

**FABRICATION OF GOLD NANOPARTICLES AS PAPER-BASED
MICROFLUIDIC DEVICE FOR COLORIMETRIC ASSAY OF HUMAN
CHORIONIC GONADOTROPIN**

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AUTHOR DECLARATION

I hereby declare that the work in this thesis is my own except for quotations and summaries, which have been duly acknowledged.

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ABSTRAK

Pengesanan gonadotropin korionik manusia (HCG) adalah salah satu daripada banyak kaedah diagnostik yang terkenal dalam obstetrik, ginekologi dan onkologi. Dalam kajian ini, gabungan nanopartikel emas (AuNPs) yang difungsikan dengan glutathion disulfida (GSSG) dan peranti analisis mikrofluidik berasaskan kertas (μ PAD) digunakan untuk mengesan dan menilai prestasinya terhadap pengesanan HCG dalam aspek kelurusan, kepekaan dan selektiviti. AuNPs yang digunakan dalam kajian ini berjaya disintesis dan dicirikan dengan rentang ukuran zarah dan hidrodinamik pada 4.89 - 34.40 nm dan 31.03 nm - 10.19 μ m masing-masing dan dihadkan dengan ion sitrat. AuNP yang difungsikan dengan GSSG berjaya ditukar sebagai sensor berasaskan AuNP untuk HCG. Walau bagaimanapun, disebabkan kekurangan penglihatan pada perubahan warna, sensor tidak dapat digunakan pada μ PAD dengan berkesan. Sementara itu, sensor berasaskan AuNP menunjukkan prestasi pengesanan yang baik kerana ia bertindak balas secara linear kepada perubahan kepekatan HCG dengan nilai R^2 bersamaan 0.9877, bersamaan dengan kepekaan yang rendah berbanding dengan kajian terdahulu dengan nilai pengesanan terendah 0.1577 ppm. Walaubagaimanapun, sensor yang dihasilkan tidak begitu selektif terhadap HCG kerana kajian gangguan sensor menunjukkan gangguan tinggi apabila dicampurkan dengan bendasing.

ABSTRACT

Detection of human chorionic gonadotropin (HCG) is one of many prominent diagnostics methods in obstetrics, gynaecology and oncology. In this study, the combination of gold nanoparticles (AuNPs) functionalized with glutathione disulfide (GSSG) and microfluidic paper-based analytical device (μ PAD) was used to detect and assess its performance on detection of HCG in the aspect of linearity, sensitivity and selectivity. AuNPs used in this study was successfully synthesized and characterized with particle and hydrodynamic size ranges at 4.89 – 34.40 nm and 31.03 nm – 10.19 μ m respectively and capped with citrate ion. GSSG-functionalized AuNPs was successfully derivatized as the AuNP-based sensor for HCG. However, due to lack of visibility on the colour changes, the sensor was unable to be applied on the μ PAD effectively. Meanwhile the AuNP-based sensor shown a good sensory performance as it responded linearly to the changing concentration of HCG with R^2 value of 0.9877. The sensor also shown high sensitivity compared to previous studies with limit of detection value of 0.1577 ppm. Yet, the sensor produced was not highly selective to HCG as interference study shown high interference from the impurities added.

AL-MULAKHKHAS

بعد الكشف عن موجهة الغدد التناسلية المشيمية البشرية أحد طرق التشخيص البارزة في أمراض النساء والتوليد والأورام. في هذه الدراسة ، تم استخدام مزيج من جزيئات الذهب النانوية الوظيفية مع ثاني كبريتيد الجلوتاثيون وجهاز تحليلي مستند إلى ورق ميكروفلويديك للكشف عن وتقييم أدائها عند الكشف عن موجهة الغدد التناسلية المشيمية البشرية في الجانب الخطي والحساسية والانتقائية. تم تصنيع الجسيمات النانوية الذهبية المستخدمة في هذه الدراسة بنجاح وتميزت بحجم الجسيمات والهيدروديناميكية في 4.89 – 34.40 نانومتر و 31.03 نانومتر – 10.19 ميكرون على التوالي وتوجت مع أيون سترات. تم اشتقاق الجسيمات النانوية المصنوعة من ثاني كبريتيد الجلوتاثيون بنجاح كجهاز استشعار قائم على الجسيمات النانوية الذهبية لمكورات الغدد التناسلية المشيمية البشرية. ومع ذلك ، بسبب عدم وضوح رؤية تغيرات الألوان ، لم يكن المستشعر قابلاً للتطبيق على الجهاز التحليلي المستند إلى ورق ميكروفلويديك بشكل فعال. وفي الوقت نفسه ، أظهر المستشعر المستند إلى الجسيمات النانوية الذهبية أداءً حسياً جيداً حيث استجاب خطياً للتركيز المتغير لمكورات الغدد التناسلية المشيمية البشرية بقيمة R^2 تبلغ 0.9877. كما أظهر المستشعر حساسية منخفضة مقارنة بالدراسات السابقة مع تحديد قيمة الكشف البالغة 0.1577 جزء في المليون. ومع ذلك ، فإن المستشعر المنتج لم يكن انتقائياً للغاية لمكورات الغدد التناسلية المشيمية البشرية حيث أظهرت دراسة التداخل تدخلاً كبيراً من الشوائب المضافة.

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LIST OF UNITS OF MEASUREMENTS

Temperature	°C
Volume of liquid	ml
Concentration of solution	mM
	%
	ppm
	mIU/ml
Wavenumber	cm ⁻¹
Wavelength	nm
Particle size	nm
Hydrodynamic size	nm
Mass	mg
	g
Amount of mole	mole
Channel width	mm
Flowrate of fluid	μl s ⁻¹
Pressure difference	Pa

LIST OF ABBREVIATIONS

HCG	Human chorionic gonadotropin
POCT	Point-of-care testing
AuNPs	Gold nanoparticles
μPADs	Microfluidic paper-based analytical devices
SPR	Surface plasmon resonance
FRET	Förster resonance energy transfer
AgNPs	Silver nanoparticles
FTIR	Fourier transform infrared
DLS	Dynamic light scattering
EDC HCl	N-(3-dimethylaminopropyl)-N'-ethylcarbodiimide hydrochloride
NHS	N-hydroxysuccinimide
GSSG	Glutathione disulfide
PdI	Polydispersity index
LOD	Limit of detection
LOQ	Limit of quantification