneogenesis of papillary blood vessels.

Future work will produce probability data for diagnostic groupings or classes with certain combinations of features to facilitate the decision process for the clinician. The definitions used were deliberately designed as Boolean operators (i.e. 'true/false' or 'yes/no') and these could be refined and stratified. As a result of the Boolean nature of the features, examination of the data did not include a 'pattern analysis' type approach to the assessment of the pigmented lesions that is so useful when performing a dermatoscopic diagnosis. Pattern analysis allows the clinician to take into account the overall appearance of the lesion and so helps to increase the specificity (reduce the false-positive rate). 18,26 For example, in our dataset, blue naevi tended to be classified as melanomas because they usually demonstrated dermal melanin, collagen holes and blood displacement. However, the uniformity and regularity of these features and their sharp demarcation with the surrounding normal skin readily identifies them as blue naevi. Furthermore, the analysis was necessarily sterile and artificial in that no clinical diagnosis or dermatoscopic appearance was taken into account. Theoretically, it is reasonable to

assume that the inclusion of such information would further improve the performance of the clinician.

A limitation of this study is the lack of direct comparison of the SIAscopy data with the dermatoscopy images. To become proficient in dermatoscopy requires formal training of at least 9 h. 19 None of the authors had been formally trained in dermatoscopy and any attempts to diagnose melanoma solely from these images could lead to a worse performance than diagnosing pigmented lesions on clinical grounds alone.19 It was felt that it would be difficult to appraise the dermatoscopy images without biasing the performance of the technique due to lack of experience. Similarly, dermatoscopy is not universally practised in the U.K. and this may be due in part to an absence of formal training courses, while those that are available in continental Europe are not undertaken in English.

With a much larger dataset and with improving experience of SIAscopy, newer features will be described and tighter definitions will be developed. Methods that employ automated imaging analysis techniques are being investigated in order to increase reliability and reproducibility in the identification of SIAscopy features such as asymmetry and collagen holes. Additional studies are planned that will directly compare the performance of dermatoscopy with SIAscopy. As further studies are required to extend the number of lesions

examined, an international study involving the U.K., Germany, Denmark and Australia is under way.

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